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EMERGING RESEARCH CONCEPT IN LIFE SCIENCES

First Edition

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Editors

Dr. R. B. Tripathi

Dr. R. Banupriya

Jayashree. R

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EMERGING RESEARCH CONCEPT IN LIFE SCIENCES



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Emerging Research Concept in Life Sciences

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**Dr. R.B. Tripathi
Dr. R. Banupriya
Jayashree. R.**

**Thanuj International Publishers,
Tamil Nadu, India**

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Preface

Emerging Research Concepts in Life Sciences have brought about remarkable progress in various fields, including Entomology, biodiversity, neuroscience, ecology, bioinformatics, microbiome therapy, food technology, microbiology, food science, forensic pathology, nanotechnology, health information science, agriculture and technology, food science, microbial ecology, medical sciences, pharmacology, rheumatology, immunotherapy, bioremediation, neurology, biotechnology and medicinal plants. We aim to foster scientific curiosity, inspire further research and contribute to the advancement of knowledge in these fields.

We are very much thankful to Thanuj International Publishers who readily accept and publish this subject. Also the author is very much thankful to Dr.G.K.Saravanan, Dr.M.D.Balakumaran, Dr.K.Arulmeha Ponradha, Shoba Gunasekaran, Raghavi TR, Rakshitha VS., Dr.P.Venkatachalam, Dr.S.Anbumalar, Mrs.S.Ajithadev, Dr.V.Subasshini, P.Srioviya, Dr.A.Balasubramanian, Dr.R.Vijaya Vahini, Divyasri,T., Sakshi Mahbubani, Dr.G.Sangeetha Vani, Ashwini,A., Mary George, Dr.V.P.Mahesh Kumar, S.Akash, R.Subaratinam, A.Suhaina Shafa, V.Sowndharya, Dr.N.Uma Maheswari, V.Dharani, M.Nanthini, M.Atchaya, K.Kaviya, S.Dharshini, Mr.Santosh, S.M.Sc., Farhana,V.S.George Miller, S.Mahendran, M.Nivetha, K.Gautam, T.Aravindhana, Poojashree, R.E.S.Janagakumar, Dr.P.K.SenthilKumar, C.Rajamani, K.Kolanjinathan, B.Sowbakkiyalakshmi, Karthick,M., Anand,M., Muthukatturaja, M.Murugan,A., Sathyananth,M., Ram Kumar, R.,A. Sasi Bharathi, S.Akash, Jansirani,D., Sabarunisha Begum, S., Mohamed Rizwana and A.Noorjahan Rehana, A for contributing their help and support for this work.

Dr. R.B. Tripathi
Dr. R. Banupriya
Jayashree. R.

About Editors



Dr.R.B.Tripathi is currently working as Assistant Professor in P.G.Department of Zoology, M.L.K.P.G.College, Balrampur-271201, Uttar Pradesh, India. He has been completed his Ph.D.in Zoology from Dr. R.M.L. Avadh University, Ayodhya, Uttar Pradesh, India. He has 23 years teaching experience in U.G and 19 years teaching experience in P.G classes, published 22 book chapters, 46 research papers in international and national reputed journals, participated and presented papers in many international and national seminars, conferences and workshops. He is Indian Zoologist, published by Surya Scientist Unique Researchers Yare Association, 2015. He is Associate Editor in International Journal of Advanced Research in Biological Sciences (ISSN:2348-8069), Editorial board member in International Journal of Advanced Multidisciplinary Research (ISSN:2393-8870), Published 11 Edited bookserved as Editor for Publication such as Recent Trends in Life Sciences Research (ISBN:978-81-947071-3-4), published by Darshan Publishers, Tamil Nadu, India, Recent Advancements and Research in Biological Sciences (ISBN:978-81-952529-1-6), Current Trends in Biological Sciences (ISBN:978-93-94638-00-6), Current Research in Life Sciences (ISBN: 978-93-94638-22-8), Recent Research in Biosciences (ISBN:978-93-94638-25-9),Current Advances in Biosciences (ISBN:978-93-94638-64-8), Advances in Pharmaceutical and Biosciences Research (ISBN:978-93-94638-87-7), Advance Research Trends in Biology (ISBN:978-93-94638-75-4), Biological Resources for Sustainable Research (ISBN:978-93-94638-90-7), Emerging Trends in Human Cardiology and Physiology (978-93-94638-42-6) and Concrpts and Approach for Biosciences Research (ISBN:978-93-94638-53-2) published by Thanuj International Publishers Tamil Nadu, India.



Dr. Banupriya R is a Research Project Officer in Research and Development, Shrimathi Devkunvar Nanalal Bhatt Vaishnav College for women (Autonomous), Chennai, Tamil Nadu. She has 1 year of teaching experience and 9 more years of research expertise in different fields of Biological sciences. She has been an author and co author of 12 scientific publications and 4 book chapters with ISBN Numbers. To the expertise she has 2 notable publications in Springer and 1 in Scopus. She completed her UG in Shrimathi Devkunvar Nanalal Bhatt Vaishnav College for women (Autonomous), Chennai, Tamil Nadu in Department of Plant Biology and Plant Biotechnology and PG from University of Madras in CAS in Botany and Ph.D in Applied Plant Science- Biotechnology, University of Madras. To her credits she has been awarded with AGAR Young Researcher Award 2024 and the Best Social Activist Award 2023, reflecting her leadership and volunteer work in educating the needy. Her academics developed due to strong collaboration with Department of School education and Tamil Nadu science forum where she was honored as an evaluator of the scientific events. Her credibility extended by attending various workshops by Edu fabrica in IIT Madras and Internships in Government Sugar factory and Orchid Pharmaceuticals.



R. Jayashree currently serves as a Lecturer in the Department of Perfusion Technology at Vinayaka Mission's School of Allied Health Sciences, Chennai, Tamil Nadu. She had dedicated experience in teaching and research and also demonstrated a strong commitment to advancing medical science and education. She completed her undergraduate studies in Perfusion Technology at Meenakshi Medical College Hospital and Research Institute, Kancheepuram, Tamil Nadu. During her final year, she gained extensive clinical exposure through internships in various departments, including Emergency Medicine, General Medicine, General Surgery, and Paediatrics. Following her graduation, she further honed her practical skills as a Perfusionist and Physician Associate at Billroth Hospitals, Chennai, and later at M.S.D Hospital, Kancheepuram.

R. Jayashree has made significant contributions to the field of medical science through her scholarly work. She has authored three notable research articles: "*A Review on Psoriasis*" by WJPPS (ISSN 2278-4357) 1. "*Obesity Paradox in Myocardial Infarction*" (ISBN: 978-93-94638-90-7), published in *Biological Resources for Sustainable Research* by Thanuj International Publishers, Tamil Nadu, India 2. "*Pro BNP Elevation: Diagnostic and Prognostic Impact in Cardiovascular and NonCardiac Events*" by JETIR (ISSN-2349-5162) 3. In addition to her research articles, she has contributed five book chapters to esteemed publications. These include: 1. "*The Impact of Ventricular Assist Devices in Modern Cardiology*" (ISBN: 978-93-9463842-6) 2. "*Robotic Assistance in Cardiac Surgery: A Glimpse into the Future*" (ISBN: 978-93- 94638-42-6) 3. "*Nutrition, Physical Activity, and Lifestyle: Shaping Cardiac Health Tomorrow*" (ISBN: 978-93-94638-42-6) 4. "*Evolving Paradigms in Cardiac Rehabilitation: Personalized Patient Care for Tomorrow*" (ISBN: 978-93-94638-42-6). These chapters were published in the book *Emerging Trends in Human Science* by Thanuj International Publishers, Tamil Nadu, India. R. Jayashree has also served as an editor for esteemed publications, including *Emerging Trends in Human Science* by Thanuj International Publishers, as well as other contributions in cardiology and physiology. Her active participation in national seminars, conferences, and workshops — where she has presented posters, organized webinars, and engaged in knowledgesharing — underscores her dedication to fostering collaboration in the medical community. With her multifaceted expertise in cardiac sciences, perfusion technology, and allied health, R. Jayashree continues to inspire innovation and excellence in her field.

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CONTENTS

S.No	Chapters	Pages
1	Insecticidal: Poisoning- Treatment Dr. R.B.Tripathi	1-4
2	Exploring Sea grasses Diversity and Distribution in Asia and Australian continents: A Sustainable Approach Dr. R. Banupriya, Dr. G.K. Saravanan and Dr. M.D. Balakumaran	5-25
3	"Neuroplasticity: Anatomical Insights into Brain Adaptability" Jayashree R	26-40
4	Ecological Resources for Sustainable Research Dr. K. Arulmeha Ponradha	41-52
5	Overview of Bioinformatics: Definition, History, Key Areas, and Applications Shoba Gunasekaran, Raghavi TR, Rakshitha V S	53-65
6	Microbiome-based therapeutics: Opportunities and challenges Dr. P.Venkatachalam, Dr. S. Anbumalar, Mrs. S. Ajithadevi	66-78
7	Almond peel: A bioactive compounds and the waste to wealth approach Dr.V. Subasshini, P. Srioviya	79-95
8	"The Vicious Cycle of Multidrug-resistant (MDR) bacteria" Dr.A.Balasubramanian	96-108
9	Sourdough: A Journey Through Tradition, Innovation, and Nutrition Dr. R. Vijaya Vahini, Divyasri T	109-126
10	From Spores to Clues: Fungi, a Potential Tool in Crime Investigations Sakshi Mahbubani, Dr. G. Sangeetha Vani	127-147

11	Recent advances in Nanomaterials for Sustainable development Dr. Ashwini. A and Dr. Mary George	148-165
12	Real-World Data Analytics for Outcome-Based Healthcare Dr. V. P. Maheshkumar	166-181
13	Assessment of compost maturity and seed germination index S. Akash	182-194
14	Palmyra Tree -An Source of Nutritious Food R. Subaratinam, A. Suhaina Shafa ,V. Sowndharya	195-208
15	Microbial Ecology: Microorganism In Natural Environment Dr.N. Uma Maheswari and V.Dharani	209-216
16	Medical science and its foundations Dr N.Uma Maheswari and M.Nanthini	217-225
17	Crop Production: Techniques and Trends Dr.N. Uma Maheswari and M. Atchaya	226-237
18	Artificial Intelligence In Sustainable Agriculture Dr.N.Uma Maheswari and K.Kaviya	238-250
19	The Role of Medicinal Plants In Modern Pharmacology Dr.N.Uma Maheswari and S.Dharshini	251-262
20	Tocilizumab, an anti il – 6 receptor monoclonal antibody to treat severe covid-19 patients during pandemic. Mr. Santhosh. S, Msc, Farhaana .v	263-282
21	Immunotherapy: Innovations and Clinical Applications S. George Miller	283-296
22	Factors Affecting Bioremediation and Its Types S. Mahendran, M. Nivetha, K. Gautam and T. Aravindhan	297-305
23	"A novel ray on the disease's path Recognition, and Rehabilitation of Myasthenia Gravis" Poojashree R.E, Jayashree R	306-313
24	Halophilic actinomycetes (<i>Streptomyces sp</i>) involved in antimicrobial resistance and its various applications S. Janagakumar & Dr. P. K. Senthilkumar	314-321

25	Extraction of secondary metabolites from halophilic actinomycetes and its characterization studies C.Rajamani, K.Kolanjinathan, B.Sowbakkiyalakshmi	322-329
26	Catastrophic Mass Mortality of <i>Hiatuladiphos</i> in the Punnaikayal Estuary: Implications of Climate Change and Extreme Weather Events Karthick M, Anand M, Muthukatturaja M, Murugan A Sathyananth M & Ramkumar R	330-337
27	Plants used as cosmetics A Sasi Bharathi and S Akash	338-354
28	Commercial Applications of Aeroponics Jansirani. D, Sabarunisha Begum. S, Mohamed Rizwana. A, Noorjahan Rehana. A	355-376

Insecticidal: Poisoning- Treatment

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Introduction

Pesticides (Insecticides, fungicides, acaricide, rodenticides etc.) are toxic not only to insects and other pests but also to human beings and domestic animals. They may affect human beings and livestock by ingestion through mouth, direct contact with bare skin or inhalation by breathing. This is only possible when they are not properly handled and proper precautions are not taken at the time of their application.

Pesticides

On the basis of mammalian toxicity pesticides are classified as:

- (I) Non hazardous pesticides (Pyrethrum products, sulphur etc.)
- (II) Moderately hazardous pesticides (DDT, BHC, malathion, carbaryl, fenitrothion, endosulfan etc.).
- (III). Dangerous pesticides (Parathion, endrin, DDVP, thimet, carbofuran etc.).

In large scale use of pesticides hazards may arise due to accidental or intentional poisoning, operational hazards during application, post application hazards due to residue etc. The following precautions should be strictly followed while handling pesticides:

1. Before use read the label on the pesticide container carefully and follow the directions strictly.
2. Pesticides should only be stored in the original packing in a locked room and not together with food stuffs.
3. Do not measure pesticides in spoons, cups etc., generally used in houses.
4. Destroy and bury empty pesticide containers and never use for storage of food stuffs or beverages.

5. Avoid skin contact specially of the concentrates and inhalation of dusts and spray mists.
6. Wash hand, face and other parts of the body thoroughly with soap and water when contaminated with pesticides.
7. Take a bath after application of more toxic pesticides.
8. Do not smoke or eat while handling pesticides.
9. Wear protective clothing, boots, rubber gloves, gloves and respiratory masks according to the toxicity of the pesticides.
10. A separate knife should be kept for opening pesticides bags or tin containers.
11. While preparing spray solutions bare hands should never be used for mixing solution. It is better to use long handled mixer to avoid splashing.
12. Spraying or dusting should immediately be stopped if a person feels tiredness, headache, nausea and vomiting.
13. In order to avoid toxic residues on crops at the time of consumption, the safety period must be strictly observed and the recommended rates of application must not be exceeded.
14. Avoid drift of pesticides to adjacent crops, streams, ponds and livestock.
15. Avoid the hottest hours of the day for pesticides application. Spraying should be done during early morning or late afternoon hours.
16. Protects the beneficial insects by applying insecticides during hours when bees are not visiting the plants and use selective insecticides.
17. Keep poultry, dairy or meat animals away from recently treated areas and observe the safety periods.
18. Call a doctor immediately when a person shows sign of poisoning.
19. Label the insecticidal treated fields.
20. First aid box should always be with the person who is giving insecticidal treatment. The box should contain the following articles:
 - a. One five ml Syringe
 - b. Ammonium carbonate, grinded mustard seed, common salt, potassium per manganate, tannic acid, vinegar, milk of magnesia and charcoal etc.
 - c. Amyl nitrite pearls, atropine sulfate, caffeine and apomorphine capsules.
 - d. A stomach cleaning tube.

Treatment in Case of Poisoning:

Following treatment should be kept in mind if poisoning takes place by mistake:

- I. Immediate efforts should be made to wash out the poison from the body.
- II. Use antidote at once.
- III. Excretory waste of the patient should be collected for medical examination.
- IV. The patient should be admitted to be nearest hospital.

Treatment in Case of swallowing poison:


- I. If the pesticides has been swallowed ,give to patient a tablespoon full of salt in a glass of warm water to induce vomittingand repeat until the vomit fluid is clear.It may also be induced by one gram zinc sulfate in a glass of water or apomorphinehydrochloride injection of 1/10 gram.
- II. The patient should lie down and be kept quiet.
- III. Never try to dispense anything to an unconscious person through the mouth.
- IV. Vomiting should not be induced if the patient feel fits or unconscious or he has taken petroleum products or concentrated acid or caustic soda.The stomach cleaning of such patients should be done under the expert guidance of a medical practitioner.
- V. Univesal antidotes (consisting of 2 part charcoal, one part tannic acid, one part of milk of magnesia) be given after vomiting or stomach cleaning.This mixture is useful in poisoning of acids, liquids, glycosides and heavy metals. Except in case of poisoning by corrosive substances it should be followed by gastric lavage.
- VI. Other antidotes are atropine sulfate, diacetylmonohexene and pyridine aldoxymet etc.

Treatment in case of inhaling the poison

- I. Carry the patient to fresh air.
- II. Open all windows and doors.
- III. Loosenall tight clothing.
- IV. Prevent the patient from chilling and wrap him in a blanket.
- IV. If breathing is irregular, apply artificial respiration and avoid vigorous application of pressure to the chest.

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Exploring Sea grasses Diversity and Distribution in Asia and Australian continents: A Sustainable Approach

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Abstract

Sea grass meadows represent highly productive and biologically affluent ecosystems that are located in tropical and temperate coastal waters across the globe. These ecosystems render indispensable services such as providing habitat for marine species, facilitating carbon sequestration, and supporting local fisheries. Asia and Australia share a geographical connection through the Indo-Pacific region, which forms a natural transition zone. This continuity makes them an excellent combined study area for assessing sea grass diversity, bio geographic patterns, and shared challenges. Thus the study focused on the two sub continents Asia and Australia for the diversity of sea grasses. In Southeast Asia, particularly in nations such as India, Sri Lanka, Indonesia, Malaysia, and the Philippines, is acknowledged as a global hot spot for sea grass biodiversity. These meadows harbor a diverse array of species, including dugongs and green sea turtles, and sustain the livelihoods of millions of individuals through their fisheries. Nevertheless, urban expansion and aquaculture present significant threats to their continued existence. In Australia, extensive sea grass meadows are situated along the coasts of Queensland and Western Australia, with notable habitats present in the Great Barrier Reef and Shark Bay. These regions are vital for biodiversity and carbon storage; however, they are becoming increasingly susceptible to the effects of climate change and natural disturbances such as cyclones. The conservation of these essential ecosystems necessitates the mitigation of both local and global hazards to guarantee their sustained role in marine biodiversity and climate regulation.

Key words: Sea grass meadows, Biodiversity hotspots, Asia, Australia, Sea grass Diversity

Introduction

Sea grasses constitute essential marine ecosystems that significantly contribute to the preservation of oceanic health and biodiversity. Located in shallow coastal waters throughout the world, these submerged flowering plants create dense underwater meadows that offer a multitude of ecological advantages, ranging from sediment stabilization and water quality enhancement and serve as a vital habitats for various marine species; thus, sea grasses are indispensable to both coastal and marine ecosystems.

The diversity and spatial distribution of sea grasses are subject to the influence of numerous factors, including climatic conditions, water salinity levels, availability of light, and nutrient concentration. Each continent offers a distinctive amalgamation of environmental circumstances and stressors, resulting in unique distributions of sea grasses and variations in species composition. For instance, tropical regions such as Southeast Asia are home to some of the most diverse sea grass ecosystems, whereas temperate zones typically exhibit a lower number of species but expansive meadows.

In addition to their ecological functions, sea grasses play a substantial role in the global carbon cycle through their capacity to sequester carbon dioxide, commonly termed "blue carbon." These ecosystems sequester carbon at rates significantly exceeding those of terrestrial forests, thereby fulfilling an essential function in the mitigation of climate change. Nevertheless, they are increasingly imperiled by anthropogenic activities, including coastal development, pollution, overfishing, and climate-induced alterations such as rising sea levels and ocean acidification.

This research investigates the Asia and Australian diversity and distribution of sea grasses, with a particular emphasis on their occurrence across the continent. Through the analysis of geographical patterns, environmental factors, and anthropogenic influences, the research seeks to underscore the significance of preserving these vital ecosystems. Comprehending the global variations in sea grass diversity and distribution is imperative for formulating strategies aimed at safeguarding and rehabilitating these delicate habitats in light of escalating environmental challenges and highlighting sea grass diversity in Asia and Australia contributes to global efforts, such as the United Nations Decade on Ecosystem Restoration and

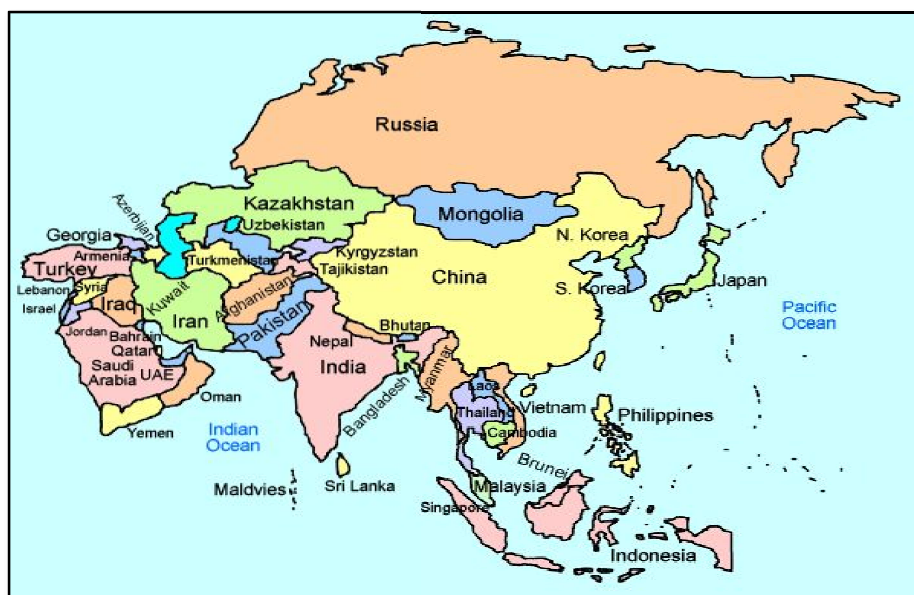
Sustainable Development Goals (SDGs), particularly SDG 14 (Life Below Water).

Biodiversity Hotspots and its impact on environment:

Sea grass meadows represent exceptionally productive and biologically diverse ecosystems, flourishing in tropical and temperate coastal marine environments worldwide. The diversity and extent of these meadows vary among continents, influenced by localized environmental conditions, oceanographic processes, and anthropogenic impacts.

Asia

Asia encompasses some of the most ecologically diverse sea grass ecosystems globally, particularly in Southeast Asia, which is frequently identified as a hotspot for sea grass biodiversity. Nations including India, Indonesia, Malaysia, and the Philippines are characterized by a remarkable variety of sea grass species. These meadows provide habitat for marine species while supporting livelihoods for fishermen. Nonetheless, urban development and aquaculture present considerable challenges to the sustainability of these ecosystems.



Map 1: Asian countries depicting the abundance of sea grasses
(<https://in.pinterest.com/pin/1618549841337384/>)

Australia

Australia is home to some of the world's most diverse and vital marine ecosystems, with sea grass playing a crucial role in maintaining the health of its coastal and marine environments. Sea grasses, which are underwater flowering plants, provide essential ecosystem services, including supporting marine biodiversity, stabilizing the seabed, improving water quality, and storing carbon. Australia's sea grass meadows, found along its vast coastline, are among the largest and most important in the world. However, sea grasses are facing growing threats from climate change, coastal development, and water pollution, making their conservation a priority for Australia's environmental efforts. Australia is recognized for its vast sea grass meadows, especially found along the coasts of Queensland and Western Australia. These regions play a crucial role in supporting biodiversity and sequestering carbon; however, they are susceptible to the effects of climate change and natural events such as cyclones.



Map 1: Australian countries depicting the abundance of sea grasses
(<https://www.worldatlas.com/maps/australia>)

List of Sea grass species in Asia:

Asia is home to diverse sea grass species, particularly in tropical and subtropical regions. The table 1 depicts the most diverse sea grass species present in Asia.

Table 1: List of most diverse Sea grass species in Asia

S.No	Sea grass	Common name	Location	References
1	<i>Thalassia hemprichii</i>	Pacific turtlegrass	Coastal waters of the Philippines, Indonesia (e.g., Sulawesi, Bali), Malaysia (e.g., Sabah and Sarawak), Thailand (e.g., Andaman Sea), Vietnam (e.g., Gulf of Tonkin)	Fortes <i>et al.</i> , 2018
2	<i>Halodule uninervis</i>	Narrow leaf Sea grass	Red Sea coastal areas (e.g., Saudi Arabia), Persian Gulf (e.g., UAE, Iran), Indian Ocean coasts (e.g., Sri Lanka, Maldives), Southeast Asia (e.g., Thailand, Indonesia, Malaysia).	Perumal Parthasarathi <i>et al.</i> , 2021
3	<i>Cymodocea rotundata</i>	Ribbon Sea grass	South east Asia (e.g., Malaysia, Indonesia, Vietnam) India (e.g., Gulf of Mannar), Andaman and Nicobar	Waycott <i>et al.</i> , 2004

Emerging Research Concept in Life Sciences

			Islands, Coastal areas of China (e.g., Hainan Island)	
4	<i>Syringodium isoetifolium</i>	Noodle Sea grass	East Africa to Southeast Asia (e.g., Kenya to Philippines) Indian coastal regions (e.g., Lakshadweep Islands), Indonesia (e.g., Bali and Kalimantan), Japan (e.g., Okinawa)	Clores and Agoo, 2013
5	<i>Halophila ovalis</i>	paddle weed, spoon grass or dugong grass	South east Asia (e.g., Thailand, Vietnam, Malaysia), Indian coasts (e.g., Palk Bay, Gulf of Mannar), Pacific Islands near Asia (e.g., Papua New Guinea), Japan (e.g., Kyushu)	Nguyen, 2014
6	<i>Enhalus acoroides</i>	Tape Sea grass	Philippines (e.g., Palawan and Mindoro), Indonesia (e.g., Java, Sulawesi), Malaysia (e.g., Sabah and Peninsular	Jillian Lean Sim Ooi <i>et al.</i> , 2011

Emerging Research Concept in Life Sciences

			Malaysia), Vietnam (e.g., Nha Trang Bay), Coastal Thailand	
7	<i>Halophila decipiens</i>	Carribean Sea grass or Paddle grass	Shallow coastal areas of Southeast Asia (e.g., Indonesia, Philippines), Indian Ocean regions (e.g., Sri Lanka, Maldives), China (e.g., South China Sea)	Nguyen, 2014
8	<i>Thalassodendron ciliatum</i>	Sickle- leaved cymodocea	Indian Ocean coastlines (e.g., Sri Lanka, Maldives), Southeast Asia (e.g., Indonesia, Malaysia), Coastal waters of the Andaman Sea	Chanda, 2022
9	<i>Halodule pinifolia</i>	Needle Sea grass	Philippines (e.g., Visayas, Palawan), Indonesia (e.g., Borneo, Sulawesi), Malaysia (e.g., Sabah), Vietnam (e.g., Mekong Delta)	Susantha Udagedara <i>et</i> <i>al.</i> , 2017

Emerging Research Concept in Life Sciences

10	<i>Zostera japonica</i>	Dwarf eelgrass or Japanese Eelgrass	Japan (e.g., Honshu, Hokkaido), Korea (e.g., Jeju Island), China (e.g., Bohai Sea and Yellow Sea)	Zhang, 2021
11	<i>Halophila minor</i>	Dwarf Sea grass	Thailand (e.g., Andaman Coast), Indonesia (e.g., Banda Sea), Philippines (e.g., Palawan and Cebu)	Kwok-Leung, 2006
12	<i>Cymodocea serrulata</i>	Serrated ribbon sea grass	India (Gulf of Mannar, Palk Bay and Andaman and Nicobar Islands), Sri Lanka, Maldives, Indonesia (Bali, Sulawesi and Maluku Islands), Thailand and Myanmar (Andaman sea), Malaysia (Peninsular Malaysia and Sabah), Philippines (Visayan Islands and Palawan), Vietnam	Arriesgado, 2015.

Emerging Research Concept in Life Sciences

			(South China Sea)	
13	<i>Halophila beccarii</i>	Ocean Turf Grass	Southeast Asia (e.g., Philippines, Indonesia), Indian Ocean (e.g., Maldives, Sri Lanka), India (e.g., Gulf of Mannar, Palk Bay)	Parthasarathy, 1988
14	<i>Zostera marina</i>	Eelgrass, Common Eelgrass, Widgeon grass, Broad-leaved grass wrack, Marlee, Sedge, Slitch, Sweet Sea-grass	Japan (e.g., Hokkaido, Honshu), South Korea, China (e.g., Bohai Sea, Yellow Sea)	Chefaoui, 2018
15	<i>Thalassia testudinum</i>	Turtle Grass	Found in the tropical regions of the Indian Ocean and parts of Southeast Asia, though it is more common in the Caribbean and the Americas.	Tussenbroek, 2007
16	<i>Halophila stipulacea</i>	Halophila seagrass or Broadleaf seagrass	Red Sea, Mediterranean and Indian Ocean, coastal areas of India (e.g., Gujarat), Sri Lanka, and	Marlene Wesselmann <i>et al.</i> , 2021

Emerging Research Concept in Life Sciences

			the Maldives	
17	<i>Cymodocea angustata</i>	-	Western Indian Ocean, Somalia, India (e.g., Gulf of Mannar), Sri Lanka, and the Maldives	Short <i>et al.</i> , 2007
18	<i>Ruppia maritima</i>	Beaked tasselweed, beaked ditchgrass, ditch grass, tassel pondweed and widgeon grass	India (e.g., Gujarat), Thailand, and southern Japan, brackish waters of the coastal areas of the Philippines and Indonesia	Bijan Mostafazadeh <i>et al.</i> , 2024
19	<i>Halodule wrightii</i>	Shoal grass or shoalweed,	Indian Ocean, India (e.g., Gujarat, Kerala), Maldives, and Sri Lanka.	Dunton <i>et al.</i> , 1994
20	<i>Syringodium filiforme</i>	Manatee grass	Philippines, Indonesia, and Malaysia.	Buzzelli <i>et al.</i> , 2012
21	<i>Thalassia ovata</i>	-	Philippines, Indonesia, and other regions of Southeast Asia	Gole Swapnali <i>et al.</i> , 2023
22	<i>Halophila baillonis</i>	Clover grass	Southeast Asia, particularly in Indonesia and the Philippines.	Van Dijk, <i>et al.</i> , 2023

Emerging Research Concept in Life Sciences

23	<i>Zostera muelleri</i>	Eelgrass Garweed	or	Southeast Asia (including the Philippines), and New Zealand	Leduc <i>et al.</i> , 2011
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List of Sea grass species in Australia:

Australia is home to a diverse range of sea grass species, particularly along its coastlines, which play crucial ecological roles in supporting marine life and coastal ecosystems.

S.No	Sea grass	Common name	Location	References
1	<i>Zostera muelleri</i>	Eelgrass or Garweed	Southern and Eastern coasts of Australia, including parts of New South Wales, Victoria, and Tasmania	Stafford-Bell <i>et al.</i> , 2019
2	<i>Posidonia australis</i>	Fibre-ball weed or ribbon weed	Temperate waters of southern Australia, particularly around Western Australia, South Australia, and Victoria.	Giulia Ferretto <i>et al.</i> , 2021
3	<i>Halophila ovalis</i>	paddle weed, spoon grass or dugong grass,	Australia's tropical and subtropical regions, including parts of Queensland and northern	Carruthers <i>et al.</i> , 2002

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			Western Australia.	
4	<i>Cymodocea nodosa</i>	Little Neptune grass	Queensland and Northern Territory of Australia	Julia Máñez-Crespo., 2020
5	<i>Halophila decipiens</i>	Carribean Sea grass or Paddle grass	Western Australia and the Northern Territory.	Western Australian Herbarium, 2024
6	<i>Thalassia testudinum</i>	Turtle Grass	Northern coasts of Queensland and the Great Barrier Reef.	Ugarelli <i>et al.</i> , 2019
7	<i>Syringodium isoetifolium</i>	Noodle Sea grass	Tropical waters of northern Australia, including the Great Barrier Reef.	Collier <i>et al.</i> , 2021
8	<i>Enhalus acoroides</i>	Tape Sea grass	Northern Australia, including parts of Queensland, Northern Territory, and Western Australia.	Rattanachot & Prathap 2011
9	<i>Halodule uninervis</i>	Narrow leaf Sea grass	Northern Australia, including parts of Queensland, Northern Territory, and Western Australia.	Wehbe <i>et al.</i> , 2024

Emerging Research Concept in Life Sciences

10	<i>Thalassia hemprichii</i>	Pacific turtlegrass	Tropical coastal waters in the northern parts of Australia, including Queensland and Western Australia	Kuo <i>et al.</i> , 1991
11	<i>Halophila beccarii</i>	Ocean Turf Grass	Shallow tropical waters of northern Queensland and the Northern Territory	Ganesh <i>et al.</i> , 2015
12	<i>Zostera marina</i>	Eelgrass, Common Eelgrass, Widgeon grass, Broad-leaved grass wrack, Marlee, Sedge, Slitch, Sweet Sea-grass	Southern and Eastern Australia, including New South Wales and Tasmania.	Yu <i>et al.</i> , 2023
13	<i>Cymodocea serrulata</i>	Serrated ribbon sea grass	Queensland and Northern Territory coasts.	Pollard and Greenway, 1993
14	<i>Ruppia maritima</i>	Beaked tasselweed, beaked ditchgrass, ditch grass, tassel pondweed and widgeon grass	Southern and Southeastern coastal regions of Australia	Jacobs, Margaret Brock, 1982
15	<i>Halophila stipulacea</i>	Halophila seagrass or Broadleaf seagrass	Northern parts of Australia, including Queensland and Great	Gambi <i>et al.</i> , 2009

Emerging Research Concept in Life Sciences

			Barrier Reef	
16	<i>Halodule wrightii</i>	Shoal grass or shoalweed	Tropical and subtropical regions of northern Australia, including Queensland and Northern Territory	Carruthers <i>et al.</i> , 2002
17	<i>Halophila minor</i>	Dwarf Sea grass	Northern coast of Australia, including Queensland and the Northern Territory	Calvin McMillan, 1983
18	<i>Cymodocea angustata</i>	-	Tropical coasts of Northern Australia, including Queensland	Calvin McMillan <i>et al.</i> , 1983
19	<i>Thalassodendron ciliatum</i>	Sickle-leaved cymodocea	Northern parts of Western Australia, Queensland, and Northern Territory	Walker and Prince, 1987
20	<i>Cymodocea rotundata</i>	Ribbon Sea grass	Tropical and subtropical regions of Australia's northern coasts, including Queensland and Northern Territory	Asch. & Schweinf, 1870

Emerging Research Concept in Life Sciences

21	<i>Syringodium filiforme</i>	Manatee grass	Northern Australia, particularly in the tropical waters of Queensland and around the Great Barrier Reef.	Ugarelli <i>et al.</i> , 2019
22	<i>Halophila australis</i>	Paddle Weed	Southern regions of Australia, particularly in temperate waters off the coasts of South Australia, Victoria, and Tasmania.	Doty and Stone, 1966
23	<i>Zostera capricorni</i>	Eel grass	Eastern coast of Australia, from southern Queensland to northern New South Wales, as well as in some southern regions like South Australia.	Surrey <i>et al.</i> , 2006
24	<i>Ruppia tuberosa</i>	Tuberous seatassel, widgeon grass	Found in estuarine and brackish waters along the coast, including in areas of Western Australia and	Kate Frahn <i>et al.</i> , 2012

			parts of Queensland	
25	<i>Posidonia oceanica</i>	Neptune grass or Mediterranean tapeweed	more common in the Mediterranean, it has been reported in some scattered regions in Australia, mainly around Western Australia	Tânia Aires <i>et al.</i> , 2011

Conclusion

The diversity and distribution of sea grasses across Asia and Australia underscore their ecological and environmental significance in these regions. Both areas host extensive sea grass meadows that support vibrant marine ecosystems, protect coastlines, and contribute to global carbon storage. Asia is recognized for its high sea grass species diversity, while Australia boasts some of the largest and most well-preserved meadows globally. Despite these natural advantages, sea grasses in both regions face growing pressures from human activities, climate change, and habitat degradation. Collaborative efforts in conservation, sustainable management, and community engagement are essential to safeguard these vital ecosystems, ensuring they continue to thrive and provide invaluable services for generations to come.

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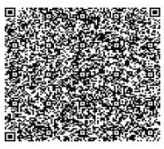
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"Neuroplasticity: Anatomical Insights into Brain Adaptability"

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Introduction

Neuroplasticity refers to the nervous system's capacity to reorganize its structure, function, and connections in response to intrinsic or extrinsic stimuli. This adaptability enables the brain to modify neural pathways, facilitating learning, memory formation, and recovery from injury.

The concept of neuroplasticity has evolved over time. In 1890, William James introduced the idea that the brain is not as unchangeable as previously thought, suggesting that neural pathways could be altered by experience. Later, Polish neuroscientist Jerzy Konorski coined the term "neuroplasticity" to describe the brain's ability to reorganize itself by forming new neural connections. These early insights laid the foundation for modern neuroscience, challenging the long-held belief that the adult brain was fixed and incapable of change.



Early Discoveries in Neuroanatomy and Brain Adaptability

- Early research in neuroanatomy provided evidence supporting the brain's adaptability.
- In the late 18th century, Italian anatomist Michele Vincenzo Malacarne conducted experiments comparing the brains of trained and untrained animals, finding that the cerebellums of trained animals were substantially larger.
- Although these findings were initially overlooked, they hinted at the brain's capacity for structural change in response to experience.
- In the early 20th century, Santiago Ramón y Cajal, a pioneering neuroscientist, used the term "neuronal plasticity" to describe nonpathological changes in the structure of adult brains.
- His neuron doctrine, which described the neuron as the fundamental unit of the nervous system, served as an essential foundation for developing the concept of neural plasticity.

Importance of Neuroplasticity in Neuroscience and Clinical Medicine

- ❖ Understanding neuroplasticity is crucial in neuroscience and clinical medicine, as it underpins the development of effective rehabilitation strategies following brain injuries. Therapies designed to encourage the brain to reorganize and compensate for lost functions, such as targeted exercises or cognitive training, leverage the principles of neuroplasticity to enhance recovery.
- ❖ Moreover, neuroplasticity plays a vital role in learning new information and skills, highlighting the importance of practice and engagement in enhancing cognitive abilities.
- ❖ By harnessing the brain's inherent adaptability, interventions can be developed to treat various neurological conditions, improve mental health outcomes, and promote overall cognitive well-being.

Structural Basis of Neuroplasticity

The CNS comprises the brain and spinal cord, with neuroplasticity predominantly occurring in regions such as the cerebral cortex, hippocampus, and subventricular zone. The cerebral cortex is involved in higher-order functions like perception and cognition, while the hippocampus plays a crucial role in memory formation and spatial navigation. The subventricular zone, lining the lateral ventricles, serves as a niche for neural stem cells, contributing to

neurogenesis. These areas are integral to the brain's capacity to adapt and reorganize in response to various stimuli.

Cellular and Molecular Mechanisms

Role of Neurons, Glial Cells, and Synapses

- ❖ Neurons are the primary signaling units in the CNS, transmitting information through synapses.
- ❖ Glial cells, including astrocytes, oligodendrocytes, and microglia, support neuronal function and modulate synaptic activity.
- ❖ Astrocytes regulate neurotransmitter levels and maintain the blood-brain barrier, while oligodendrocytes are responsible for myelination, enhancing signal conduction. Microglia act as immune cells, responding to injury and disease.
- ❖ The dynamic interplay between neurons and glial cells facilitates synaptic plasticity, essential for learning and memory.

Axonal Sprouting and Dendritic Remodeling

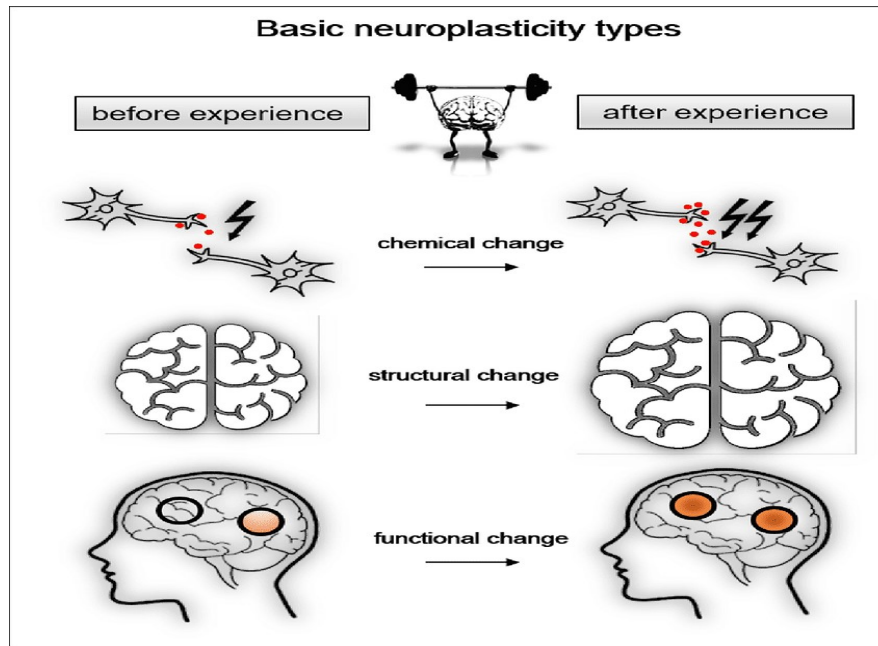
Axonal sprouting involves the growth of new axon terminals, enabling neurons to form additional synapses and re-establish connections after injury. Dendritic remodeling refers to changes in the structure of dendrites, including the formation or elimination of dendritic spines, which are small protrusions where synapses occur. These processes are fundamental to structural neuroplasticity, allowing the nervous system to adapt to new experiences or recover from damage.

Neurogenesis: Evidence from the Hippocampus and Subventricular Zone

Neurogenesis, the generation of new neurons from neural stem cells, persists in specific regions of the adult brain, notably the hippocampus and subventricular zone. In the hippocampus, particularly the dentate gyrus, new granule neurons are continuously integrated into existing neural circuits, contributing to learning and memory processes. Similarly, the subventricular zone supplies new neurons that migrate to the olfactory bulb, playing a role in olfactory functions.

These findings underscore the significance of neurogenesis in maintaining neural plasticity and cognitive function throughout life.

1. Types of Neuroplasticity:



Neuroplasticity refers to the brain's ability to reorganize itself by forming new neural connections throughout life. This adaptability allows neurons to adjust their activities in response to new experiences or changes in the environment. The primary types of neuroplasticity include.

Functional Plasticity

Functional plasticity involves the brain's capacity to move functions from damaged areas to undamaged regions. This reorganization enables the restoration of abilities compromised by injury or disease. For instance, following a stroke, language functions may shift from the left hemisphere to the right to compensate for damage. This adaptability is crucial for recovery and rehabilitation.

Structural Plasticity

Structural plasticity refers to the brain's ability to change its physical structure in response to learning, experience, or environmental changes. This includes the formation or elimination of synapses, dendritic branching, and alterations in gray matter density. These modifications facilitate the development of new skills and memories. For example, studies have shown that extensive practice in musicians leads to increased gray matter volume in regions associated with finger movements.

Developmental Plasticity

Developmental plasticity occurs during critical periods of development, particularly in early childhood, when the brain is highly malleable. During this time, neurons rapidly form new connections, and synaptic pruning eliminates weaker connections, optimizing neural networks for efficient functioning. This process is essential for normal cognitive and behavioral development. Disruptions during these periods can lead to developmental disorders.

Adaptive Plasticity

Adaptive plasticity refers to the brain's ability to adjust to new experiences or changes in the environment by reorganizing its functions and structure. This form of plasticity underlies learning and memory formation and allows individuals to acquire new skills or adapt to sensory changes. For instance, in individuals who become blind, the visual cortex can be repurposed to process tactile information, enhancing other senses.

2. Neuroplasticity in Learning and Memory

Neuroplasticity in Learning and Memory

Neuroplasticity, the brain's ability to reorganize itself by forming new neural connections, is fundamental to learning and memory. This adaptability involves specific brain regions, synaptic mechanisms, and structural changes that facilitate the acquisition and retention of information.

Brain Regions Involved in Learning and Memory

The hippocampus and prefrontal cortex are pivotal in learning and memory processes. The hippocampus is essential for forming and retrieving declarative memories, such as facts and events, and plays a significant role in spatial navigation. The prefrontal cortex is involved in working memory, decision-making, and the integration of information necessary for complex cognitive behavior. The interaction between these regions enables the encoding, storage, and retrieval of memories.

Role of Long-Term Potentiation (LTP) and Long-Term Depression (LTD)

LTP and LTD are enduring changes in synaptic strength that serve as cellular mechanisms underlying learning and memory. LTP enhances synaptic transmission following high-frequency stimulation, leading to a sustained increase in signal strength between neurons. Conversely, LTD results from low-frequency stimulation, causing a prolonged decrease in synaptic strength. These processes modulate synaptic efficacy, thereby encoding information and facilitating synaptic plasticity essential for memory formation.

Structural Adaptations During Skill Acquisition and Habit Formation

Learning new skills and forming habits induce structural changes in the brain, including synaptogenesis (formation of new synapses), dendritic branching, and alterations in gray matter density. For example, motor skill learning is associated with increased dendritic spine formation in the motor cortex, enhancing neural circuitry involved in executing movements.

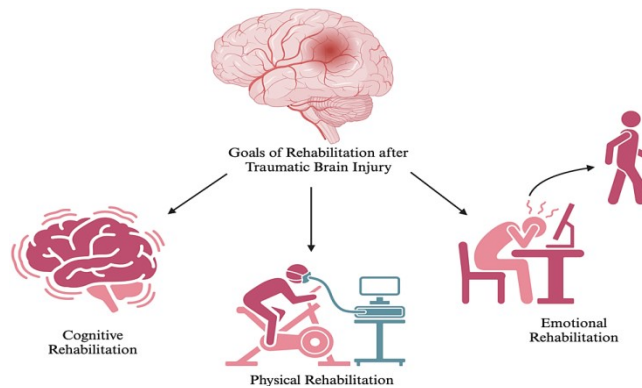
Recovery After Brain Injury

Neuroplasticity plays a pivotal role in recovery following brain injuries such as stroke and traumatic brain injury (TBI). Understanding the mechanisms that drive this adaptability, alongside effective rehabilitation strategies and emerging technologies, is essential for optimizing patient outcomes.

Mechanisms of Neuroplasticity in Stroke and Traumatic Brain Injury (TBI)

Following a brain injury, the central nervous system exhibits a remarkable capacity to reorganize itself. This reorganization involves the formation of new neural connections and the strengthening or weakening of existing ones, facilitating the restoration of lost functions. In stroke and TBI, neuroplasticity encompasses processes such as axonal sprouting, dendritic branching, and synaptogenesis, which collectively contribute to functional recovery. The extent and efficacy of these neuroplastic changes are influenced by factors including the injury's severity, the individual's age, and the timing and intensity of rehabilitation interventions.

Role of Rehabilitation in Promoting Plasticity:



Physical Therapy

Physical therapy (PT) is integral to neurorehabilitation, aiming to enhance motor function, balance, and coordination. Through repetitive, task-specific exercises, PT promotes neuroplasticity by reinforcing neural pathways associated with movement.

Cognitive Therapy

Cognitive therapy focuses on improving mental functions such as memory, attention, and problem-solving skills. By engaging patients in targeted cognitive exercises, this therapy leverages neuroplasticity to rewire neural circuits, thereby enhancing cognitive abilities impaired by brain injury.

Emerging Technologies to Enhance Recovery

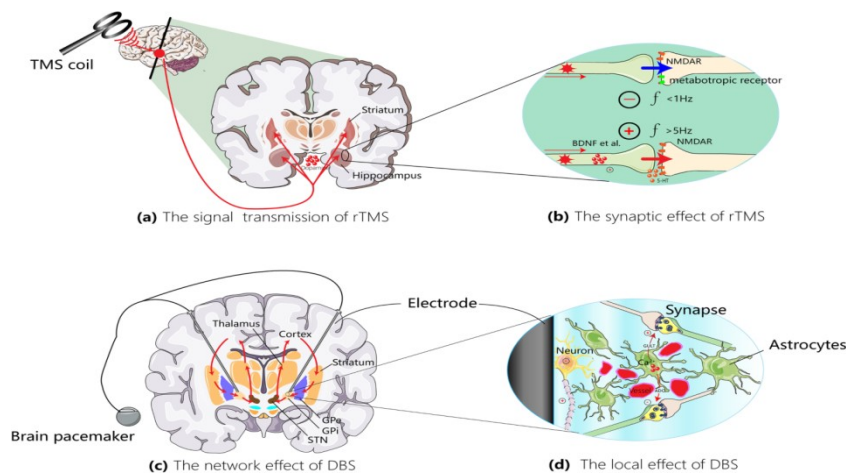
Brain stimulation (TMS, DBS)

- Transcranial Magnetic Stimulation (TMS) and Deep Brain Stimulation (DBS) are neuromodulation techniques employed to facilitate neuroplasticity.
- TMS uses magnetic fields to non-invasively stimulate specific brain regions, modulating neural activity to promote recovery.
- DBS involves the surgical implantation of electrodes to deliver electrical impulses to targeted brain areas.

Virtual Reality (VR)-based neurorehabilitation

- Virtual Reality (VR) offers immersive, interactive environments for rehabilitation, providing patients with engaging platforms to practice motor and cognitive tasks.
- VR-based interventions can simulate real-life scenarios, enhancing the relevance and effectiveness of therapy. Research indicates that VR can induce neuroplastic changes, improving functional outcomes in patients with brain injuries.

3. Neuroplasticity in Neurodegenerative Diseases:



Neuroplasticity, the brain's ability to reorganize itself by forming new neural connections, plays a significant role in the progression and potential treatment of neurodegenerative diseases such as Alzheimer's disease (AD) and Parkinson's disease (PD).

Role in Alzheimer's Disease

Early compensatory mechanisms

In the initial stages of Alzheimer's disease, the brain attempts to compensate for neuronal loss and synaptic dysfunction through neuroplastic mechanisms. These compensatory changes may involve the strengthening of existing synapses or the formation of new synaptic connections to maintain cognitive function. However, as the disease progresses, these mechanisms become overwhelmed, leading to cognitive decline.

Synaptic Failure And Neuroplasticity Deficits

Alzheimer's disease is characterized by the accumulation of amyloid-beta plaques and tau tangles, which disrupt synaptic function and impair neuroplasticity. Synaptic failure is a hallmark of AD, leading to deficits in learning and memory. The loss of synaptic connections correlates strongly with cognitive decline in AD patients.

Parkinson's Disease and Compensatory Plasticity

In Parkinson's disease, the degeneration of dopaminergic neurons in the substantia nigra leads to motor deficits. The brain exhibits compensatory plasticity by reorganizing neural circuits to mitigate these deficits. For instance, increased activity in the premotor and parietal cortices has been observed, which may help compensate for impaired motor function. However, these compensatory mechanisms are often insufficient to fully restore motor control.

Potential Therapeutic Approaches to Enhance Plasticity in Neurodegeneration

Enhancing neuroplasticity presents a promising therapeutic avenue for neurodegenerative diseases. Several strategies are under investigation:

Pharmacological interventions

Research is exploring drugs that can modulate neuroplasticity-related pathways. For example, certain cancer drugs targeting the enzyme indoleamine-2,3-dioxygenase 1 (IDO1) have shown potential in restoring

memory and brain function in Alzheimer's models by improving glucose metabolism in astrocytes.

Brain Stimulation Techniques

Techniques such as Transcranial Magnetic Stimulation (TMS) and Deep Brain Stimulation (DBS) are being investigated for their ability to enhance neuroplasticity. DBS, for instance, is a recognized therapeutic method that can improve neuroplasticity in neurodegenerative diseases.

Rehabilitation And Cognitive Training

Intensive amplitude-specific exercise-based therapeutic approaches that adhere to the principles of neuroplasticity have been introduced as possible solutions to improve rehabilitation outcomes for people with Parkinson's disease.

Innovative Therapies

Innovative therapeutic methods, such as deep brain stimulation (DBS), have been developed to improve neuroplasticity in neurodegenerative diseases.

External Influences on Neuroplasticity

External Influences on Neuroplasticity

Neuroplasticity—the brain's ability to reorganize itself by forming new neural connections—is influenced by various external factors, including environmental stimuli, lifestyle choices, and pharmacological agents. Understanding these influences is crucial for developing strategies to enhance brain health and cognitive function.

Environmental Factors

Enrichment And Stimulation

Exposure to enriched environments, characterized by increased sensory, cognitive, and social stimuli, has been shown to promote neuroplasticity. Such environments can lead to structural changes in the brain, including increased dendritic branching and synaptogenesis, thereby enhancing cognitive functions.

Studies have demonstrated that environmental enrichment positively affects neural development and repair, highlighting its potential therapeutic applications.

Stress And Trauma

Conversely, chronic stress and traumatic experiences can adversely affect neuroplasticity. Elevated levels of stress hormones, such as cortisol, can impair neurogenesis and synaptic plasticity, particularly in regions like the hippocampus. This impairment may contribute to cognitive deficits and mental health disorders. Research indicates that stress and trauma can negatively impact neural development and repair, underscoring the importance of stress management for brain health.

Lifestyle Factors

Exercise And Its Effects On Brain Structure And Function

Regular physical activity is a potent modulator of neuroplasticity. Exercise has been associated with increased hippocampal volume, enhanced synaptic plasticity, and improved cognitive functions. It promotes the release of neurotrophic factors, such as brain-derived neurotrophic factor (BDNF), which support neuronal growth and survival. Studies have shown that exercise can induce structural and functional brain plasticity, thereby enhancing cognitive performance.

Nutrition And Neuroplasticity

Dietary habits significantly influence neuroplasticity. Nutrient-rich diets, particularly those high in omega-3 fatty acids, antioxidants, and vitamins, support brain health by reducing inflammation and oxidative stress, which can impair neuroplasticity. Conversely, diets high in processed foods and sugars may have detrimental effects on brain function. Research indicates that a healthy diet positively affects cognitive reserve and neural plasticity.

Pharmacological Agents and Their Role in Modulating Plasticity

Certain pharmacological agents can modulate neuroplasticity. For instance, antidepressant medications have been shown to enhance neuroplasticity in regions such as the hippocampus, potentially contributing to their therapeutic

effects. Additionally, substances like nicotine and alcohol can have complex effects on neuroplasticity, with nicotine potentially enhancing and alcohol impairing neural plasticity. Understanding the impact of various pharmacological agents on neuroplasticity is essential for developing effective treatments for neurological and psychiatric conditions.

4. Advances in Research and Technologies

Advances in Research and Technologies

Recent advancements in neuroimaging, artificial intelligence (AI), and cutting-edge tools have significantly enhanced our understanding of neuroplasticity—the brain's ability to reorganize itself by forming new neural connections. These developments offer deeper insights into neural circuits and hold promise for therapeutic interventions.

Brain Imaging Techniques for Studying Neuroplasticity

Functional Magnetic Resonance Imaging (fMRI)

- ❖ fMRI measures brain activity by detecting changes in blood flow, providing insights into functional neuroplasticity. Studies have utilized fMRI to observe gray matter changes during learning and task-related recovery, highlighting its role in tracking neuroplasticity.

Diffusion Tensor Imaging (DTI)

- ❖ DTI assesses the integrity of white matter tracts by measuring the diffusion of water molecules, aiding in the study of structural neuroplasticity. It has been instrumental in examining changes in white matter associated with learning and recovery from injury.

Cutting-Edge Tools in Neuroanatomical Research



Optogenetics
Connectomics

5. Challenges and Controversies in Neuroplasticity Research

Neuroplasticity—the brain's ability to reorganize itself by forming new neural connections—is a dynamic field of study. However, several challenges and controversies persist, particularly concerning the limitations of current models and techniques, ethical considerations, and the variability of neuroplasticity across individuals and populations.

Conclusion

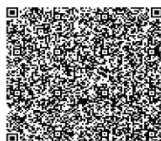
Neuroplasticity—the brain's capacity to reorganize itself by forming new neural connections—is fundamental to learning, memory, and recovery from injury. Understanding its mechanisms has profound implications across healthcare, education, and society.

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Ecological Resources for Sustainable Research

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Introduction

In recent years, the incorporation of ecological resources into sustainable research has become crucial for solving environmental concerns such as climate change, biodiversity loss, and resource depletion. Forests, water bodies, soils, and biodiversity are examples of ecological resources. These resources are critical for maintaining ecological balance and can have a substantial impact on the success of sustainable research projects. This chapter delves into the concept of ecological resources, their function in sustainable research, and how they might be effectively employed to address global environmental and socioeconomic challenges.

Sustainable research attempts to reduce environmental impact while promoting social fairness and economic prosperity. Understanding and utilizing ecological resources allows research to enhance biodiversity conservation, pollution reduction, and resource efficiency. However, this necessitates careful planning, resource management, and adherence to ethical norms. As the globe faces the accelerating challenges of climate change, environmental degradation, and resource depletion, there has never been a greater need for research that not only advances knowledge but also ensures ecosystems' long-term health.

Ecological resources, which range from water and soil to forests and biodiversity, constitute the foundation of sustainable development and are essential to human life and health. Sustainable research, therefore, requires a thorough understanding of how human activities interact with these resources and how they might be used in ways that do not jeopardize their future availability. By focusing on incorporating ecological principles into study methods, scientists can develop solutions that not only address important global concerns but also help to preserve and restore ecological systems. This

strategy not only seeks to reduce the environmental impact of research, but also encourages the development of sustainable technologies and practices that benefit both ecosystems and the communities that rely on them.

1. Understanding Ecological Resources

Ecological resources are vital components of the Earth's ecosystems, sustaining life and promoting environmental health. These resources supply the fundamental services that humans and all other living organisms require for survival, such as food, water, air, shelter, and energy-producing materials. Ecological resources are often divided into two categories: natural resources and ecosystem services. Both categories are essential to the survival of life on Earth and play an important role in sustainable research.

1.1 Natural Resources: The Building Blocks of Ecological Systems

Natural resources are raw materials provided by the Earth that can be used in a variety of ways to sustain life and economic activity. These resources are divided into renewable and non-renewable categories based on their availability and replenishment rates.

- **Renewable Resources**

Renewable resources, such as sunshine, wind, water, and biological organisms, renew naturally on a human timeline. Examples include forests, crops, and fish. Renewable resources can be handled in a sustainable way so that they are used without depleting their availability for future generations. The essential problem here is to use renewable resources in a way that does not surpass their natural regeneration potential. For example, solar energy is unlimited and limitless, making it a perfect renewable resource for long-term energy research.

- **Non-Renewable Resources**

Non-renewable resources are limited and cannot be replenished during a human lifetime. These include minerals like gold, copper, and rare earth metals, as well as fossil fuels like coal, oil, and natural gas. Overuse of these resources can result in their depletion, making their use in research and technology a delicate balancing act. Sustainable research aims to find alternatives or lessen reliance on non-renewable resources by emphasizing renewable technologies and conservation techniques.

1.2 Ecosystem Services: Nature's Contribution to Human Well-Being

Provisioning Services:

These are the direct results of ecosystems. They consist of food, water, timber, medicinal plants, and genetic resources. Sustainable research in agriculture, forestry, and fisheries frequently focuses on improving the delivery of these services while ensuring that they are extracted in an environmentally responsible way. For example, sustainable agriculture research could focus on increasing crop yields while preserving soil fertility and reducing pesticide use.

Regulating Services:

Climate regulation, water purification, flood management, disease regulation, and carbon sequestration are all examples of services that assist regulate environmental conditions. Forests, marshes, and oceans are especially important for their regulatory functions. Environmental science and climate change research frequently focuses on understanding and protecting these regulating services in order to offset the effects of human-caused climate change. Wetlands, for example, play an important role in water filtration and flood management, whereas woods serve as carbon sinks, absorbing significant volumes of CO₂ from the atmosphere.

Cultural Services:

These services offer non-material advantages such as recreational, artistic, spiritual, and cultural qualities. Research on ecosystem restoration and conservation is increasingly recognizing the value of these services, particularly in terms of biodiversity protection and community well-being. Natural landscapes, such as national parks and nature reserves, benefit both human health and cultural identity. Ecotourism research, for example, encourages the sustainable use of natural regions while preserving their cultural and aesthetic values.

Supporting Services:

These are the fundamental processes that enable all other ecosystem services, such as nitrogen cycling, soil formation, primary production, and biodiversity. The loss of supportive services can jeopardize the delivery of other services. Soil health, for example, is critical for agricultural output, while biodiversity conservation supports ecosystem resilience to perturbations. Ecological study frequently seeks to understand these processes and how they might be conserved through sustainable land use and conservation activities.

1.3 Ecological Footprint and Sustainability

The ecological footprint is a measure of the demands made on Earth's ecosystems by human activity, such as the resources needed to feed human populations and absorb garbage. It is an effective tool for analysing sustainability since it compares human use to the Earth's potential to replenish resources. Sustainable research seeks to lessen the ecological footprint by supporting methods that are compatible with ecological boundaries.

- **Carbon Footprint:** This is a subset of the ecological footprint that quantifies the quantity of CO₂ emitted into the atmosphere as a result of human activity, specifically the use of fossil fuels. Renewable energy research, such as wind and solar power, is critical for lowering carbon emissions and mitigating climate change.
- **Water Footprint:** This metric assesses the amount of water utilized to produce goods and services. Agriculture is the largest consumer of water, thus research on water-saving methods and technology is critical to ensuring that water resources are used sustainably.
- **Biodiversity Footprint:** The biodiversity footprint refers to the effects that human actions have on species and ecosystems. Sustainable conservation biology study aims to reduce biodiversity loss and maintain habitats through responsible land use and resource exploitation.

1.4 The Role of Ecological Resources in Sustainable Research

Ecological resources are crucial to sustainable research, which seeks to strike a balance between economic development and environmental protection. Research in subjects such as agroecology, ecological economics, environmental science, and sustainable engineering frequently focuses

- **Resource Conservation:** Sustainable research aims to conserve natural resources including water, soil, and forests. For example, sustainable agricultural research encourages approaches such as crop rotation, organic farming, and integrated pest control, all of which preserve soil health and reduce the need for chemical inputs.
- **Restoration and Rehabilitation:** Human activity has degraded ecosystems around the world. Ecological restoration research seeks to rebuild these ecosystems by restoring habitat, replenishing biodiversity, and reestablishing ecological functioning. This research is crucial for reversing the damage caused by deforestation, desertification, and pollution.

- **Sustainable Resource Use:** It also aims to develop efficient technologies and practices that minimize waste and reduce the environmental impact of production and consumption. Innovations in sustainable manufacturing, green chemistry, and energy-efficient technologies are crucial for minimizing non-renewable resource consumption.

1.5 The Interdependence of Ecological Resources

Ecological resources are interdependent. Ecosystem equilibrium is dependent on intricate interactions between species, climate, soil, and water systems. A single component failure might have far-reaching consequences for the entire system. For example, the loss of pollinators can affect food production, while soil erosion can contribute to poor water quality and flooding. Understanding these relationships is crucial for creating holistic, sustainable solutions. Ecology and environmental science research is critical for better understanding these complicated linkages and developing more effective conservation and resource management techniques. Sustainability encourages us to consider not only the direct usage of resources, but also their interdependence and the long-term health of the entire ecosystem.

2. Integrating Ecological Resources into Research Design

Integrating ecological resources into research design entails using ecological principles to ensure that research procedures, outcomes, and technology are not harmful to the environment. It is critical that study designs include both the direct and indirect effects of human activities on ecosystems, resulting in research that not only increases knowledge but also promotes environmental sustainability. This approach acknowledges the critical role that natural resources play in the health of the world and attempts to develop research approaches that respect and safeguard these resources.

2.1 Ecological Principles in Research Design

Integrating ecological concepts into study design requires a fundamental understanding of ecosystem function and the interactions between species, habitats, and resources. A research strategy that combines these concepts is more likely to account for the complexity and interdependence of ecological systems.

- **Holistic Systems Thinking:** Understanding the interdependence of ecological systems is an essential component of ecological research. Research designs should incorporate systems thinking, taking into account not only the research's immediate impact but also its larger

ecological implications. For example, ecological studies in agricultural systems must take into account soil health, water availability, biodiversity, and climate change, all of which have an impact on agricultural techniques' effectiveness.

- **Ecological Sustainability:** This concept ensures that research procedures do not harm ecological resources or destabilize ecosystems. For example, studies in forestry research should incorporate strategies that protect soil health, sustain biodiversity, and reduce deforestation. To maintain soil fertility and reduce pesticide use, agricultural researchers should incorporate sustainable farming strategies such as crop rotation and agroforestry into their designs.
- **Resilience and Adaptive Capacity:** Another principle is to conduct research that improves resilience in ecosystems. For example, climate change adaptation research could focus on generating crop types that are more resistant to temperature extremes and drought, so ensuring food security while protecting ecological systems.

2.2 Participatory Approaches and Local Ecological Knowledge

Integrating local ecological knowledge (LEK) into study design is an effective technique for promoting sustainability. Indigenous groups and local populations have a wealth of information about the natural environment, which is often passed down through generations and can provide vital insights into how ecosystems work and how to manage resources sustainably.

- **Community-Based Research:** Engaging local populations in the planning and implementation of research initiatives ensures that ecological resources are managed in accordance with traditional knowledge and cultural traditions. Local people, for example, may assist forestry researchers in identifying culturally acceptable and effective sustainable forest management strategies for biodiversity preservation.
- **Collaborative Decision-Making:** Collaborative research designs that incorporate local ecological knowledge and include local stakeholders in decision-making processes ensure that research initiatives are both environmentally sustainable and socially acceptable. This strategy contributes to study findings that can be more effectively implemented in local contexts and communities.

2.3 Incorporating Ecosystem Service Valuation into Research

Integrating ecosystem service valuation into study design entails calculating the worth of natural resources and ecosystem services in terms of their impact on human well-being and economic systems. This method assists academics in understanding the full advantages of preserving healthy ecosystems and offers a foundation for policy decisions.

- **Economic Valuation of Ecosystem Services:** Valuing ecosystem services, such as carbon sequestration, water purification, and pollination, can assist make the case for sustainable resource management. For example, a study project that focuses on the restoration of a wetland might quantify the benefits of flood control, water filtering, and biodiversity protection, which can be quantified and compared to the restoration costs.
- **Cost-Benefit Analysis in Ecological Research:** Integrating cost-benefit analysis into study design guarantees that a project's environmental benefits are addressed alongside economic expenses. In agricultural research, this could imply weighing the long-term environmental benefits of sustainable farming practices (such as reduced soil erosion and increased biodiversity) against the short-term economic gains of intensive farming.

2.4 Sustainable Methodologies for Data Collection

The methods used to collect data in ecological study have a substantial environmental impact. Traditional research methods frequently demand significant resource consumption, which can contribute to ecological damage if not adequately managed. As a result, ecological researchers should use sustainable data collection approaches.

- **Non-invasive research methods:** Non-invasive data gathering tools, such as remote sensing and satellite images, can help to reduce the environmental impact of research projects. For example, using drones to monitor wildlife populations or deforestation provides vital data without disrupting the habitats of the species under study.
- **Minimizing Resource Consumption:** Minimizing non-renewable resource consumption is vital in field research. For example, researchers can use renewable energy sources, such as solar power, to power research equipment in remote places, lowering their research's environmental impact.

- **Long-term Monitoring:** Long-term monitoring of ecological processes is crucial for understanding the effects of human activity on ecosystems. Long-term ecological monitoring in research designs allows for more precise assessments of environmental changes over time, such as biodiversity shifts or climate change's effects on ecosystems.

2.5 Integrating Life Cycle Analysis (LCA) in Research Design

Life cycle analysis (LCA) is a technique that assesses the environmental implications of a product's whole life cycle, from raw material extraction to disposal. Integrating LCA into research design allows researchers to assess the environmental effect of the goods, materials, and technologies they investigate or develop.

- **Product Development and Resource Efficiency:** LCA can be used in research on new technologies or goods, such as bioplastics or sustainable energy systems, to examine the environmental impact of these innovations across their whole life cycle. This ensures that the items under investigation do not have unforeseen negative environmental repercussions, such as increased resource use or waste generation.
- **Minimizing Environmental Impact:** LCA can also assist researchers explore ways to reduce the environmental impact of their own research efforts. For example, in the creation of sustainable materials, LCA can be used to discover the most resource-efficient and least polluting manufacturing processes.

2.6. Policy Implications and Integrating Ecological Resources into Policy Research

Research designs that include ecological resources have the ability to affect policy decisions. Sustainable research can give the evidence needed to design policies that protect ecosystems while encouraging economic growth.

- **Research to Inform Environmental Policy:** Ecosystem, biodiversity, and resource management research can help policymakers make informed decisions by providing scientific evidence of the value of ecosystem services and the necessity for sustainable practices. For example, research into the effects of deforestation could help develop forest management policies that prioritize conservation and long-term use of forest resources.
- **Policy Evaluation Tools:** Incorporating ecological resource evaluation methodologies into policy research assists policymakers in understanding the environmental impact of proposed legislation or regulations. For

example, research that simulates the impact of a new energy policy on ecosystems can inform decisions that ensure the long-term viability of both energy production and the natural environment.

2.7 Adaptive Management in Research

Adaptive management is a method for responding to uncertainties in ecological research by employing iterative, flexible management procedures that can be modified in response to new data or changing environmental conditions.

- **Responding to Ecological Changes:** Research must adapt to changing environmental conditions. This necessitates continual monitoring and flexible research design. For example, biodiversity conservation research may need to be revised in response to unexpected changes in species populations or new threats, such as invading species.
- **Building Resilience through Adaptive Management:** Adaptive management strengthens natural systems against perturbations like harsh weather and invasive species. By utilizing flexible, adaptive research methodologies, scientists can help ecosystems recover and retain their ecological functions in the face of these difficulties.

3. Case Studies in Sustainable Research

Conservation Research and Biodiversity

Sustainable conservation research strives to maintain biodiversity while reducing ecological footprints. The Wildlife Conservation Society (WCS) is one important example of an organization that works to safeguard endangered species by researching habitat loss, poaching, and the effects of climate change. Their research blends ecological data with community-based management, thereby promoting sustainable land use.

Sustainable Agriculture and Food Security

Scientists do sustainable agriculture research to develop farming systems that make efficient use of ecological resources. A significant example is agroforestry, which combines trees with crops or livestock to increase biodiversity, improve soil fertility, and minimize water consumption. The Food and Agriculture Organization (FAO) promotes sustainable food production research, including organic farming, agroecology, and the adoption of robust crop types.

Renewable Energy Research

Renewable energy is critical for minimizing reliance on non-renewable ecological resources. Research into sun, wind, and biofuel has resulted in the development of more efficient energy systems. Wind farms, for example, are now being developed on a global scale, with research focused on optimizing turbine positioning and design to maximize energy capture. In bioenergy, scientists are looking into algae-based biofuels and their ability to replace fossil fuels while lowering carbon emissions.

4. Challenges and Limitations

Resource Depletion and Overexploitation

Research efforts frequently involve the extraction of natural resources, which can lead to overexploitation. Overfishing, deforestation, and soil degradation are all instances of how human activity, including study, may devastate ecosystems. The aim is to establish research approaches that do not compound these issues, but rather encourage sustainability. Depletion hazards can be mitigated by implementing conservation biology concepts and sustainable harvesting practices.

Climate Change and Uncertainty

Climate change has a substantial impact on the availability of ecological resources. For example, shifting rainfall patterns have an impact on the availability of water for research initiatives. Temperature variations affect plant growth and animal migration. To accommodate for these uncertainties, research designs must include flexible, adaptive procedures that can evolve as environmental conditions change.

Ethical Considerations

The utilization of natural resources in research must be conducted ethically. Issues such as the right to land and access to resources, indigenous knowledge systems, and the effects on local populations must be considered. Ethical research approaches include honoring local customs, guaranteeing equitable benefit sharing, and limiting environmental damage.

5. Future Directions and Opportunities:

Innovation in Ecological Resource Management:

The future of sustainable research depends in the development of new technologies and techniques that improve ecological resource management. New bioremediation strategies, such as employing plants or bacteria to clean

up polluted settings, are emerging. Smart agricultural research, such as precision farming, is boosting water efficiency and lowering chemical inputs.

Collaborative Research Models:

Sustainable research requires interdisciplinary collaboration. Partnerships between scientists, local communities, NGOs, and policymakers can ensure that research is both scientifically sound and practically applicable. Citizen science and co-management are models where local knowledge is integrated into formal research processes, ensuring that ecological resources are used sustainably.

Policy and Governance:

Research can help shape policies that promote sustainable resource usage. International frameworks, such as the United Nations Sustainable Development Goals (SDGs) and the Paris Agreement, offer standards for ecological study. Research on policy effectiveness helps shape global environmental governance and guarantees that ecological resources are preserved for future generations.

Conclusion


Ecological resources are essential for the success of sustainable research. Scientists and researchers can help to preserve ecosystems, use resources more sustainably, and mitigate environmental concerns by including ecological factors into research project design and implementation. To ensure the planet's long-term health, the research community must take a comprehensive approach that balances scientific advancement with environmental care. Sustainable research not only benefits the environment, but it also improves social fairness and economic resilience, laying the groundwork for a sustainable future.

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Overview of Bioinformatics: Definition, History, Key Areas, and Applications

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Bioinformatics – Definition

Bioinformatics is a dynamic and interdisciplinary scientific field that integrates biology, computer science, mathematics, and statistics to analyze, interpret, and manage biological data. It bridges the gap between experimental biology and computational analysis, enabling researchers to uncover insights into complex biological systems. By leveraging advanced computational tools and techniques, bioinformatics facilitates the exploration of vast datasets generated by modern biological research, such as genome sequencing, transcriptomics, proteomics, and metabolomics.

At its core, bioinformatics focuses on understanding biological processes at the molecular level. For example, it plays a crucial role in gene sequencing, where it helps to decipher the nucleotide sequences of DNA and RNA, aiding in the identification of genes and their functions. In protein structure analysis, bioinformatics tools predict and model the three-dimensional structures of proteins, which are essential for understanding their functions and interactions within biological pathways. Additionally, bioinformatics supports evolutionary biology by analyzing genetic and protein sequences to elucidate relationships among species and trace their evolutionary histories.

In essence, bioinformatics acts as a management information system for molecular biology, enabling the systematic organization, storage, and retrieval of biological data. This capability has far-reaching practical applications, including drug discovery, personalized medicine, agricultural biotechnology, and environmental science. For instance, bioinformatics helps identify potential drug targets by analyzing protein-ligand interactions and predicting the effects of genetic mutations. In agriculture, it contributes to crop improvement by studying genetic variations and enhancing traits like yield and disease resistance.

As a rapidly evolving field, bioinformatics continues to transform the landscape of biological research by providing innovative solutions for handling and interpreting complex biological data. Its interdisciplinary nature and diverse applications make it an indispensable tool in advancing our understanding of life sciences and addressing critical challenges in health, agriculture, and the environment.

History of bioinformatics

The history of bioinformatics is intertwined with the advancements in molecular biology, genetics, computer science, and information technology. Here's a chronological overview:

Early Developments (1950s-1970s): The advent of molecular biology and genetics set the stage for the development of bioinformatics. In the 1950s, the groundbreaking discovery of the DNA structure by Watson and Crick, along with the deciphering of the genetic code, provided a fundamental understanding of the molecular basis of life. This breakthrough laid the groundwork for further research, leading to the rise of computational methods in biology. By the late 1960s and early 1970s, early computational tools, including sequence alignment algorithms and protein structure prediction techniques, began to emerge, marking the beginning of bioinformatics as a field.

Birth of Bioinformatics (1980s): Bioinformatics emerged as a distinct field in the 1980s, driven by the rapid growth of genetic data resulting from DNA sequencing projects and the increasing need for computational tools to analyze and interpret this information. While the term "bioinformatics" was first coined in the early 1970s, it was in the 1980s that the field truly began to take shape, fueled by the surge of biological data from these sequencing initiatives.

Key milestones in this development included the creation of major genetic databases like GenBank (1982) and the European Molecular Biology Laboratory (EMBL) nucleotide sequence database (1982), which provided researchers with invaluable access to vast amounts of genetic data. To manage and analyze this data, computational tools and algorithms for tasks like sequence alignment, homology searching, and database management became indispensable, marking the beginning of bioinformatics as an essential discipline in biological research.

Rapid Expansion (1990s): The 1990s marked a period of rapid growth for bioinformatics, driven by significant advancements in DNA sequencing technologies and the completion of transformative projects like the Human

Genome Project (HGP). This era saw an explosion in the generation of biological data, leading to the development of advanced computational tools designed to manage, analyze, and interpret the ever-increasing volume of information.

During this time, bioinformatics expanded rapidly with the creation of more sophisticated algorithms and software for tasks such as sequence analysis, molecular modeling, and phylogenetic studies. The establishment of key institutions, such as the National Center for Biotechnology Information (NCBI) and the European Bioinformatics Institute (EBI), provided centralized resources and databases that became vital for researchers worldwide, offering access to biological data and powerful analytical tools. This period laid the foundation for bioinformatics as a cornerstone of modern biological and biomedical research.

Genomics Era (2000s): The Genomics Era of the 2000s was defined by groundbreaking advancements in genomic sequencing technologies, transformative discoveries in genomics, and the widespread application of genomic data across biological research, medicine, and other fields. Central to this era was the crucial role of bioinformatics in analyzing and interpreting the vast amounts of genomic data produced by the Human Genome Project (HGP) and subsequent genome sequencing efforts.

With the advent of high-throughput sequencing technologies, particularly next-generation sequencing (NGS), the volume of genomic data increased exponentially. This surge in data production necessitated the development of more sophisticated bioinformatics tools and algorithms for efficient data analysis, storage, and interpretation. Bioinformatics enabled researchers to decipher complex genomic information, identify genetic variations, and explore gene functions, contributing significantly to advancements in personalized medicine, disease research, and our overall understanding of genomics. This period marked the maturation of bioinformatics as a critical discipline in modern biology.

Integration with Systems Biology (2010s-present):

In recent years, bioinformatics has increasingly integrated with systems biology, with the goal of understanding biological systems as intricate networks of interacting molecules. This integration has led to the development of computational models and simulations that allow researchers to study biological processes at a systems level, combining data from genomics, proteomics, metabolomics, and other omics disciplines into cohesive frameworks.

The rise of big data and the application of machine learning techniques have further transformed bioinformatics, enabling the analysis of vast and complex biological datasets. These technological advancements have facilitated the discovery of novel insights into biological systems, including the identification of new biomarkers, drug targets, and mechanisms of disease. By leveraging machine learning and other AI-driven approaches, bioinformatics is unlocking previously uncharted areas of biology, offering exciting new possibilities for personalized medicine, disease prevention, and therapeutic development.

Need for bioinformatics

Bioinformatics serves as a universal language for expressing and interpreting disparate biological data, harnessing cutting-edge molecular biology and supercomputing capabilities to store, process, analyze, and simulate biological information. The wealth of data generated, particularly from genomics, presents both opportunities and challenges for pharmaceutical and biotechnology companies. While genomics offers a multitude of potential drug targets, the sheer volume of data overwhelms individual researchers. However, Bioinformatics companies are poised to assist in navigating this complexity. As the pharmaceutical industry seeks new avenues for growth, Bioinformatics companies are increasingly integral to drug discovery, offering essential tools and services. Originally emerging in the late 1980s to interpret sequence analysis data, Bioinformatics has evolved to encompass all facets of molecular biology. Market projections suggest significant growth in the Bioinformatics sector, with annual revenues estimated to reach \$110 million by 2004, and the potential to surpass \$5 billion in the next five years according to some forecasts.

The need for bioinformatics arises from the exponential growth of biological data generated by advancements in high-throughput technologies, such as genomics, transcriptomics, proteomics, and metabolomics. These technologies produce vast amounts of raw data, which require sophisticated computational methods for storage, management, analysis, and interpretation.

Bioinformatics provides the tools and techniques to analyze complex biological systems comprehensively, integrating data from multiple sources to gain insights into gene functions, regulatory networks, and biological processes. By leveraging computational algorithms and statistical models, bioinformatics enables researchers to extract meaningful information from large-scale datasets, uncover patterns, correlations, and associations, and generate testable hypotheses for further experimental validation.

In biomedical research and healthcare, bioinformatics is essential for understanding the genetic basis of diseases, identifying biomarkers for diagnosis and prognosis, and developing targeted therapies tailored to individual patients' genetic profiles. By analyzing genomic data, bioinformatics helps unravel the molecular mechanisms underlying diseases, identify potential drug targets, and predict drug responses, ultimately improving patient care and treatment outcomes.

In agriculture and environmental science, bioinformatics aids in crop improvement, biodiversity conservation, and environmental monitoring. By analyzing genomic data from plants and animals, bioinformatics guides breeding programs to develop high-yielding, stress-tolerant crop varieties and sustainable agricultural practices. In environmental monitoring, bioinformatics tools analyze DNA barcodes and metagenomic data to assess ecosystem health, track species distributions, and identify environmental contaminants, supporting biodiversity conservation efforts and ecosystem management strategies.

Overall, bioinformatics is indispensable in modern biological research and applications, facilitating data-driven discoveries, accelerating innovation, and addressing complex challenges in healthcare, agriculture, biotechnology, and environmental science. By integrating computational and biological approaches, bioinformatics empowers researchers to explore the intricacies of living systems, unravel the mysteries of life, and apply this knowledge to benefit society and the environment.

Key areas of bioinformatics

Genomics: Genomics is a comprehensive field of biology that focuses on the study of entire genomes, encompassing the complete set of genetic material within an organism. It involves processes such as genome sequencing, which determines the precise order of nucleotides in DNA, and genome assembly, where this data is organized into a complete genetic blueprint. Annotation plays a crucial role by identifying genes, regulatory regions, and functional elements within the genome, shedding light on gene expression and regulatory mechanisms. Comparative genomics extends this understanding by analyzing and comparing genomes across different species to uncover evolutionary relationships, conserved genes, and unique genetic traits. Through these efforts, genomics provides valuable insights into genetic variation, biological functions, and the molecular basis of diseases, driving innovations in medicine, agriculture, and biotechnology. This field continues to revolutionize our understanding of life by unraveling the complexities of genetic information.

Transcriptomics: Transcriptomics is the branch of molecular biology that focuses on studying RNA molecules to understand gene expression and its regulation in cells and tissues. This includes analyzing messenger RNA (mRNA), non-coding RNA, and splice variants to gain insights into how genes are transcribed and expressed under various conditions. By examining mRNA, transcriptomics reveals the active genes in a cell at a specific time, reflecting its functional state. The study of non-coding RNA highlights regulatory molecules that play crucial roles in gene silencing, RNA stability, and chromatin modification. Additionally, transcriptomics investigates alternative splicing events, where a single gene can produce multiple RNA variants, significantly expanding the diversity of proteins and functions encoded by the genome. This field provides a comprehensive view of the regulatory networks governing cellular processes, enabling a deeper understanding of development, disease mechanisms, and responses to environmental stimuli.

Proteomics: Proteomics is a comprehensive field dedicated to the large-scale study of proteins, which are essential molecules responsible for the structure, function, and regulation of cells and organisms. It involves identifying and quantifying proteins within a biological system, analyzing their structures, and examining their post-translational modifications (PTMs), such as phosphorylation, glycosylation, or ubiquitination. These modifications often play critical roles in regulating protein activity, stability, and interactions. Proteomics also investigates protein-protein interactions and interactions with other biomolecules like DNA, RNA, and small metabolites, offering insights into the intricate networks that drive cellular processes. By deciphering protein functions and their dynamics in different states, such as during health, disease, or environmental changes, proteomics provides a deeper understanding of biological systems. It also complements genomics and transcriptomics, bridging the gap between genetic information and phenotypic outcomes. Advances in proteomics have profound applications in drug discovery, biomarker identification, and personalized medicine, making it a cornerstone of modern biological research.

Metabolomics: Metabolomics is a field of study that focuses on the comprehensive analysis of small molecules, known as metabolites, within biological samples such as cells, tissues, or biofluids. These metabolites, which include sugars, amino acids, lipids, and other small organic compounds, serve as the end products of cellular processes and provide a direct reflection of an organism's metabolic state. By profiling and quantifying these molecules, metabolomics offers valuable insights into the biochemical pathways and processes that sustain life. This field plays a critical role in understanding how

metabolic pathways are regulated and how they respond to various stimuli, such as environmental changes, disease progression, or therapeutic interventions. It also helps identify specific metabolic signatures or biomarkers associated with physiological or pathological states, enabling the development of diagnostic tools and personalized treatment strategies.

Structural Bioinformatics: Structural bioinformatics is a branch of bioinformatics that focuses on the prediction, modeling, and analysis of the three-dimensional (3D) structures of biomolecules, such as proteins, nucleic acids, and small molecules. Understanding the 3D structure of these molecules is crucial because their shape directly influences their function and interactions within biological systems. Structural bioinformatics uses computational techniques to predict how biomolecules fold, how their structures relate to their biological activities, and how they interact with other molecules, including other proteins, DNA, RNA, and ligands.

Systems Biology: Systems biology is an interdisciplinary field that combines experimental and computational approaches to study complex biological systems in their entirety. Rather than focusing on individual components, systems biology seeks to understand how genes, proteins, metabolites, and other biomolecules interact and function together within a biological context. It aims to map the intricate networks of molecular interactions and regulatory pathways that govern cellular functions, development, and disease. At the core of systems biology is the concept of *emergent properties*, which refers to the idea that biological systems exhibit behaviors and characteristics that arise from the interactions of their parts, rather than from the properties of individual components.

Phylogenetics and Evolutionary Genomics: Phylogenetics is a field of biology that focuses on reconstructing the evolutionary relationships among species, genes, or other biological entities based on molecular data, such as DNA or protein sequences. By analyzing genetic similarities and differences, phylogenetics helps trace the lineage of organisms or genes, creating a "tree of life" that illustrates how species have evolved and diverged over time. Evolutionary genomics examines the genomic changes and processes of adaptation that occur across different evolutionary timescales. It delves into how genomes evolve, how genetic variations contribute to survival and fitness, and how organisms adapt to changing environments. Evolutionary genomics explores various aspects, such as the role of natural selection, genetic drift, and gene flow, in shaping the genome over time. By comparing genomic data from diverse species or populations, evolutionary genomics uncovers the molecular

basis of evolutionary processes, including the identification of conserved genes and regulatory pathways that are critical for adaptation.

Clinical and Medical Bioinformatics: Clinical and medical bioinformatics is a specialized field that applies computational techniques and tools to analyze and interpret a wide range of genomic, clinical, and healthcare data. The goal is to enhance patient care by providing insights that enable personalized medicine, more accurate disease diagnosis and prognosis, and improved drug development and treatment strategies. One of the primary focuses of clinical bioinformatics is personalized medicine, where genomic data, such as sequencing information, is used to tailor medical treatments to an individual's genetic makeup. This allows for more precise therapies, reducing the risk of adverse effects and improving treatment outcomes. By integrating genomic data with clinical and healthcare information, clinical and medical bioinformatics enables the development of more effective, targeted, and individualized treatments, ultimately advancing precision medicine and transforming patient care in modern healthcare systems.

Structural and Functional Genomics: Structural and functional genomics is an interdisciplinary field that combines genomic and functional data to elucidate the relationship between genetic variation and phenotypic traits. This field aims to bridge the gap between genotype and phenotype by studying how variations in the DNA sequence translate into observable traits, including disease susceptibility, drug response, and complex biological characteristics. Structural genomics focuses on the three-dimensional structures of proteins, nucleic acids, and other biomolecules, often using techniques like X-ray crystallography, NMR spectroscopy, and computational modeling. It aims to understand the architecture of the genome, including gene organization, regulatory elements, and functional sequences that govern cellular and molecular processes. Functional genomics complements structural genomics by studying gene expression, regulatory networks, and cellular interactions. It employs various high-throughput technologies, such as RNA sequencing (RNA-seq), proteomics, and metabolomics, to identify the functional roles of genes and their products in specific biological contexts. By linking genetic variation to phenotypic outcomes, functional genomics helps to identify genes associated with diseases, drug response, and other complex traits, providing insights into the mechanisms underlying these conditions. This integration of genomic and functional data allows researchers to uncover not only the genetic basis of health and disease but also the molecular pathways involved in drug action and resistance. Structural and functional genomics thus contribute to the development of precision medicine, where treatments and interventions can be

tailored based on an individual's genetic makeup, enhancing the effectiveness and minimizing side effects.

Environmental Bioinformatics: Environmental bioinformatics is a specialized branch of bioinformatics that applies computational tools and techniques to analyze biological data in the context of environmental factors. It focuses on understanding the interactions between living organisms and their environments, particularly how microbial communities and ecosystems respond to various ecological and environmental changes. By examining biological data, environmental bioinformatics helps in monitoring and managing environmental health, biodiversity, and sustainability. One key area of environmental bioinformatics is the study of microbial communities in different ecosystems, such as soil, water, or the human microbiome. Through data analysis, bioinformatics provides a deeper understanding of how contaminants spread through ecosystems and how they can be mitigated. Additionally, bioremediation is an area where environmental bioinformatics plays a critical role in identifying and utilizing microorganisms that can degrade pollutants, thereby cleaning up contaminated environments. By understanding the genes and pathways involved in biodegradation, bioinformatics aids in the development of more efficient bioremediation strategies. Overall, environmental bioinformatics is a powerful tool for addressing pressing environmental issues, from preserving biodiversity to managing pollution and improving sustainability practices in ecosystems.

Applications of bioinformatics

Genomic Sequencing and Annotation: Bioinformatics facilitates genomic sequencing by analyzing raw data from various sequencing platforms, assembling sequences, and annotating genes and functional elements. It enables accurate genome assembly, gene prediction, and functional annotation, providing insights into genetic variation, regulatory regions, and evolutionary relationships across different species. Comparative genomics tools aid in understanding genome structure, gene content, and evolutionary conservation, while variant analysis identifies genetic variations associated with phenotypic traits or diseases. Metagenomic analysis uncovers microbial diversity and functional potential in complex ecosystems, contributing to environmental and biomedical research. Overall, bioinformatics drives genomic sequencing and annotation, empowering diverse applications in biological research, medicine, agriculture, and environmental science.

Drug Discovery and Development: Bioinformatics accelerates drug discovery by identifying potential targets and lead compounds through computational

analysis of biological data. It aids in virtual screening and lead optimization by predicting compound interactions with target proteins and optimizing drug-like properties. Chemoinformatics and QSAR studies guide compound design, while ADME/Tox prediction assesses pharmacokinetic and safety profiles. Bioinformatics enables drug repurposing, personalized medicine, and pharmacogenomics by integrating genomic data to tailor therapies to individual patients' genetic profiles. Big data analytics and machine learning drive insights from large-scale datasets, while collaborative platforms and open data sharing foster innovation and knowledge exchange, collectively advancing drug discovery and development efforts.

Personalized Medicine: Bioinformatics plays a pivotal role in personalized medicine by integrating genomic and clinical data to tailor medical treatments to individual patients' genetic profiles. It enables the analysis of genetic variants associated with diseases, drug response, and adverse reactions, guiding treatment decisions and optimizing therapeutic outcomes. Pharmacogenomic studies identify genetic factors influencing drug metabolism, efficacy, and toxicity, facilitating the selection of appropriate medications and dosages for patients. Bioinformatics also supports risk prediction, disease prognosis, and preventive interventions based on genetic predispositions, empowering precision medicine approaches for improving patient care and outcomes.

Disease Diagnosis and Biomarker Discovery: Bioinformatics drives disease diagnosis and biomarker discovery by analyzing genomic, transcriptomic, and proteomic data to identify genetic variants, gene expression patterns, and protein biomarkers associated with diseases. Computational algorithms prioritize candidate biomarkers and assess their diagnostic or prognostic value, aiding in early detection, disease monitoring, and treatment response prediction. Integrative analyses uncover molecular signatures and pathways underlying disease pathogenesis, guiding the development of diagnostic assays and targeted therapies. Bioinformatics also facilitates the discovery of circulating biomarkers in biofluids and non-invasive diagnostic approaches, advancing precision medicine and personalized healthcare strategies for improved disease management and patient outcomes.

Comparative Genomics and Evolutionary Biology: Bioinformatics drives comparative genomics and evolutionary biology by analyzing genomic sequences across different species to elucidate evolutionary relationships, genetic diversity, and adaptive evolution. Computational methods identify conserved regions, gene families, and synteny, revealing evolutionary patterns and functional conservation. Phylogenetic reconstruction tools reconstruct

evolutionary trees and infer ancestral relationships based on molecular data, providing insights into speciation events, lineage divergence, and evolutionary dynamics. Comparative genomics also highlights genomic innovations, gene gains and losses, and adaptive changes underlying species divergence and adaptation to diverse environments, fostering a deeper understanding of evolutionary processes and biodiversity conservation.

Functional Genomics and Systems Biology: Bioinformatics facilitates functional genomics and systems biology by integrating genomic, transcriptomic, proteomic, and metabolomic data to elucidate gene functions, regulatory networks, and biological processes at a systems level. Computational methods analyze high-throughput data to identify genes, pathways, and regulatory elements associated with cellular functions and phenotypic traits. Network-based approaches model molecular interactions and signaling pathways, uncovering emergent properties and system-level behavior. Functional genomics tools assess gene expression, protein abundance, and metabolite profiles under different conditions, providing insights into cellular responses, disease mechanisms, and drug targets. Systems biology frameworks integrate multi-omics data to characterize complex biological systems comprehensively, advancing our understanding of cellular physiology, disease pathogenesis, and therapeutic interventions.

Phylogenetics and Taxonomy: Bioinformatics drives phylogenetics and taxonomy by analyzing molecular data, such as DNA or protein sequences, to reconstruct evolutionary relationships and classify organisms into taxonomic groups. Computational algorithms infer phylogenetic trees and ancestral relationships based on sequence similarity and evolutionary distance, providing insights into species divergence, speciation events, and evolutionary history. Phylogenetic analyses inform taxonomy by defining evolutionary relationships among taxa, resolving taxonomic uncertainties, and identifying cryptic species. Bioinformatics tools aid in biodiversity assessment, species conservation, and ecosystem management by characterizing microbial diversity, tracking invasive species, and studying evolutionary patterns across diverse taxonomic groups.

Biomedical Research and Clinical Trials: Bioinformatics plays a vital role in biomedical research and clinical trials by analyzing high-throughput data from genomics, transcriptomics, proteomics, and clinical studies to uncover molecular mechanisms underlying diseases, identify biomarkers for diagnosis and prognosis, and evaluate therapeutic interventions. Computational methods integrate multi-omics data to characterize disease signatures, identify potential drug targets, and stratify patient populations for personalized treatment

strategies. Bioinformatics facilitates the design and analysis of clinical trials by optimizing study protocols, identifying patient cohorts, and evaluating treatment efficacy and safety outcomes. Collaborative platforms and data sharing initiatives enable knowledge exchange and reproducibility, accelerating translational research and precision medicine applications in healthcare.

Agricultural Genomics and Crop Improvement: Bioinformatics revolutionizes agricultural genomics and crop improvement by analyzing plant genomes, identifying genes associated with agronomic traits, and developing molecular markers for breeding programs. Computational tools predict gene functions, regulatory elements, and genetic variations underlying traits such as yield, disease resistance, and stress tolerance. Genome-wide association studies (GWAS) and marker-assisted selection (MAS) leverage genomic data to accelerate breeding efforts, improve crop productivity, and develop climate-resilient varieties. Bioinformatics also facilitates comparative genomics and evolutionary studies to understand crop domestication, genetic diversity, and adaptation to diverse environments, enabling sustainable agriculture practices and food security initiatives.

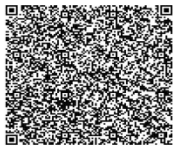
Environmental Monitoring and Biodiversity Conservation: Bioinformatics contributes to environmental monitoring and biodiversity conservation by analyzing DNA barcodes, tracking species distributions, and studying microbial communities in ecosystems. Computational methods analyze environmental DNA (eDNA) to identify species presence, abundance, and diversity, providing insights into ecosystem health and biodiversity hotspots. Metagenomic analysis uncovers microbial diversity and functional potential in soil, water, and air samples, informing environmental management and conservation strategies. Bioinformatics tools assess the impact of human activities, climate change, and habitat degradation on biodiversity, guiding conservation efforts and ecosystem restoration initiatives for preserving Earth's ecosystems and biodiversity.

Conclusion

In conclusion, bioinformatics is an indispensable field that bridges the gap between biology and computational analysis, facilitating the interpretation of vast biological datasets. Its evolution from the late 1980s, driven by advancements in molecular biology, genetics, and information technology, has led to its current role in genomics, transcriptomics, proteomics, and more. With its ability to process and analyze complex biological data, bioinformatics plays a critical role in numerous applications, including drug discovery, personalized medicine, disease diagnosis, agriculture, and environmental conservation. As

Emerging Research Concept in Life Sciences

biological data continues to grow in scale and complexity, the role of bioinformatics will only become more central to scientific advancement and innovation.

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Microbiome-based therapeutics: Opportunities and challenges

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Introduction

The term "**human microbiome**" refers to the trillions of bacteria and their genomes in the human body, predominantly prevalent in the gastrointestinal system and on the skin. Microorganisms have a role in almost every biological system, for instance, in supporting metabolic functions, warding off pathogens and interacting with the immune system.

Various disorders can be caused by imbalances in the microbiome, called **dysbiosis**. This can be brought on by factors including viruses, antibiotics and unhealthy diets. Hence, microbiome – based treatments have the potential to treat a wide range of symptoms, with mounting evidence that focus on and modify the microbiome. Microbiome could enhance human health and treat more than 25 conditions across different therapeutic areas. These include infectious diseases, especially *Clostridioides difficile* infections, and autoimmune diseases such as Inflammatory Bowel Disease (IBD), Ulcerative colitis, Crohn's disease, and even cancer.

In recent years microbiome – based therapeutics, also referred to as microbiome modulators, have made rapid advances, with several promising candidates moving forward through the clinical pipeline. Two breakthrough approvals in the past twelve months have been Seres Therapeutics' Rebyota®, which got the green light from the US Food and Drug Administration (FDA) in November 2022, and Ferring Pharmaceuticals' VOWST™, which was approved in April 2023. Both products promise to offer safe and effective treatments for patients with recurrent *Clostridioides difficile* infection (rCDI), which in the US alone is a billion dollar market.

In this rapidly evolving landscape, Pharma and Life Sciences (PLS) companies and investors are focused on microbiome – based therapeutics. This chapter deals the accelerating rise of microbiome based therapeutics. Current research on the human microbiota has become much more sophisticated and more comprehensive. Therefore, we propose that research should focus on the host-microbe interaction and on cause-effect mechanisms, which could pave the way to an understanding of the role of gut microbiota in health and disease, and provide new therapeutic targets and treatment approaches in clinical practice.

The human microbiota in health

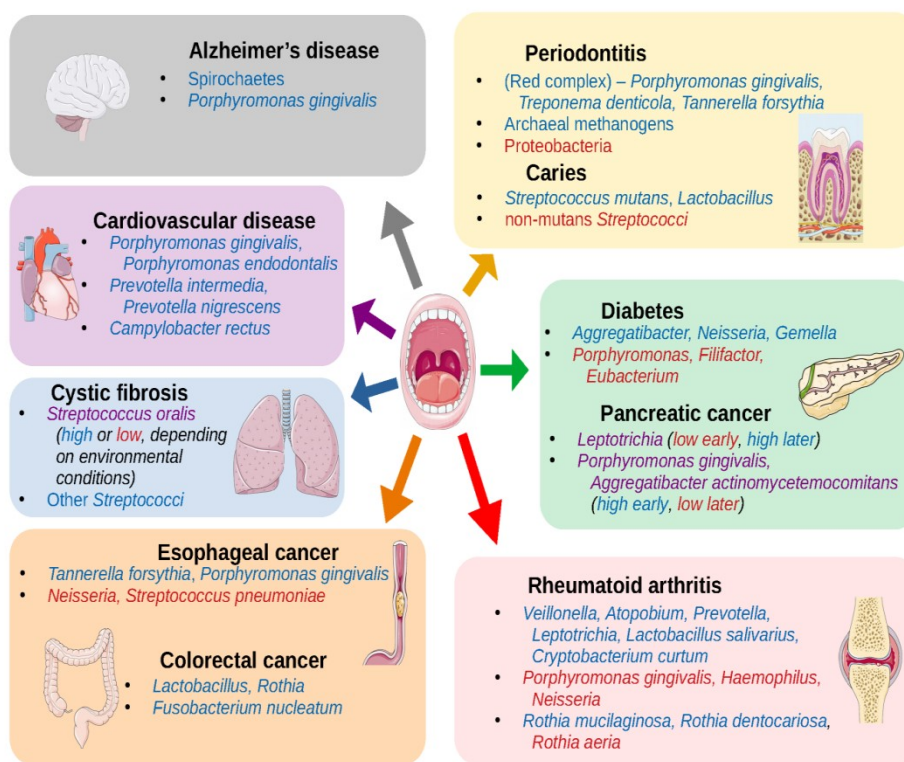
The human microbiota affects host physiology to a great extent. Trillions of microbes colonize the human body, including bacteria, [archaea](#), viruses, and eukaryotic microbes. The body contains at least 1000 different species of known bacteria and carries 150 times more microbial genes than are found in the entire human genome. Microbiotic composition and function differ according to different locations, ages, sexes, races, and diets of the host.

Commensal bacteria colonize the host shortly after birth. This simple community gradually develops into a highly diverse ecosystem during host growth. Over time, host-bacterial associations have developed into beneficial relationships. Symbiotic bacteria metabolize indigestible compounds, supply essential nutrients, defend against colonization by opportunistic pathogens, and contribute to the formation of intestinal architecture. For example, the intestinal microbiota is involved in the digestion of certain foods that cannot be digested by the stomach and small intestine, and plays a key role in maintaining energy homeostasis. These foods are primarily dietary fibers such as xyloglucans, which are commonly found in vegetables and can be digested by a specific species of *Bacteroides*. Other non-digestible fibers, such as fructo – oligosaccharides and oligosaccharides, can be utilized by beneficial microbes, such as *Lactobacillus* and *Bifidobacterium*. Studies have clarified the role of the gut microbiota in lipid and protein homeostasis as well as in the microbial synthesis of essential nutrient vitamins. The normal gut microbiome produces 50–100 mmol·L⁻¹ per day of short-chain fatty acids (SCFAs), such as acetic, propionic, and butyric acids, and serves as an energy source to the host intestinal epithelium. These SCFAs can be quickly absorbed in the colon and serve many diverse roles in regulating gut motility, inflammation, glucose homeostasis, and energy harvesting. Furthermore, the gut microbiota has been shown to deliver vitamins to the host, such as folates, vitamin K, biotin, riboflavin (B₂), cobalamin (B₁₂), and possibly other B vitamins. A

previous study demonstrated that B₁₂ can be produced from delta-aminolevulinate (ALA) as a precursor.

In addition, gut-colonizing bacteria stimulate the normal development of the humoral and cellular mucosal immune systems. The signals and metabolites of microorganisms can be sensed by the hematopoietic and non-hematopoietic cells of the innate immune system and translated into physiological responses. Studies comparing normal mice with GF mice have found that GF mice show extensive defects in the development of gut-associated lymphoid tissue and antibody production. A report has also demonstrated that the gut microbiota generates a tolerogenic response that acts on gut dendritic cells and inhibits the type 17 T-helper cell (Th17) anti-inflammatory pathway.

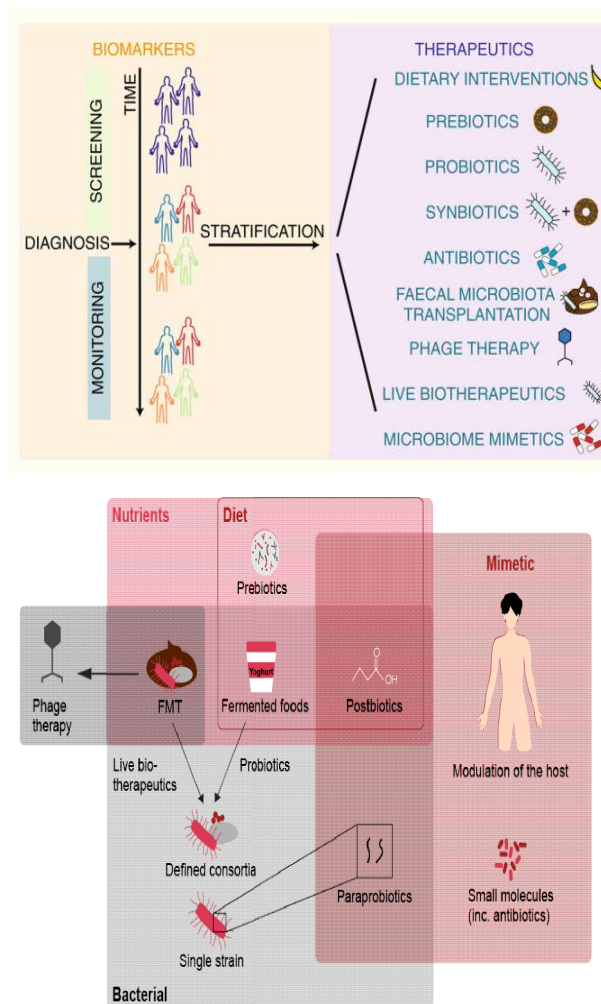
Figure: 1: Microbiota in health and disease



Microbiome based therapeutic approaches and driving development

The earliest reported basic research on using bacteria to treat disease was in 1891, when William Coley, a surgical oncologist at New York City's Hospital for Special Surgery developed a mixture containing toxins filtered from the dead bacteria of *Streptococcus pyogenes* and *Serratia marcescens* to treat cancer. However, it was not until the late 20th century that the rise of sequencing technologies and synthetic biology intensified systematic microbiome research.

Figure 2 Classification of microbiome-based therapeutics Gulliver, E. L. *et al.* (2022)



Between 2007 and 2012, the EU-funded Metagenomics of the Human Intestinal Tract (MetaHIT) programme marked a new stage in collaborative research, involving 15 institutes from eight countries. The program explored the genetic potential of human microbial companions and their impact on health and wellbeing, significantly increasing our understanding of the human microbiome importance, way beyond the digestive system

Drawing on these insights and advances in modern synthetic biology, researchers have been able to create microbial strains with unique and complex characteristics. These include altered metabolic pathways and new capabilities such as secretion of specific compounds. The examples are listed below

Table 1: Merits of microbiome based therapeutic approaches

	Therapeutic	Advantages	Disadvantages	Future directions/implications
Probiotics	Live microorganisms which, when administered in adequate amounts, confer a health benefit on the host	<ul style="list-style-type: none">• Relatively safe• Readily available as standardised mix	<ul style="list-style-type: none">• Not targeted to a disease or patient• Dependant on specific microbe colonisation• Dependant on gut micro-environment• Therapeutic response temporary• Viability not requirement of regulatorc	Efficacious following antibiotics and in the prevention of NEC. Potential as non-specific treatments to increase bacterial diversity
Synbiotics	Probiotics that are synergistically combined with prebiotics (components in food supporting beneficial bacteria)	<ul style="list-style-type: none">• Relatively safe• Includes all components for efficacy	<ul style="list-style-type: none">• Therapeutic response temporary• Require a specific gut micro-environment• Potential adverse responses (e.g. post-antibiotics)	Efficacious in the treatment of metabolic diseases. Further combinations should be explored for the treatment of other diseases

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Antibiotics	Substances that destroy or slow down growth of bacteria	<ul style="list-style-type: none"> • Safe • Cheap • Approved medication • Existing regulatory framework 	<ul style="list-style-type: none"> • Potential off-target effects (antibiotic resistance, disruption of colonisation resistance) • Limited to disruption of the microbiota 	Examination for use in targeted microbiome manipulation; however, caution is required to avoid off-target, adverse effects
Phage therapy	Viruses that exclusively infect bacterial cells	<ul style="list-style-type: none"> • Highly specific 	<ul style="list-style-type: none"> • Limited to disruption of the microbiota • Targets require specific development • Emerging therapy 	Examination for use in altering microbiome structure due to their highly specific nature
FMT	FMT (Faecal Microbiota Transplantation) can restore bacterial diversity and health-associated functions such as colonization resistance by introducing a faecal-associated microbiota from a healthy individual	<ul style="list-style-type: none"> • Contains all microbes and nutrients • Proven efficacious for <i>Clostridioides difficile</i> treatment 	<ul style="list-style-type: none"> • Donor variability • Requires rigorous pre-screening • Efficacy only seen for some conditions • Some administration costly • Inability to standardise composition 	Further work is required to determine causality in FMT treatment. This will allow for FMT to be considered for the treatment of other diseases
Live Biotherapeutics	Live organisms (such as bacteria) that are applicable to prevention, treatment or	<ul style="list-style-type: none"> • Approved for specific indications 	<ul style="list-style-type: none"> • Requires maintenance of bacterial viability • Potential adverse long-term health effects • Difficulty determining causal 	Determination of causality required to allow for development

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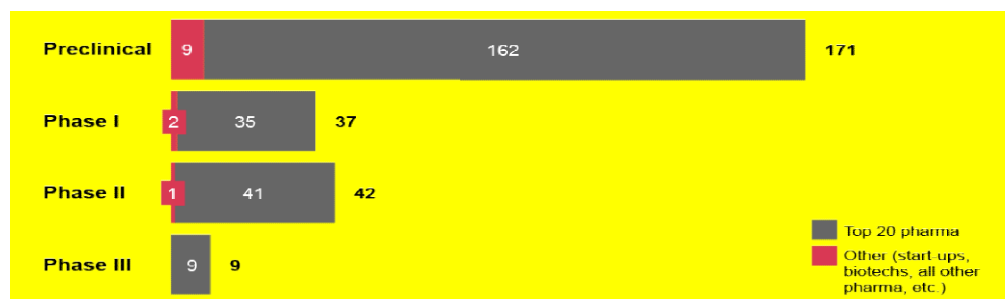
	cure of a disease		relationship	
Microbiome mimetics	Microbiome mimetics describes any intervention that replicates the interaction between the microbiome and the host, that yields a therapeutically beneficial outcome. This can include bacterial derived products, small molecules, conventional therapeutics or host derived products	<ul style="list-style-type: none"> Not reliant on current microbiome state 	<ul style="list-style-type: none"> Limited research to develop mimetics 	More research required to identify candidates as mimetics and mechanisms of delivery, including diet, should be explored

Source: Gulliver, E. L. et al. (2022). Review article: the future of microbiome-based therapeutics. *Aliment Pharmacol Ther.* 2022;56:192–208

Fecal microbiota transplantation (FMT) and live biotherapeutics are currently attracting the most attention of pharma companies in the prescription-based market. Besides the two recent breakthrough approvals mentioned above, we see promising opportunities in advanced FMT assets in ongoing Phase III clinical trials conducted by MaaT Pharma for the treatment of Graft-versus-host disease and by Mikrobiomik Healthcare for the treatment of *Clostridioides difficile* infections. Advanced live biotherapeutics in Phase III clinical trials include therapeutics targeting conditions such as Hyperoxaluria (OxThera) and chronic renal failure (Kibow Pharmaceuticals). Currently, microbiome R&D is driven by small and medium-sized companies. Among the more than 100 companies actively developing a total of 243 microbiome

modulator candidates as of August 2023, only five of top 20 pharma companies are represented. Furthermore, looking at investment deals for microbiome assets since 2015, start-ups and biotechs have been the technological originators in 56 percent of the cases, with Big Pharma only accounting for 3 percent

Figure 3 Number of microbiome-based therapeutics candidates at each clinical stage



Source: Citeline Pharma Intelligence – Pharmaprojects

Potential market opportunity for microbiome-based therapeutics

Prescription-based products

The commercial market for microbiome-based treatments is still at an early stage, with widely varying assessments of its global value. Strategic Market Research estimates that the prescription-based market was only worth \$115 million in 2021, but forecasts that it will reach \$1.3bn by 2030. Another report from Research and Markets predicts that the worldwide microbiome market will increase from \$390 million in 2022 to \$570 million by the end of 2023.

Despite these variations, all market size estimates for microbiome-based therapeutics are rather small compared with other major therapeutics markets such as oncology. However, such estimates usually do not consider the potential for microbiome-based therapeutics to replace conventional therapeutics in healthcare. This is a likely scenario has given the expectation of microbiome-based prescription products to be a global multi-billion-dollar market by 2030.

Non-prescription-based products

These products have a far longer history focused on more consumer-oriented products such as probiotic supplements for gut health, which dominate the sector. The market is still substantially larger than for prescription-based products and more mature, as indicated by moderate forecast growth. For example, one estimate predicts the market will increase by a Compound Annual Growth Rate (CAGR) of 6.4 percent in the next four years to be worth \$85.4bn by 2027.

Major market trends for microbiome-based therapeutics

Three major trends are driving the increasing market potential of microbiome-based therapeutics.

1. Applications beyond the digestive system

Probiotics and prebiotics in food and food supplements have long been known to be beneficial for health. As far back as 1917, for example, the probiotic Mutaflor was commercialized, mostly for treating digestive disorders. In recent decades, advances in genomic sequencing technologies and research milestones like the MetaHIT project have opened up the market for microbiome-based therapeutics beyond the digestive system.

One emerging trend that has seen several new R&D partnerships involving some leading PLS names this year is research related to skin microbiome-based therapeutics. Johnson & Johnson is working with Sequential Skin Inc. to develop new methods for non-invasive genomic-based skin testing. Meanwhile, Unilever has partnered with the University of Liverpool's Microbiome Innovation Centre and Brain and Behaviour Laboratory to research the link between changes in the skin microbiome and wellbeing over time, and find new product and technology approaches

The potential market for skin microbiome products has attracted other biotechs and mid-sized pharma companies. One example is Boston-based AOBiome, which is entirely focused on skin microbiome therapeutics. The company currently has multiple ongoing phase II clinical trials for treating conditions such as Atopic Dermatitis, Acne and Rosacea

2. More clinical trials

The intensification of basic research and clinical development can be seen in the increasing number of academic articles on the microbiome globally, from 1,177 in 2010 to 20,867 in 2020. In the same period, the number of microbiome-related clinical studies in the US increased from 31 to 632. The

US is the leading region for microbiome R&D, partly because the FDA has issued clear guidelines that therapeutics are regulated as a drug, in contrast to Europe, where national regulators generally regard it more vaguely as a drug "or something else". The FDA's tighter definition means the US is a more popular location for clinical studies.

3. Increasing consumer awareness

Beyond treating symptoms, maintaining good health and preventing illness are increasingly important priorities for today's consumers, as we highlighted in a recent report "From Healthcare to Life Care". Most current microbiome-based therapeutics is consumer products such as probiotics in the form of yoghurt drinks and supplements. One market study in 2020 suggests that 52 percent of global consumers purchased probiotics in the previous 12 months with the goal of improving their digestive health and ensuring they had enough "good" bacteria in the body

Challenges in microbiome based therapeutics

We believe the following challenges demand the most urgent attention:

- **Evidence generation**

Microbiome therapeutics do not have the same pharmacokinetic profile as conventional "non-living" therapeutics. In addition, each individual's microbiome has unique characteristics due to several factors such as genetics and differences in diet. Academic and clinical studies must account for this to ensure that they are representative of large patient populations and generate sufficiently reliable data to meet robust safety and efficacy standards.

- **Technology for R&D and manufacturing**

R&D in microbiome therapeutics relies heavily on both sequencing and bioinformatics capabilities to generate genetically modified microbes with therapeutic characteristics, but it is still rare to find companies that are good in both areas. On the production side, manufacturing and quality control processes require specialized expertise and infrastructure that most contract development and manufacturing organizations (CDMOs) do not yet have to a sufficient degree, and therefore struggle to maintain the microorganism's viability throughout the entire manufacturing process.

- **Local regulatory challenges**

The contrast between National regulators in the EU, which individually decide if microbiome-based therapies should be classified as a drug or not, and

the US FDA, which narrowly defines them as drugs, has created uncertainty for PLS companies and investors, especially in Europe. However, a recently proposed EU directive for medicinal products covering microbiome-based therapies, which was proposed in April 2023, may bring more clarity. If passed, it could encourage more marketing approvals for microbiome-based therapies in the EU, and by extension in other non-member European countries such as the UK and Switzerland.

• Need for stakeholder education

The rapid pace of innovation in microbiome-based therapies means stakeholders, from healthcare providers and regulators to investors and patients often do not fully understand the nuances of different approaches. PLS companies should educate their target customers and clearly convey a therapy's mechanism of action and value proposition to encourage its adoption and realize its full market potential.

To overcome these challenges, carefully designed, large-scale clinical trials that generate robust, objective data on patient outcomes, combined with an increasing body of real-world evidence, will likely help convince stakeholders about a therapy's efficacy and encourage regulatory clarity and consistency across markets. In addition, the proliferation of microbiome R&D partnerships between industry, academic institutions and governments can help continue to boost both specialist knowledge and public awareness about microbiome-based therapies. The crucial roles of the human microbiota should be investigated at a much deeper level, and microbiome-based diagnosis and treatment strategies will be used for future personalized medicine work.

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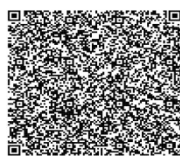
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Almond peel: A bioactive compounds and the waste to wealth approach

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Abstract

Almonds (*Prunus dulcis*) rank among the most abundant natural food sources of vitamin E and polyphenols. The scope of the nut industry, as well as the significant biomass of these skins, must be reconsidered. These skins exhibit a high antioxidant capacity and are abundant in phenolic compounds and dietary fibre. Historically, nut skins and other by-products have been used for animal feed and as raw materials for energy generation. However, recent investigations have confirmed that they are an affordable and valuable source of natural antioxidants for nutraceutical and pharmaceutical applications. Nuts' skin contains more antioxidants, including phenolic compounds. As a result, they can be potentially applied in food valorization, food protection, and as pharmaceutical or colouring agents. Notably, the skins of almonds present a promising source of bioactive compounds, including vitamins, free amino acids, minerals, and polyphenols, particularly flavanols. These nuts and its byproducts possess antioxidant, anti-inflammatory, and cardioprotective properties, primarily due to their rich content of vitamin E and unsaturated fatty acids. They help regulate glucose, potentially reducing type-2 diabetes, cholesterol, blood pressure. Furthermore, almond by-products present a challenge in waste management; however, they can be transformed into

revalorized products or ingredients, thereby reducing waste and minimizing environmental impact. Many of these by-products have been analysed, revealing various bioactive properties, including phenolic compounds that are significant in the prevention of degenerative diseases and other health conditions. In order to enhance the value of almond by-products and mitigate both economic and environmental challenges, it is possible to extract bioactive compounds or utilize the by-products directly as food ingredients.

Keywords: Almond, Almond peel, By-product valorization, Nutritional benefits and functional benefits.

Introduction

The almond tree, scientifically known as *Prunus amygdalus*, or *Amygdalus communis* L., is one of the most extensively cultivated nut trees around the world, especially in dry regions, due to its robustness and ability to withstand water shortages and insufficient irrigation. Over the past decade, there has been a notable rise in nut consumption, largely fueled by heightened awareness of their potential health benefits. This surge in nut intake is associated with a clear increase in the production of by-products and waste.

Nut peels, frequently regarded as waste, are in fact a rich source of nutrients and bioactive compounds. They are abundant in dietary fibre, antioxidants, vitamins, and minerals, which can greatly enhance the nutritional profile of various food products. By integrating these peels into items such as baked goods, snacks, beverages, and dairy products, we not only augment their health benefits but also foster sustainable practices by minimizing food waste. The principles of waste reduction and the valorization of by-products are fundamental to achieving effective management and improved sustainability within the food industry.

The circular economy is based on the theories and principles of industrial ecology, which seeks to establish a closed-loop system for materials and substances, thereby reducing resource consumption and environmental emissions. Within industrial ecology, the notion of industrial metabolism highlights the operation of industrial systems in a manner similar to that of natural ecosystems. This model of the circular economy is fundamentally restorative and aims to replicate natural processes by improving and optimizing systems. Furthermore, utilizing nut peels can provide economic advantages for food producers by generating new revenue opportunities.

The popularity of bakery products is on the rise, attributed to their improved nutritional attributes. By incorporating healthful and medicinal plant-

based components, the industry can adeptly meet consumer demands. Fortifying essential cereal items with these ingredients presents a practical strategy for fulfilling nutritional needs and enhancing overall health. Therefore, it can be concluded that repurposing nut peels is a wise and eco-friendly strategy for optimizing the advantages of nuts while promoting a more sustainable food system.

Almond peel as a functional ingredient:

The substantial consumption of dried fruits is evidently associated with an increased generation of by-products, which include the shell, skin, and hull resulting from the cleaning and processing of these fruits. In almonds, the portion that is consumable constitutes approximately 24% of the total fruit, indicating that 76% is considered waste.

➤ To advance sustainable agricultural and food systems, valorization of Almond skins which account for 70 to 100% of the total phenolic content found in the entire almond nut. Numerous studies have indicated that these skins possess advantageous phytochemical properties, serving as an excellent source of phenolic compounds.

➤ There natural compound has the potential to be utilized in the development of nutritional supplements that offer improved health benefits. Almond skins are a valuable functional ingredient due to their high content of bioactive phenolic compounds, such as flavonoids and phenolic acids.

➤ Utilizing the byproduct of almond after processing by value adding new products. Therefore, the by-products of almonds are currently a concern in terms of waste management.

➤ By creating a sustainable system by repurposing of food industry by products that reduce environmental harm and resource wastage is crucial. Enhance sustainability in the food sector through the principles of a circular economy. They can potentially be repurposed into revalorized products or ingredients, which would help reduce their presence and mitigate their environmental effects

Therapeutic potential of almond peel:

In recent years, the demand for food products with added value has expanded beyond simply satisfying hunger or providing nutrients in low concentrations, as the connection between diet and health has been thoroughly demonstrated. Consequently, the term "fortified" has shifted to "functional," reflecting the trend of incorporating specific components into food products that provide physiological benefits and lower the risk of diseases.

Food technologists are increasingly focused on enhancing the nutritional profile of products by integrating healthier components, including dietary fibres, vitamins, minerals, and essential oils. The consumption of bioactive compounds, such as antioxidants, total phenols, flavonoids, and flavonol, plays a crucial role in promoting health and safeguarding against diseases like cancer, cardiovascular issues, and various other degenerative conditions.

The skin of almonds, also known as the seed coat, can be removed and discarded following the blanching process. The proximate composition of almond skin reveals that total dietary fibre content is 47.5 g/100 g for blanched skin and 45.1 g/100 g for natural skin. Additionally, the soluble dietary fibre amounts to 2.7 g/100 g in blanched skin and 3.8 g/100 g in natural skin. For blanched skin, the lipid content is noted to be 22.2 g per 100 g, whereas the other sample shows a lipid content of 24.2 g per 100 g for natural skin, while the protein levels are 12.8 g/100 g in blanched skin and 10.3 g/100 g in natural skin.

Almonds and their derivatives have been recognized for their prebiotic properties, particularly almond skins, which contribute to the enhancement of intestinal microbiota diversity and the overall improvement of gastrointestinal function. Research indicates that the consumption of almonds, including their skins, leads to an increase in β -galactosidase activity, a marker associated with beneficial colonic bacteria like bifidobacteria and lactobacilli. This effect is significant in enhancing carbohydrate metabolism in individuals suffering from Long-term health issues including Crohn's disease and ulcerative colitis.

Furthermore, polyphenols derived from almond skins exhibit synergistic effects when paired with vitamins C and E, as they offer protection against LDL oxidation and enhance antioxidant defences. In addition to polyphenols, almond skins contain varying amounts of triterpenoids, particularly betulinic acid, oleanolic acid, and ursolic acid, which have been recognized for their anti-inflammatory, anticancer, and antiviral properties, particularly against the human immunodeficiency virus (HIV).

Polyphenols found in almond skin exist primarily as flavonoids in both aglycone and glycoside forms, with aglycones demonstrating superior radical scavenging capabilities compared to their glycoside counterparts. The predominant phenolic compounds in almond skin include flavanols and flavonol glycosides. Among the flavanols, catechin and epicatechin are the most prevalent, while isorhamnetin-3-O-rutinoside and kaempferol-3-O-rutinoside are the key flavonol glycosides identified.

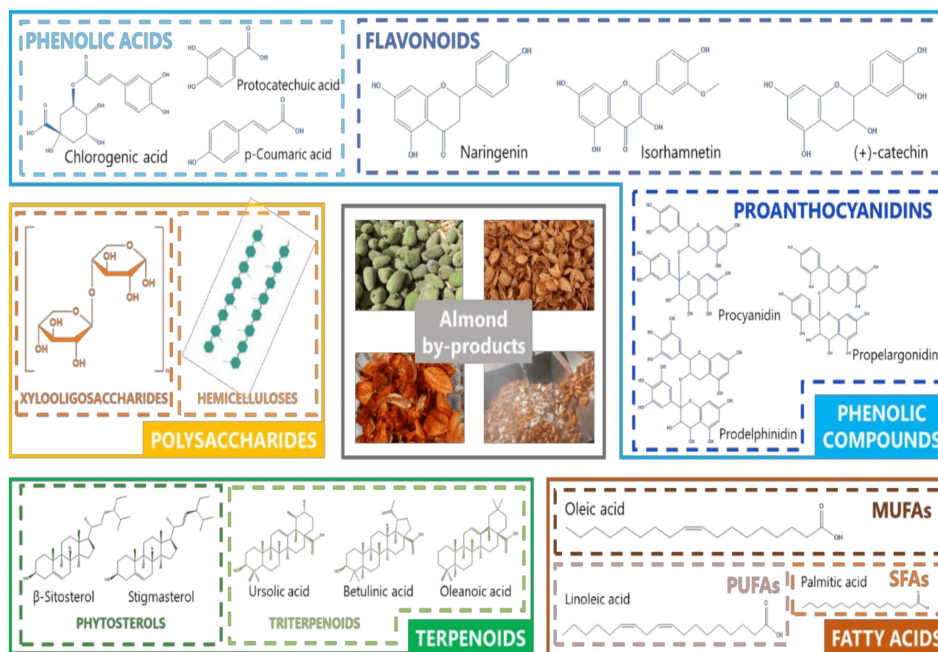


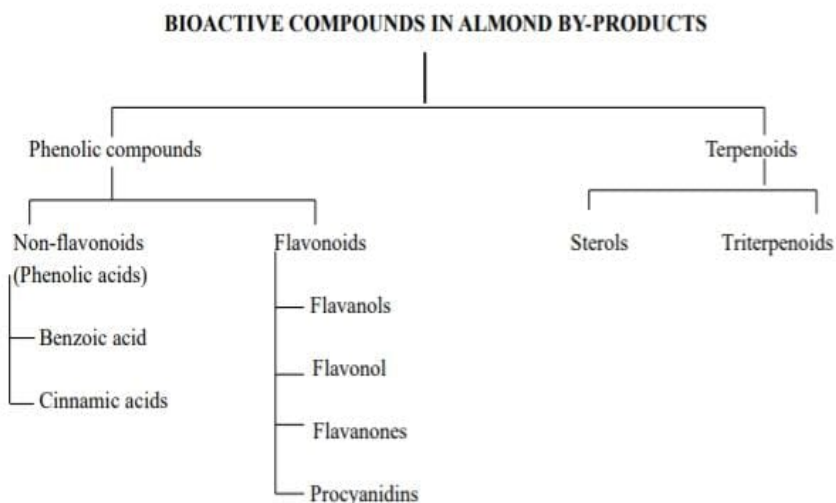
Fig 1., Phytochemical components Source: (Garcia-Perez *et al.*, 2021)

Bioactive potential of almond peel:

A total of 31 phenolic compounds were identified, which included flavanols (constituting the largest proportion at 33.0–56.0% of the overall phenolic content), flavonol glycosides, hydroxybenzoic acids and aldehydes, flavonol aglycones, flavanone glycosides, flavanone aglycones, hydroxycinnamic acids, and dihydroflavonol aglycones. The primary flavonoids identified in almond skin include catechin, epicatechin, kaempferol, and isorhamnetin, which may exist in the forms of 3-O-rutinoside or 3-O-glucoside.

It was stated that isorhamnetin, in its 3-O-glucoside or 3-O-rutinoside forms, constitutes around 70.0% of the overall flavonoid content is found in different almond varieties, with a measurement of $97.1 \pm 2.3\%$ of these

compounds sourced from the skin. In a study, catechin was found to account for 10.0–23.0% of the identified phenolics, while isorhamnetin-3-O-rutinoside represented 6.8–17%. The concentration and presence of each phenolic compound in almond skin may vary based on factors such as the industrial processing methods employed, storage conditions, and extraction techniques. The main phenolic acids, which are non-flavonoids, found in numerous plant-based foods are derivatives of hydroxybenzoic and hydroxycinnamic acids. In almonds, derivatives of cinnamic acids, including caffeic, ferulic, p-coumaric, and sinapic acids, have been identified, alongside hydroxybenzoic acids such as protocatechuic, p-hydroxybenzoic, and vanillic acid, as well as chlorogenic acid as the most abundant hydroxycinnamic acid in almond skin.



Almond skins are acknowledged as a possible functional component in food products because of their antioxidant polyphenols and prebiotic Fiber. Pharmacological research has demonstrated that *P. dulcis* exhibits a range of biological activities, including prebiotic, antimicrobial, antioxidant, anti-inflammatory, anticancer, laxative, hepatoprotective, cardiometabolic protective, nootropic, anxiolytic, sedative, hypnotic, and neuroprotective properties.

Bioactivity of almond peel:

Antioxidant activity:

The polyphenolic profile of almond skin consists of flavonoids, phenolic acids, and proanthocyanins. Moreover, the importance of polyphenolic and polysaccharides compounds and the antioxidant and antiproliferative effects have been emphasized. Flavonoids are considered potent antioxidants, functioning by neutralizing reactive oxidants, binding transient metals, and/or enhancing the body's natural antioxidant defence systems.

The significant polyphenol content in almonds and their derivatives demonstrates strong antioxidant properties, which operate through various mechanisms. These include the scavenging of free radicals, the induction of antioxidant enzymes, the modulation of genetic responses related to antioxidants, and the regulation of biomarkers associated with oxidative stress and lipid peroxidation. In the context of almond products, research has primarily focused overall on the evaluation of seeds, skins, husks, and blanching water was conducted to determine their antioxidant activity through in vitro and biochemical testing.

Polyphenols, as widely distributed secondary metabolites, are commonly found in high concentrations in seed coats or skins acting as phytoalexins to safeguard nutrients in the seed. The kernel undergoes processes of oxidation and microbial activity prior to germination. Numerous meta-analytic studies consistently demonstrate that flavonoid-rich foods can improve cardiovascular disease risk factors and decrease overall mortality rates. The benefits of flavonoids for health are significant & attributed to their diverse biological actions, including antioxidation, anti-inflammation, vasodilation, anti-hypertension, platelet function regulation, insulin sensitization, and cholesterol reduction.

Anti-inflammatory activity:

The polyphenols found in almond skins are readily absorbed by the body and exhibit antioxidant properties both in laboratory settings and in living organisms, effectively reducing oxidative stress in the blood. Additionally, these bioactive compounds in almond skins have been suggested for use in treating intestinal inflammatory diseases. Furthermore, almond skins are a valuable source of Fiber, which can have a positive impact on the gut microbiome and are recommended for daily intake by adults. Incorporating almond skins into food products could enhance their nutritional value by

increasing their Fiber content, representing a sustainable approach to food production.

The relationship between inflammation, oxidative stress, and carcinogenesis highlights the necessity of investigating the anti-inflammatory effects of almonds and their by-products. Such analysis is important for enhancing the value of these food products. The elevated concentrations of bioactive compounds, predominantly polyphenols, unsaturated fatty acids, and protein hydrolysates, are considered the main factors contributing to this bioactivity.

Antimicrobial activity:

Polyphenols found in almond skin are bioavailable in the upper gastrointestinal tract, suggesting their potential for absorption during the digestive process. Additionally, the antimicrobial properties of almond skin indicate that this by-product may serve as a valuable source of natural antimicrobials. The antimicrobial characteristics of fractions obtained from almond skin were assessed against both Gram-positive and Gram-negative bacteria, as well as yeast. The fractions, whether natural or blanched, demonstrated antibacterial effects against two Gram-positive bacteria, namely *Listeria monocytogenes* and *Staphylococcus aureus*, while the natural almond skin exhibited activity against the Gram-negative bacterium *Salmonella enterica*.

Hepatoprotective activity:

Catechins constitute over 40% of the polyphenolic flavonoid content found in almond skin. These compounds are also present in various plants. They are generally regarded as non-toxic and have shown protective properties against oxidative stress. Recent studies indicate their capacity to form adducts with Di carbonyls. Additionally, catechins have been found to safeguard hepatocytes from cytotoxicity and lipid peroxidation induced by tert-butyl hydroperoxide (TBH). The bioactive constituents of almond skin in the non-lipophilic polyphenol extract have proven to be particularly effective in protecting hepatocytes from oxidative stress caused by hydroperoxide and in mitigating cytotoxicity induced by Di carbonyls.

Prebiotic activity:

In addition, studies have confirmed that almond skin exhibits prebiotic effects. The evaluation of the prebiotic potential of both natural and blanched almond skin was conducted using a complete gastrointestinal tract model,

which included in vitro gastric and duodenal digestion, followed by colonic fermentation with faecal bacterial cultures. Moreover, the prebiotic effects of almond and almond skin consumption were investigated in healthy subjects, with faecal samples analysed for microbiota composition and selected indicators of microbial activity.

Almond skins exhibit a beneficial probiotic index, similar to that of fructo-oligosaccharides, indicating a favourable relationship between the growth of beneficial bacteria and less desirable ones. The results from these studies consistently indicated that almond skin stimulates typical prebiotic effects, likely due to its high dietary fibre content that is resistant to digestive enzymes, enabling it to reach the large intestine where it interacts with microbiota to support gut health.

The dietary polyphenols present in almond skin may also contribute positively to gut health, as they can alter gut microecology and increase the abundance of beneficial microbiota. Phenolic compounds may interact with proteins in the food matrix, and although the bio accessibility of each antioxidant varies significantly, the potential synergistic effects among polyphenols are likely to enhance their bioactivity and influence glycoprotein transporters across the mucosal epithelium.

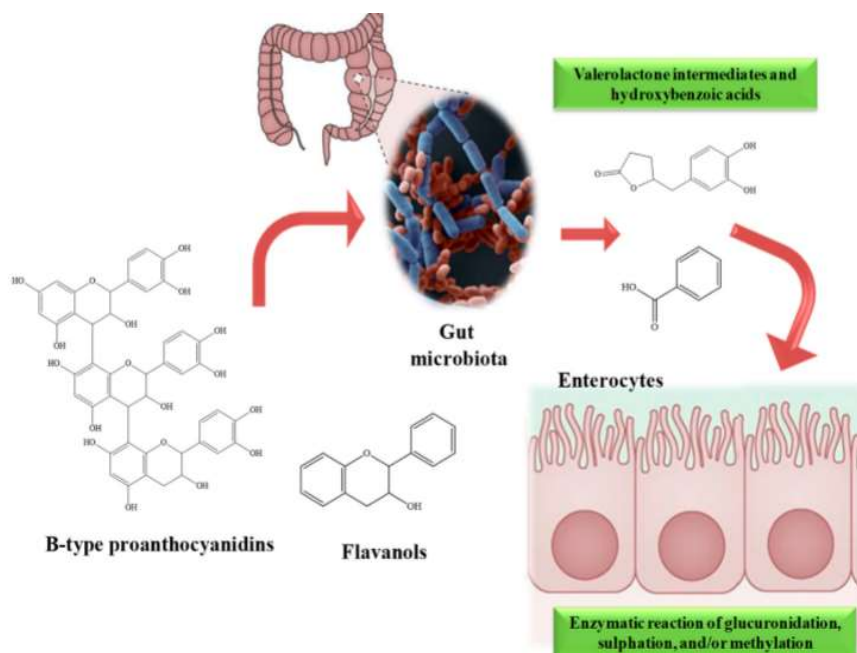


Fig 2., Prebiotic activity Source: (Barreca *et al.*, 2020)

Anticancer activity:

The anticancer properties of almonds and their by-products have been evaluated through both in vitro and in vivo studies, particularly focusing on the roles of oleic and linoleic acids. These compounds demonstrated a significant antiproliferative effect on two colon carcinoma cell lines, namely the primary line (Colo-320) and the metastatic line (Colo-741), in a manner that is dependent on both dosage and duration of exposure.

In recent years, extracts rich in polyphenols from almond skins have been recognized as promising components for the development of health products aimed at enhancing human well-being, primarily due to their significant antioxidant properties. Furthermore, employing antioxidants to combat oxidative stress is viewed as a novel approach to mitigate dysmetabolic diseases. The measurement of antioxidant capacity has become a standard method for identifying foods and beverages that are high in antioxidants. The antioxidant effects can result from the direct neutralization of various reactive oxygen species, including $\text{HO}\bullet$, $\text{ROO}\bullet$, $\text{NO}\bullet$, and $\text{LOO}\bullet$, with different antioxidants exhibiting varied responses to distinct radical or oxidant sources.

Various researchers have emphasized the beneficial effects of almond by-products on several chronic conditions, such as dyslipidaemia, diabetes, and cardiovascular diseases (CVD), in addition to their neuroprotective and hepatoprotective properties.

By product utilization of almond peel:

The concept of a circular economy encompasses the practices of reusing, repairing, refurbishing, and recycling existing materials and products, transforming what was once considered waste into valuable resources. Within the context of the food system, a circular economy aims to minimize waste generation, promote the reuse of food, and enhance the utilization of by-products and food waste, alongside nutrient recycling and the adoption of more diverse and efficient dietary patterns.

The nut industry generates a significant amount of byproducts, principally shells and skins. These wastes represent a significant store of bioactive substances that can be collected and used to improve a variety of products. In the confectionary sector, the processing of blanched almonds resulted in significant numbers of almond skins, which are generally used for animal feed and composting.

The industrial procedure for preparing almonds for consumption entails a systematic removal of the outer layers of the almond, resulting in the extraction of the almond meat or kernel. This is followed by various processes, including roasting, where the almond skin is loosened but not completely removed, and blanching, which produces peeled almonds by discarding the skin and creating a by-product referred to as blanch water.

Almond skin represents only around 4.0% of the total weight of the almond and is typically associated with a low economic value. Recent studies have indicated that this, however, remains a point of interest skin contains about 60.0% to 80.0% of the total phenolic compounds found in the nut. It is vital to acknowledge that the phenolic compounds in almond skin are significantly responsible for the high radical scavenging activity demonstrated by these materials, and they may also be linked to the positive health effects associated with the consumption of almonds.

These by-products are recognized as valuable and cost-effective sources of nutraceuticals and functional compounds, particularly bio-phenols. These can be repurposed in the creation of highly nutritious and health-promoting foods with an extended shelf life. Furthermore, re-utilizing materials discarded during industrial processing can help mitigate food waste, support environmental sustainability, and enhance the long-term viability of the food production system.

Concurrently, the rise in almond by-product production is associated with a corresponding increase, which adversely affects not only the local environment but also the economic stability of related industries. In this context, understanding the biochemical profile and functionality of these materials will facilitate their efficient utilization through innovative valorization strategies, thereby supporting the concept of a "zero waste economy."

Therefore, it can be concluded that the waste and by-products produced by the agri-food sector have recently become a focal point of scientific inquiry. There is a considerable transformation taking place which is occurring towards the recycling of these by-products to decrease waste and mitigate disposal costs.

Significance of value-added products & valorization of almond peel:

Food technologists are increasingly focused on enhancing the nutritional profile of products by integrating healthier components, including dietary fibres, vitamins, minerals, and essential oils. The consumption of

bioactive compounds, such as antioxidants, total phenols, flavonoids, and flavonol, plays a crucial role in promoting health and safeguarding against diseases like cancer, cardiovascular issues, and various other degenerative conditions.

These natural compounds can also be classified as nutraceutical ingredients or supplements, facilitating the creation of products that offer improved nutritional value, possible health advantages, extended shelf-life, and an appealing sensory profile. Byproducts consist of polysaccharides, organic acids, proteins, and various other compounds, which, without incurring extra production costs and at a lower industrial expense, provide a valuable source of natural compounds. These can potentially be utilized in the food industry as sources of food additives.

Almond by-products are currently a concern in waste management, as they can be repurposed into valuable products or ingredients. This transformation would facilitate waste reduction and mitigate their environmental effects. In addition to the significance of almonds and their by-products in food applications, these resources have been investigated for innovative strategies aimed at establishing a robust circular economy within this productive agri-food sector.

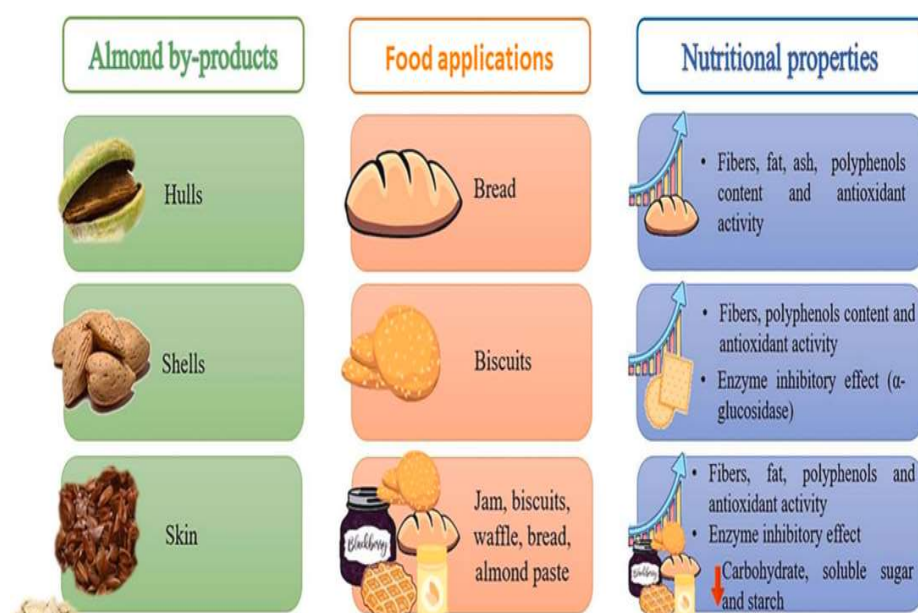


Fig 3.,Application of almond by products Source:(Lacivita *et al.*, 2024)

Experimental studies of almond peel valorization:

Gaglio *et al.*, (2023) developed food products enriched with almond skins involve the modification of a classic sourdough bread recipe, substituting semolina with 5% and 10% almond peel. Similar outcomes were observed in baked goods when almond skins were incorporated into waffle recipes. Additionally, a higher concentration of polyphenols, fibre, and enhanced antioxidant properties were recorded in functional blackberry jam that included 20% powdered almond skin, compared to formulations with 10% and 15% fortification. Biscuits that promote health have been formulated by integrating almond skins powder, which replaces 10 to 20 grams of wheat flour for every 100 grams used was developed by **(Pasqualone *et al.*, 2020)**.

A novel application for almond skins has been introduced by **Loizzo *et al.*, (2021)**, aiming to enhance the formulation of blackberry jam. This approach utilizes different concentrations of dry powder, specifically 10%, 15%, and 20% w/w.

Oliveira *et al.*, (2023) studied on Waffles enhanced with almond skin at four varying concentrations—1%, 2%, 5%, and 10%—demonstrated increased levels of phenolic compounds and antioxidant activities in comparison to the control sample. Over the past few decades, there has been a notable rise in almond consumption, attributed to its health-promoting properties and nutrient density. This surge in demand has resulted in an expansion of the almond market, accompanied by an increase in the production of associated by-products.

By adopting a circular economy framework for the management of almond by-products, it is possible to recover valuable natural ingredients that can be used in the creation of new nutritional, cosmetic, and pharmaceutical offerings. Such a strategy would not only address future consumer preferences concerning environmental impact but also prioritize human health. These are few research which includes the application of almond peel on new product development.

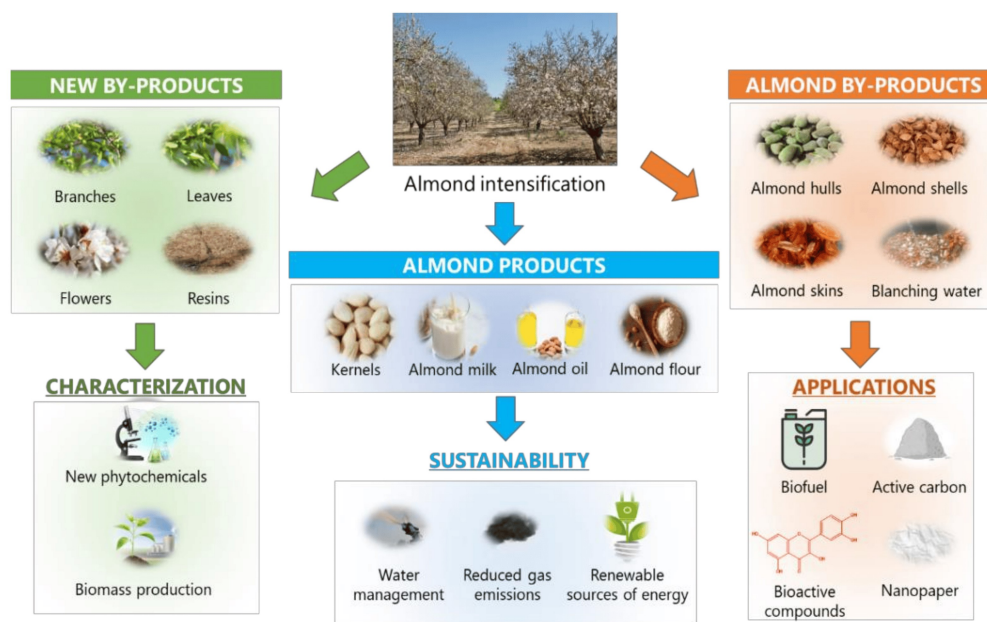


Fig 4., Valorization of Almond & its by-products Source:(Garcia-Perez *et al.*, 2021)

Conclusion

The flesh of the almond seed is surrounded by a brown, leathery covering known as the seedcoat, which serves to protect the almond from oxidation and microbial contamination. The skins of almonds, which are by-products of agricultural processing, are abundant sources of phenolic compounds. Almond seedcoat are readily accessible sources of these phenolics. These phenolic compounds play a crucial role in inhibiting lipid oxidation by scavenging free radicals, chelating metals, activating antioxidant enzymes, reducing tocopherol radicals, and inhibiting enzymes responsible for oxidation reactions. The significant polyphenol levels found in almonds and their by-products provide these natural matrices with strong antioxidant properties, which are associated with beneficial health effects. The implementation of a circular economy model for the use of almond by-productscould facilitate the extraction of natural ingredients, enabling the development of innovative Nutritional, cosmetic, and pharmaceutical products. This strategy signifies a progressivethinking strategy that aligns with future consumer demands

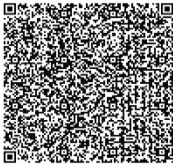
regarding environmental sustainability and health. Creating novel products from almond residue can reduce environmental effect and increase value in almond cultivation. So, it is concluded that almond peel contains an effective bioactive component and can be used in product development.

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"The Vicious Cycle of Multidrug-resistant (MDR) bacteria"

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Introduction

Multidrug-resistant (MDR) bacteria are well-recognized to be one of the most important current public health problems. The Infectious Diseases Society of America (IDSA) recognises antimicrobial resistance as “one of the greatest threats to human health worldwide” (Spellberg *et al.*, 2015). Several issues underlie the critical danger that is posed by the rise of MDR bacteria. First and most importantly, outcomes in patients infected with MDR bacteria tend to be worse as compared to patients infected with more susceptible organisms (Vardakas *et al.*, 2013). In this way, rising rates of antibacterial resistance have an impact on all aspects of modern medicine and threaten to decrease the yield of many accomplishments such as cancer care, transplantation and surgical procedures (Perez *et al.*, 2020). Second, tremendous added costs are associated with these infections. In the US, associated annual additional costs of infections caused by resistant organisms as compared to susceptible organisms are estimated between \$21 billion and \$34 billion (Spellberg *et al.*, 2015). Third, the prevalence of specific MDR bacteria is closely linked to the use of broad-spectrum antibiotics, both for empiric as well as for definitive therapy. This increased use in turn leads to even higher rates of MDR bacteria, thus creating a vicious cycle.

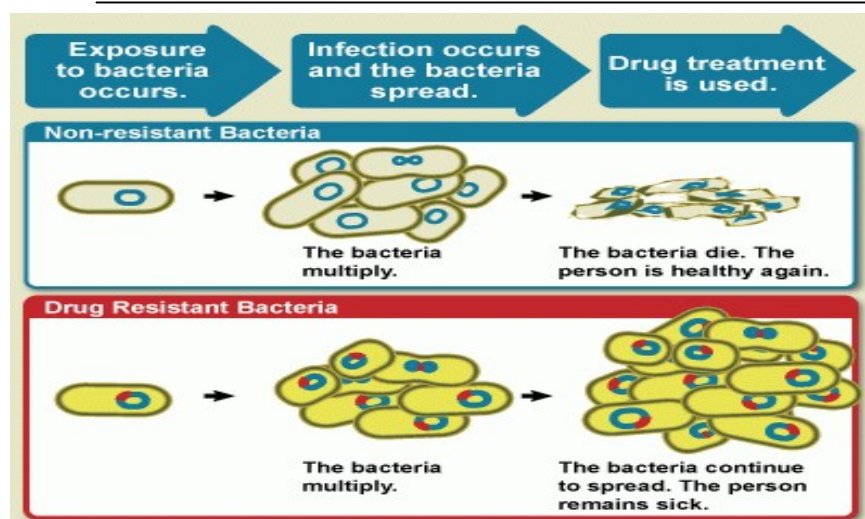


Figure 1: the difference between non-resistant bacteria and drug-resistant bacteria. Non-resistant bacteria multiply, and upon drug treatment, the bacteria die. Drug-resistant bacteria multiply as well, but upon drug treatment, the bacteria continue to spread (*NIAD*).

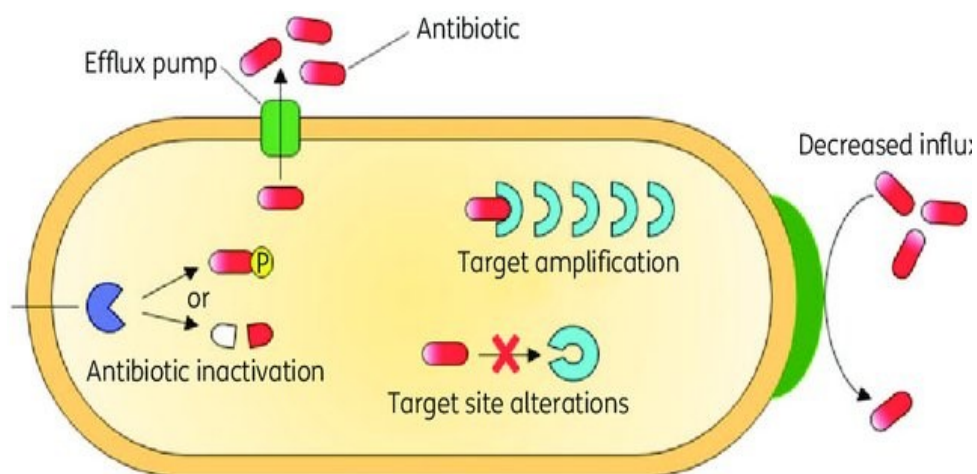


Figure 2. Schematic diagram highlighting the antibiotic resistance mechanisms utilized by bacteria. MDR pathogens can employ one or more of these mechanisms to become resistant to a diverse array of antibiotics. This figure appears in colour in the online version of JAC and in black and white in the print version of JAC. (Alav *et al.*, 2018)

Typically, MDR bacteria are associated with nosocomial infections. However, some MDR bacteria have become quite prevalent causes of community-acquired infections. This is an important development as community spread of MDR bacteria leads to a large increase of the population-at-risk, and subsequently an increase in the number of infections caused by MDR bacteria. In addition, when the incidence of a certain resistance pattern in bacteria causing community-acquired infections exceeds a specific threshold, broader spectrum antibacterials and/or combination antibacterial therapy are indicated for the empiric treatment of community-acquired infections.

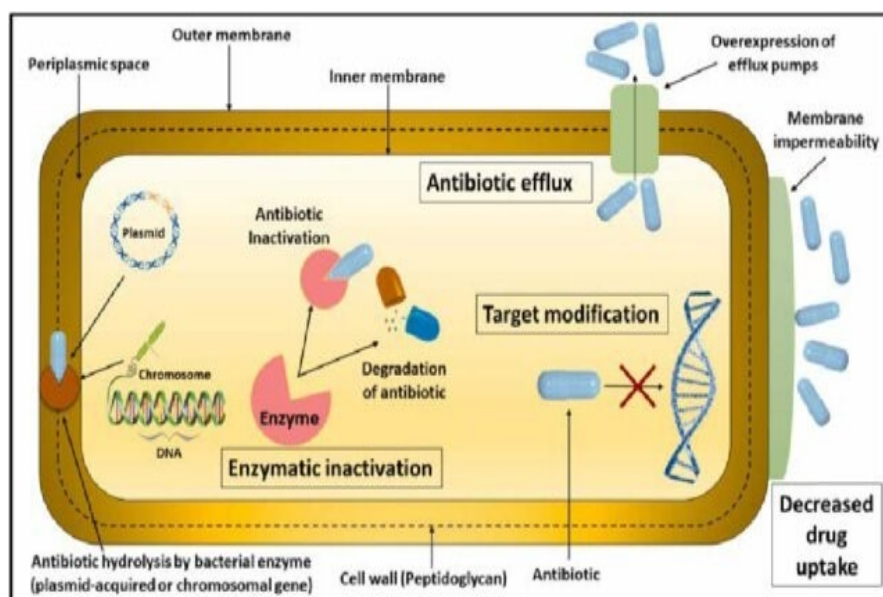


Figure 3. Diagrammatic presentation of multi-drug-resistance mechanism in bacteria (Ciamak Ghazaei, 2022)

Previous Definitions Applied to Bacteria Resistant to Multiple Antimicrobial Agents

MDR

In literal terms, MDR means 'resistant to more than one antimicrobial agent', but no standardized definitions for MDR have been agreed upon yet by the medical community. Many definitions are being used in order to characterize patterns of multidrug resistance in Gram-positive and Gram-negative organisms (Cohen *et al.*, 2008). The absence of specific definitions

for MDR in clinical study protocols gives rise to data that are difficult to compare.

One of the methods used by various authors and authorities to characterize organisms as MDR is based on IN VITRO antimicrobial susceptibility test results when they test 'resistant to multiple antimicrobial agents, classes or subclasses of antimicrobial agents' (Kallen *et al.*, 2010). The definition most frequently used for Gram-positive (Pillar *et al.*, 2008) and Gram-negative (Fallon *et al.*, 2009) bacteria is 'resistant to three or more antimicrobial classes'. An overview of the variability of these definitions is provided in a comprehensive review of MDR in *P. AERUGINOSA* and *A. BAUMANNII* where the authors note that a sizeable number of studies do not propose any specific definitions for MDR, but the majority define MDR as 'resistant to three or more antimicrobial classes'.

Another method used to characterize bacteria as MDR is when they are 'resistant to one key antimicrobial agent' (Hidron *et al.*, 2008). These bacterial isolates may have public health importance due to resistance to only one key antimicrobial agent, but they often demonstrate cross or co-resistance to multiple classes of antimicrobials, which makes them MDR. Creating an acronym for a bacterium based on its resistance to a key antimicrobial agent (e.g. methicillin resistance in *S. AUREUS*, i.e. MRSA) immediately highlights its epidemiological significance; the advantage of using this approach for surveillance purposes is that it can be easily applied.

XDR

Bacteria that are classified as XDR are epidemiologically significant due not only to their resistance to multiple antimicrobial agents, but also to their ominous likelihood of being resistant to all, or almost all, approved antimicrobial agents. In the medical literature XDR has been used as an acronym for several different terms such as 'extreme drug resistance', 'extensive drug resistance', 'extremely drug-resistant' and 'extensively drug-resistant' (Tseng *et al.*, 2009).

Initially, the term XDR was created to describe extensively drug-resistant *Mycobacterium tuberculosis* (XDR MTB) and was defined as 'resistance to the first-line agent's isoniazid and rifampicin, to a fluoroquinolone and at least one of the three-second-line parenteral drugs (i.e. amikacin, kanamycin or capreomycin)' (CDC 2006). After this, definitions for strains of non-mycobacterial bacteria that were XDR were constructed according to the principle underlying this definition for XDR MTB (i.e. describing a resistance profile that compromised most standard

antimicrobial regimens). Two sets of criteria have mainly been used to characterize bacteria as XDR. The first is based on the number of antimicrobials or classes or subclasses to which a bacterium is resistant, and the second is on whether they are ‘resistant to one or more key antimicrobial agents’ (Tavares *et al.*, 2019).

PDR

From the Greek prefix ‘pan’, meaning ‘all’, pan drug-resistant (PDR) means ‘resistant to all antimicrobial agents’. Definitions in the literature for PDR vary even though this term is etymologically exact and means that, for a particular species and a bacterial isolate of this species to be characterized as PDR, it must be tested and found to be resistant to all approved and useful agents. Examples of current definitions are: ‘resistant to almost all commercially available antimicrobials’, ‘resistant to all antimicrobials routinely tested’ and ‘resistant to all antibiotic classes available for empirical treatment’ (Kuo *et al.*, 2015), making the definition of PDR subject to inconsistent use and liable to potential misinterpretation of data.

Common antibiotic-resistant bacterial species

Globally, infections linked to health care increase rates of morbidity and mortality. Antimicrobial resistance, which restricts the use of antibiotics and makes it more challenging to treat infections brought on by multiresistant microbes, is directly linked to the rise in mortality. Infections with gram-negative bacteria that are resistant to carbapenem, primarily Enterobacteria, emerged as a significant public health concern at the start of the twenty-first century (Tacconelli *et al.*, 2014). Nosocomial infections are thought to be mostly caused by MDR gram-negative bacteria, such as *A. baumannii*, *Pseudomonas aeruginosa*, Enterobacteria that produce extended-spectrum beta-lactamase (ESBL), and Enterobacteria that are resistant to carbapenem (Teerawattanapong *et al.*, 2017). “The World Health Organization (WHO) has identified the genera *Pseudomonas*, *Acinetobacter* and *Enterobacter* as those belonging to the Gram-negative family of bacteria for which new and effective medications are desperately needed. They are used as “the last line of antibiotic defence” against resistant organisms because, among other things, they produce an extended spectrum of β -lactamases (ESBLs) that confer resistance to antimicrobials like cephalosporins, penicillins, and monobactams. Additionally, they include an increasing number of strains that are resistant to carbapenem” (Bush & Jacoby, 2010, Miyagi & Hirai, 2019). Concerningly, during the past 10 years, there has been a noticeable global rise in nosocomial CRB (carbapenem-resistant bacterium) infections; infections

caused by *Acinetobacter* and *Pseudomonas* have been linked to 40–80% mortality in intensive care units (Joshi & Litake, 2013 and Safaei *et al.*, 2017).

According to current reports, the most prevalent bacterial diseases are vancomycin-resistant *Enterococcus* (VRE) and methicillin-resistant *S. aureus* (MRSA). Hospitals have also been shown to harbour animal products, water, and animals. Although some strains of MDR *P. aeruginosa*, Carbapenem-resistant Enterobacteriaceae, and *A. baumannii* have also been recovered from foods, animals, and water, clinical samples have been the primary source of these germs. Many of these organisms are opportunistic infections that contaminate the ill or immunocompromised. Many of these bacteria seem to be found in large quantities in nature, and a contaminated environment can promote their proliferation. Public health is also greatly concerned about the Gram-positive methicillin-resistant *Staphylococcus aureus* (MRSA) and vancomycin-resistant *Enterococcus* (VRE) bacteria. It is well-recognised that both can spread epidemics (Arora *et al.*, 2014).

The World Health Organization released the first-ever list of antibiotic-resistant "priority pathogens," which is a catalogue of 12 bacterial families that are the biggest threats to human health. According to how urgently new antibiotics are needed, the WHO list is split into three categories: critical, high, and medium priority (WHO, 2017).

Priority 1: CRITICAL

- ❖ *Acinetobacter baumannii*, carbapenem-resistant
- ❖ *Pseudomonas aeruginosa*, carbapenem-resistant
- ❖ *Enterobacteriaceae*, carbapenem-resistant, ESBL-producing

Priority 2: HIGH

- ❖ *Enterococcus faecium*, vancomycin-resistant
- ❖ *Staphylococcus aureus*, methicillin-resistant, vancomycin-intermediate and resistant •
- ❖ *Helicobacter pylori*, clarithromycin-resistant
- ❖ *Campylobacter spp.*, fluoroquinolone-resistant
- ❖ *Salmonellae*, fluoroquinolone-resistant
- ❖ *Neisseria gonorrhoeae*, cephalosporin-resistant, fluoroquinolone-resistant

Priority 3: MEDIUM

- ❖ *Streptococcus pneumoniae*, penicillin-non-susceptible

- ❖ *Haemophilus influenzae*, ampicillin-resistant
- ❖ *Shigella spp.*, fluoroquinolone-resistant

There has been no standard approach for determining the types, classes or groups of antimicrobial agents that should be used when defining MDR, XDR and PDR. Frequently, chemical structures for antimicrobial classes (e.g. cephalosporins) (Prescott, 2013), antimicrobial subclasses, (e.g. third-generation cephalosporins) (Dubois *et al.*, 2001) or specific antimicrobial agents (e.g. ceftazidime) (Pagani *et al.*, 2005) have been used to define these terms. This approach is not always conclusive and makes it difficult to compare results between studies. The expert group, therefore, constructed ‘antimicrobial categories’ for each of the organisms or organism groups with the intent of placing antimicrobial agents into more therapeutically relevant groups.



Figure 4: Multidrug resistance Bacteria-A threat

Prevention

Prevention of further spread of MDR bacteria in the community is one of the most urgent public health challenges. Unfortunately, national or even regional data on antibiotic susceptibilities are often limited. In addition, when these data are available in some form, the accompanying epidemiologic metadata is usually too restricted to determine which isolates are truly community-associated. Furthermore, clinical infections are generally the tip

of the proverbial iceberg. Once a signal is generated that is sufficient in amplitude to get the attention of policy-makers, subclinical spread has already occurred.

Any successful prevention strategy must consist of a multi-pronged approach and involve all stakeholders. In addition to human clinical antimicrobial stewardship, we need to remove antibiotics from the food chain. Furthermore, we need to limit the number of xenobiotics such as quaternary ammonium compounds that reach the environment (Hawkey and Jones, 2009). Another challenging step in limiting exposure of bacteria to antibiotics is the treatment of contaminated wastewater such as that generated by pharmaceutical factories and medical facilities. For instance, a study evaluated samples collected from a wastewater treatment plant in India that received water from 90 regional bulk drug manufacturers containing – amongst other compounds – higher concentrations of ciprofloxacin than are generally found in the blood of patients who are being treated with this agent. Bacteria recovered from this water were tested against 39 antibiotics. Approximately 30% of bacteria were resistant to 29–32 antibiotics tested, and another ~20% were resistant to 33–36 antibiotics (Marathe *et al.*, 2013). The magnitude of this effect, combined with the knowledge that soil-dwelling bacteria will pass on resistance genes to more clinically relevant bacteria, illustrates the importance of limiting this contamination (Forsberg *et al.*, 2019).

Antimicrobial stewardship is developing rapidly as a hospital speciality. Stewardship teams often will combine strengths from Infectious Disease medical specialists and doctors of pharmacy to evaluate the appropriateness of choice and duration of antibiotic strategies (Wagner *et al.*, 2014). However, most antibiotics are prescribed in ambulatory care and more attention is needed in this realm to impact overall community antibiotic exposure (van Duin & Paterson, 2020). This will not only require a paradigm shift in the behavior of prescribers but also a cultural shift in the public on the risks and benefits of antibiotics. Rapid diagnostic testing to identify MDR bacteria more quickly and thus limit the empiric use of unnecessarily broad antibiotics will be of great significance. Also, rapid testing to diagnose alternative, non-bacterial etiologies is important.

An important question is whether any interventions can address the issue of chronic colonization with MDR bacteria. Decolonizing these patients would decrease the risk of transmission. Also, the burden on the individual patient with this condition should not be underestimated. In many healthcare systems, patients with MRSA or CRE are “labelled” as carriers for life, resulting in the institution of isolation precautions whenever they are admitted to the hospital. This has multiple adverse effects and leads to decreased patient satisfaction (Vinski *et al.*, 2012). For these reasons, decolonization is a theoretically attractive option. However, most decolonizing strategies involve the use of antibiotics. For MRSA decolonization, most strategies involve some combination of intranasal mupirocin with topical chlorhexidine. This approach is effective in decreasing infections after surgery (Chen *et al.*, 2022). However, the effect is generally short-lived and recurrence of colonization is the rule. For enteric bacteria, no good options are currently available. Various selective gut decontamination strategies have been described, but none have shown true promise. In addition, with growing knowledge of the role of the gut microbiome in the defence against MDR bacteria, it would seem counter-intuitive to give even more antibiotics. Modulating the gut microbiome either through probiotics or faecal microbiota transplantation is a promising but experimental method of decolonising patients.

Conclusion

In Conclusion, antibiotic resistance poses a formidable threat to global public health, requiring urgent and concerted efforts from the scientific, medical, and policy-making communities. Antibiotic resistance could usher in a post-antibiotic era, where common infections and minor injuries become life-threatening. Addressing antibiotic resistance demands a multifaceted approach. First and foremost, there is an urgent need for global cooperation to curb the inappropriate use of antibiotics in human medicine, agriculture, and animal husbandry. Public awareness campaigns can play a crucial role in educating the public and healthcare professionals about responsible antibiotic use. Additionally, fostering the development of new antibiotics and alternative treatment strategies is essential to stay ahead of evolving bacterial resistance.

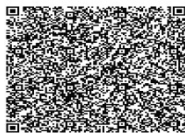
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Sourdough: A Journey Through Tradition, Innovation, and Nutrition

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Abstract

Sourdough bread is an exceptional culinary creation that blends ancient traditions with modern nutritional understanding. Celebrated for its rich flavour, distinctive texture, and artisanal quality, sourdough is made through the natural fermentation of flour and water by *wild yeasts* and *lactic acid bacteria*. This fermentation process not only contributes a delicious taste and aroma but also improves its health benefits, such as enhanced digestibility, a lower glycemic index, and better gut health. Valued for its ability to reduce anti-nutrients and boost nutrient bioavailability, sourdough presents a novel solution to contemporary dietary challenges. As its popularity increases in both global and Indian markets, sourdough rises beyond mere sustenance, becoming a representation of wellness, artistry, and heritage. Ranging from traditional whole-grain types to gluten-free options, sourdough links the past with innovation, embodying the lasting importance of nutrition and craftsmanship.

Key words: sourdough bread natural fermentation, nutritional value, and health benefits.

Introduction

Sourdough bread, celebrated for its unique flavor, texture, and numerous health benefits, occupies a prestigious role in culinary practices across the globe. Its distinct production method relies on natural fermentation with *wild yeast* and *lactic acid bacteria*, which together leaven the dough and provide a tangy taste alongside a chewy consistency. This fermentation approach sets sourdough apart from bread created with commercial yeast. Dating back over 6,000 years to around 3700 BC, sourdough is recognized as one of the earliest variations of leavened bread, likely discovered inadvertently when *wild yeast* and *bacteria* fermented a blend of flour and water, leading to the creation of sourdough starters essential in ancient cooking.

The story of sourdough began in the Fertile Crescent and Ancient Egypt, where it became a crucial food source. Egyptian bakers were famed for producing substantial amounts of sourdough bread to feed workers involved in monumental construction endeavors like the pyramids. The Roman Empire played a significant role in spreading sourdough culture, promoting wheat cultivation and advanced baking techniques across its territories. This transition transformed bread from merely a staple into a powerful emblem of nourishment, community, and cultural identity. Throughout the Middle Ages, sourdough retained its significance, especially during times of famine, commonly baked in communal ovens, thus reinforcing its place in community and spiritual life, as highlighted by biblical references like Bethlehem, which means "House of Bread."

The Industrial Revolution brought about a pivotal shift in bread-making practices. The advent of mechanized processes and refined yeast began to eclipse traditional sourdough techniques, as mass production prioritized speed and efficiency instead of traditional artistry. Nonetheless, the remarkable attributes and rich heritage of sourdough contributed to its endurance. In recent years, there has been a resurgence of interest in sourdough, driven by the artisanal baking movement and increased acknowledgment of its health benefits. Unlike many commercially produced breads, sourdough is praised for its enhanced digestibility and lower glycemic index, making it a favored option for health-conscious buyers. Today, sourdough represents a lasting connection between ancient culinary customs and modern food culture, signifying not only a kind of bread but also an art form that celebrates life, renewal, and community.

The process of making sourdough encourages bakers of all skill levels to engage with its intricacies, investigate various techniques, and experience the satisfaction of producing a loaf rich in thousands of years of history, linking the past with the present through its flavorful, nutritious, and meaningful essence.

Sourdough In Indian Market

The emergence of sourdough bread in India is a relatively recent development, influenced by global culinary trends and shifting food preferences. Traditionally, Indian cuisine has centred on leavened breads like naan and roti, with sourdough having little historical presence. However, growing interest in artisanal baking and natural fermentation has spurred curiosity among Indian consumers. Exposure to international food culture and the demand for healthier alternatives have spotlighted sourdough's distinctive taste, improved digestibility, and lower glycaemic index compared to conventional bread.

Responding to this shift, urban artisanal bakeries have started incorporating sourdough into their offerings. Many bakers blend traditional methods with local ingredients, experimenting with flours like millet and Indian spices to craft unique variations that appeal to regional tastes. Simultaneously, the popularity of sourdough-focused baking workshops and classes has nurtured a community of enthusiasts eager to master this age-old craft at home. This growing trend reflects a harmonious fusion of culinary innovation and traditional fermentation, positioning sourdough as a favoured choice for health-conscious and discerning Indian consumers.

Reasons For The Limited Sourdough Market In India

Sourdough (SD) bread faces several challenges in India due to its high production costs, labour-intensive preparation processes, and the need for prolonged fermentation and regular maintenance of starters. The bread's heavy texture and unique tangy flavour often fail to appeal to local consumers, which limits its acceptance. Additionally, the complex microbiological interactions can lead to inconsistent quality, and the lengthy, meticulous fermentation process presents difficulties for large-scale commercial production. Collectively, these factors, along with lower affordability and demand, make SD bread less feasible and marketable compared to traditional yeast bread in the Indian market.

Sourdough As A Leavening Agent In Bread Production

Natural sourdoughs rely on wild fermentation facilitated by a diverse array of microorganisms that generate various organic acids, alcohols, esters, and other compounds. However, the unpredictable nature of this fermentation can create inconsistency in the production of carbon dioxide, acids, and alcohol, which impacts the rising of the dough and the overall quality of the bread. Factors such as the types and concentrations of microorganisms, as well as fermentation time, temperature, and pH, influence this process. To achieve consistency, strict sterilization practices, including sterilizing equipment and filtering air are implemented, and fermented mixtures are converted into products like compressed or active dry yeast through techniques such as centrifugal separation and drying.

Sourdough, recognized for its unique sour flavour, has been a fundamental aspect of bread-making since its inception in ancient Egypt around 1500 BC, acting as the first leavening agent for bakers. Its significance was particularly notable in the production of rye bread, as traditional baker's yeast is not suitable for rye due to problems such as ropiness. The sensory and functional attributes of sourdough have ensured its continued relevance, with uses extending from traditional millet bread in Africa and the Middle East to contemporary dried sourdough products that offer convenience in baking.

The fermentation process in sourdough occurs through alcoholic, lactic, and hetero-fermentative pathways, with indigenous yeasts converting glucose into ethanol and carbon dioxide. Despite variations in raw materials and surrounding conditions, the stability of sourdough ecosystems is maintained through metabolic adjustments and the production of antimicrobial compounds by lactobacilli and yeasts. These microbial interactions contribute to a consistent quality, solidifying sourdough's role as a reliable and enduring component in bread-making.

Classification of Sourdough

Sourdough can be classified into four types based on its production method and dough consistency. The consistency is measured by the dough yield, which represents the ratio of flour to water and is calculated using the formula: $\text{Dough Yield (\%)} = (\text{flour amount} + \text{water amount}) \times 100 / \text{flour amount}$. During the microbial adaptation phase, various environmental conditions and technological factors play a crucial role in determining the selection and performance of the microorganisms.

Table 1 Classification of Sourdough

Process	Traditional Method	Industrial Method	Advanced Industrial	Lab-Bakery Method
Raw Material	Water + Flour	Water + Flour	Starter Culture Isolation	Mature SD + Starter Culture
Fermentation	20-30°C	Fast (30°C)	Type II SD → Dehydration	Mixed SD Type IV
Storage	Requires Refreshing	Stored for 2-5 Days	Long Shelf Life	Periodic Propagation
Examples	Pain au Levain, Panettone	White Pan Bread, Pizzas	Breads, Waffles	Similar to Type I Products

Classification of sourdough (francieli b. Siepmann et., Al. 2017)

Classification of starter cultures according to their production technology

The distinctive taste and texture of sourdough bread are mostly attributed to sourdough starter cultures, which are composed of wild yeasts and bacteria. Bakers can choose the ideal starter for reliable, superior results by grouping these cultures according to their microbial make-up and production techniques.

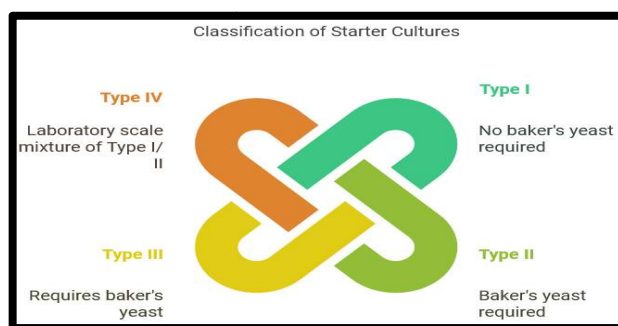


Fig. 1 classification of starter culture

Characteristics of sourdough starter cultures

Table 2 characteristics of sourdough starter culture

Natural Flora Starter Cultures (Type I)	<p>Produced using environmental <i>yeasts</i> and <i>bacteria</i> under ideal settings. Utilized in artisanal bakers most of the time. Need back slopping, or constant refreshing. Kept at the room temperature (20–30°C).</p> <ul style="list-style-type: none">❖ Starts with near-neutral pH❖ Gradual decline in pH due to organic acids (lactic acetic acid)❖ Matured Type I starters are highly acidic with dis smell, flavour, and bubbles. <p>Use Example (Arabic Bread): Traditional preparation involves whole wheat flour and water mixed daily for 6 days until sour smell and bubbles appear.</p>
Synthetic Starter Cultures (Type II)	<p>Natural starters usually contain compounded acids. Higher temperatures during fermentation accelerated the synthesis of bacterial acid. For uniformity and particular flavour qualities, it is utilized on a big scale.</p> <ul style="list-style-type: none">❖ Reduced pH system❖ Altered crumb texture or loaf volume❖ The addition of particular bacteria for a more sour flavor❖ The addition of baker's yeast or acid-tolerant yeast to guarantee rise.
Mixed Microbiologically Selected Starter Cultures (Type III)	<p>A blend of certain <i>lactobacilli</i> that have been dehydrated to preserve their activity. Usually used to stop <i>Bacillus subtilis</i> from making ropiness problems. Dry-resistant and able to endure in sourdough conditions.</p> <ul style="list-style-type: none">❖ Yeast and lactic acid bacteria strains resistant to drying are found in the dried form of Type II.❖ It encourages competition between the inoculated and naturally occurring microorganisms in the sourdough ecosystem.

Microbial Composition of Sourdough	Comprises a range of microorganisms, including <i>molds</i> , <i>yeasts</i> , and <i>bacteria</i> ; <i>lactic acid bacteria</i> and wild yeast strains are particularly significant.
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Process of Starter Culture

The procedure starts with ingredient selection, which guarantees the use of premium wheat and water for the best possible fermentation. First, a jar is filled with 100 millilitres of water and 100 grams of flour that have been carefully mixed. To produce a controlled fermentation environment, the jar is then covered. The first fermentation is given a full day to continue. Next, 100 millilitres of water and another 100 grams of flour are added to the mixture, and it is thoroughly combined once again. To guarantee optimal development, the fermentation process is observed every day for seven days while the jar is covered again. Following this time frame, the combination is prepared for usage.

Sourdough Fermentation

Bread is an essential food enjoyed around the world, primarily composed of cereal flour, water, salt, and leavening agents. The process of fermentation, particularly through sourdough (SD), significantly improves both the taste and nutritional value of bread. There are four principal types of sourdough fermentation methods. Type I fermentation follows a time-honored technique known as back slopping, which utilizes natural microbial cultures. While this method enhances flavor, it can be labor-intensive and is not easily scalable for mass production. Type II, on the other hand, employs specific strains of *Lactic Acid Bacteria*

(LAB) to boost acidity and flavor, making it more suitable for large-scale baking. Type III simplifies storage and transportation by using dehydrated sourdough, while Type IV merges starter cultures with traditional back slopping techniques.

LAB stands as the primary microbial agent in sourdough, contributing to flavor and nutritional enhancements by producing organic acids, peptides, and aromatic compounds. *Yeasts*, most notably *Saccharomyces cerevisiae*, serve as the main leavening agents, generating carbon dioxide that helps the bread rise while also enhancing flavor and improving gluten structure. The collaboration between *LAB* and *yeasts* results in bread that is not only more

digestible and has a longer shelf life but also offers a more appealing taste and texture.

Role Of Lactic Acid Bacteria (Lab)

Lactic Acid Bacteria (LAB) are key players in the fermentation process of sourdough, significantly enhancing the flavor, texture, and nutritional profile of bread. They create both lactic acid, which offers a fresh acidity, and acetic acid, which adds a sharper tang, giving sourdough its characteristic flavor. In addition to these acids, LAB generate amino acids, peptides, and various flavor compounds that enrich the aroma and umami taste of the bread.

LAB also play an important role in making minerals like calcium, iron, and zinc more accessible by breaking down anti-nutritional substances such as phytic acid. This process improves the overall nutritional value of the bread. Furthermore, LAB produce antimicrobial agents that can help prolong the shelf life of bread and mitigate gluten toxicity, making it easier to digest for those who are sensitive to gluten.

Role of Yeast

Yeasts, particularly *Saccharomyces cerevisiae*, are crucial for leavening bread by fermenting the dough and producing carbon dioxide (CO₂), which makes the dough rise. They also improve flavor by creating organic acids, alcohols, and esters during fermentation. Moreover, yeasts contribute to the strengthening of the gluten structure, enhancing gas retention and the overall texture of the dough.

In addition, yeasts boost the bioavailability of key minerals and release phenolic compounds that possess antioxidant properties, while also increasing the levels of vital vitamins such as B2 (riboflavin) and vitamin D. Their proteolytic function further improves protein digestibility and diminishes allergenic components like gluten, enhancing the nutritional value of the bread.

The collaboration between *lactic acid bacteria* (LAB) and yeasts results in bread with superior taste, texture, digestibility, and an extended shelf life, making it a well-balanced staple food.

Microorganism of Lab And Yeast

Microorganisms play a vital role in food fermentation and include various species of *lactic acid bacteria* (LAB) and yeast. The **obligately heterofermentative lab** are (*Lactobacillus acidifarinae*, *Lb. brevis*, *Lb. buchneri*, *Lb. cellobiosus*, *Lb. crustorum*, *Lb. curvatus*, and *Lb. fermentum*). The **facultatively heterofermentative lab** comprise (*Lb. alimentarius*, *Lb.*

buchneri, *Lactococcus lactis*, *Lb. paracasei*, *Lb. kimchi*, *Lb. paralimentarius*, and *Lb. pentosus*). The **obligately homofermentative lab** include (*Lb. acidophilus*, *Lb. amylolyticus*, *Lb. amylophilus*, *Lb. amylovorus*, *Lb. bulgaricus*, *Lb. farciminis*, and *Lb. johnsonii*). Lastly, the **YEAST** species are (*Saccharomyces cerevisiae*, *S. bayanus*, *Kazachstania exigua*, *K. humilis*, *K. servazzi*, and *Pichia kudriavzevii*). These microorganisms are essential in the production of various fermented foods, as they help enhance texture, develop unique flavors, and improve preservation. Their diverse functionalities make them integral to a wide range of food products enjoyed globally.

Different Types of Sourdough Breads

Various kinds of sourdough bread present a diverse array of flavors, textures, and health advantages, ranging from classic whole grain and rye sourdough to contemporary gluten-free alternatives. These different styles, which include flavored loaves and multifunctional rolls, showcase the flexibility of sourdough in both nutritious and gourmet baking.

Table 3 Different Types of Sourdough Breads

Whole Grain Sourdough Bread	Crafted with whole wheat, spelt, or a blend of multigrain flours, this sourdough bread prioritizes health advantages by providing increased fiber levels and vital nutrients. It's perfect for those in search of a more wholesome bread choice.
Rye Sourdough Bread	This classic European variant, rye sourdough, is recognized for its unique flavor and dense consistency. Its fermentation method improves digestibility and imparts a tangy, rich taste.
Gluten-Free Sourdough Bread:	This option utilizes alternative flours such as rice, millet, or buckwheat, serving those who are gluten-sensitive or have celiac disease. It maintains the signature tangy taste of sourdough while being suitable for restricted dietary needs.
Flavored or Specialty Sourdough Bread	Enriched with ingredients like rosemary, garlic, onion, olives, or cheese, these breads provide a gourmet flair. They are excellent for accompanying soups, stews, or enjoyed on their own as snacks.

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Sourdough Rolls	Ideal for sandwiches or as a complement, sourdough rolls are smaller, softer, and more adaptable versions of the traditional loaf
Sourdough Pancakes and Pizza Crusts	In addition to classic loaves, sourdough can also be made into pancakes, waffles, and pizza crusts. These dishes capitalize on the starter's fermentation for both flavor and leavening.

Difference Between Normal Baker's Yeast Bread & Sourdough Bread

Sourdough and conventional yeast bread are two widely enjoyed varieties of bread, each defined by its fermentation method and distinct features. Conventional yeast bread utilizes commercial yeast (*Saccharomyces cerevisiae*) for quick fermentation, resulting in a subtle flavor and tender texture. On the other hand, sourdough bread is leavened through natural fermentation by wild yeasts and lactic acid bacteria, which enhance its rich flavor and chewy consistency. The extended fermentation process in sourdough also provides advantages such as better digestibility, a reduced glycemic index, and an extended shelf life. This comparison examines the main differences in their microbial populations, nutritional advantages, and overall health impacts, highlighting the individual qualities of both types of bread.

Table 4 Difference Between Normal Baker's Yeast Bread And Sourdough Bread

ASPECT	NORMAL BAKER'S YEAST BREAD	SOURDOUGH BREAD
LEAVENING AGENT	Uses commercial yeast (<i>Saccharomyces cerevisiae</i>) for leavening.	Leavened through natural fermentation with wild yeasts and lactic acid bacteria.
FERMENTATION PROCESS	Quick fermentation, usually taking just a few hours.	Slow fermentation, which can range from several hours to days.

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MICROBIAL DIVERSITY	Limited, mostly dominated by a few commercial yeast strains.	Rich microbial community, including various <i>wild yeasts</i> and lactic acid bacteria (<i>LAB</i>), contributing to complex flavours.
FLAVOUR PROFILE	Mild flavour, without sourness.	Distinct and complex flavour with a characteristic tanginess from organic acids.
NUTRITIONAL BENEFITS	Offers fewer improvements in nutrient absorption compared to sourdough.	Better digestibility and nutrient absorption due to natural fermentation processes.
GLYCAEMIC INDEX (GI)	Typically has a higher GI, causing rapid blood sugar spikes.	Lower GI, promoting more stable blood sugar levels.
SHELF LIFE	Shorter shelf life, as it lacks acidity to preserve it.	Longer shelf life due to its acidity and organic acids, which help prevent mold growth
TEXTURE	Soft and uniform texture, without much chewiness.	Chewier texture with a more open crumb, due to gas retention during fermentation.
HEALTH EFFECTS	Minimal probiotics, which might not be gut-friendly for some.	May offer gut health benefits due to the presence of probiotics and post biotic compounds.

Health Benefits of Sourdough Manage The Obesity And Metabolic Diseases

Sourdough bread may aid in weight management and potentially help avert metabolic diseases due to its distinct fermentation process:

Lowered Glycemic Index: The fermentation of sourdough results in a reduced glycemic index (GI) compared to other bread varieties. The creation of organic acids, like acetic and lactic acid, slows down carbohydrate digestion, leading to a more gradual release of glucose.

Hormonal Appetite Control: The organic acids and certain fibers found in sourdough bread might enhance feelings of fullness by influencing hormones such as ghrelin and peptide YY (PYY), which play a role in hunger regulation.

Enhanced Lipid Profiles: Research suggests that consuming sourdough can lower LDL cholesterol and triglyceride levels, helping to reduce the risk of metabolic syndrome.

Decreased Inflammation Markers:

Chronic inflammation is a key characteristic of metabolic diseases. Lactic acid bacteria (LAB) in sourdough generate bioactive peptides that help diminish inflammation.

1. Improve the gut health

Gut health

Probiotics in sourdough:

❖ The main probiotic species found in sourdough are *Lactobacillus plantarum*, *Lactobacillus brevis*, and *Lactobacillus reuteri*.

❖ LAB generates lactic acid, which reduces pH levels and creates a hostile environment for harmful pathogens.

❖ The fermentation process boosts the production of bioactive peptides that have anti-inflammatory effects.

Prebiotics in Sourdough

Sourdough bread is also rich in prebiotic substances such as resistant starch and fermentable fibers, which serve as nourishment for gut probiotics and enhance their activity.

Gut Micro-biota Balance

Sourdough encourages the proliferation of beneficial gut bacteria, including *Bifido-bacteria* and *Lactobacillus species*.

Digestive Support

Enzymes generated during fermentation assist in breaking down starches and proteins, easing the digestive load on the gastrointestinal tract.

Strengthened Gut Barrier

Short-chain fatty acids (such as acetate and propionate) produced by probiotics help to maintain the integrity of the gut barrier, preventing harmful substances from entering the bloodstream.

Gastrointestinal

Sourdough bread, produced through fermentation, provides several health advantages, particularly for individuals suffering from digestive issues such as Celiac Disease (CD), Irritable Bowel Syndrome (IBS), Small Intestinal Bacterial Overgrowth (SIBO), and Inflammatory Bowel Disease (IBD) and Lactose intolerance. The fermentation process enhances digestion, decreases anti-nutrients, and boosts nutrient bioavailability, making sourdough a favorable option for many.

Celiac Disease (CD)

- ❖ **Decreased Gluten:** The fermentation of sourdough may lower gluten levels, making it more digestible for those with mild gluten sensitivities (though it's not gluten-free).
- ❖ **Relief for Digestion:** Fermentation assists in breaking down gluten proteins, which may help mitigate gut inflammation.
- ❖ **Enhanced Nutrient Absorption:** It minimizes anti-nutrients, facilitating better absorption of minerals such as calcium and zinc.

Irritable Bowel Syndrome (IBS)

- ❖ **Reduced FODMAPs:** The fermentation process lessens fermentable sugars, relieving bloating and discomfort for those with IBS.
- ❖ **Simplified Digestion:** The pre-digestion of starches and proteins within sourdough could lessen cramping.
- ❖ **Gut Health:** The prebiotic fiber found in sourdough fosters beneficial gut bacteria, aiding digestive function.

Small Intestinal Bacterial Overgrowth (SIBO)

- ❖ **Lower FODMAP Content:** Fermentation decreases FODMAPs, making sourdough gentler on the digestive system for individuals with SIBO.
- ❖ **Restored Gut Flora:** Probiotics present in sourdough assist in reinstating a healthy balance of gut bacteria.
- ❖ **Digestive Comfort:** The fermentation involved in sourdough contributes to the breakdown of carbohydrates and proteins, lessening digestive discomfort.

Inflammatory Bowel Disease (IBD)

- ❖ **Anti-Inflammatory Properties:** Sourdough consists of bioactive compounds that may help alleviate gut inflammation.
- ❖ **Support for Gut Barrier:** Probiotics help preserve the integrity of the gut barrier, which may prevent inflammation.
- ❖ **Enhanced Nutrient Absorption:** The fermentation process reduces anti-nutrients, thereby promoting better nutrient absorption for individuals with IBD.

Lactose Intolerance and Sourdough Decreased Lactose

- ❖ **The process of sourdough fermentation** diminishes lactose, making it more manageable for individuals with mild lactose intolerance.
- ❖ **Enhanced Digestion:** Extended fermentation further decreases lactose levels, improving digestibility.
- ❖ **Probiotic Benefits:** Probiotics present in sourdough, such as *Lactobacillus*, can assist in breaking down any leftover lactose, alleviating symptoms.

2. Regulate The Glycaemic Response And Blood Glucose Levels

Sourdough bread positively influences glycemic control because of its fermentation-related properties.

Starch Alteration: The fermentation process changes the structure of starch, making it less susceptible to digestion by enzymes, which decreases the glycemic response.

- ❖ **Diminished Postprandial Glucose Peaks:** Research indicates that eating sourdough bread leads to lower blood glucose levels after meals compared to consuming regular bread.

❖ **Enhanced Insulin Sensitivity:** Regular consumption of sourdough bread over time may improve insulin sensitivity, lowering the risk of developing type 2 diabetes.

3. High In Dietary Fiber And Reduce The Anti-Nutrients

Sourdough fermentation alters the fiber content and diminishes anti-nutritional components, enhancing the overall nutritional value:

Enhanced Resistant Starch: The fermentation process boosts resistant starch, which is not digestible and functions similarly to dietary fiber.

Decrease in Phytic Acid: *Lactic acid bacteria* (LAB) produce phytase, an enzyme that degrades phytic acid, an anti-nutrient that binds to minerals like iron and zinc, hindering their absorption.

Liberation of Bound Nutrients: Fermentation frees vitamins and minerals that are attached to complex structures in the flour, improving their bioavailability.

4. Enhance The Digestion, Bioavailability, And Bio-Accessibility

Sourdough bread offers enhanced digestibility and improves nutrient absorption thanks to its fermentation process

Protein Degradation: Proteolytic enzymes produced by lactic acid bacteria break down gluten proteins, making sourdough bread easier to digest for those with non-celiac gluten sensitivity.

Mineral Absorption: The decrease in phytic acid and the release of essential micronutrients such as calcium, magnesium, and iron enhance their absorption.

Accessibility of Polyphenols: The fermentation involved in sourdough increases the bioavailability of polyphenols, which are antioxidants associated with various health advantages.

Decrease in Digestive Discomfort: The gradual fermentation process leads to a more equilibrated pH and lowers the presence of FODMAPs (fermentable oligosaccharides, disaccharides, monosaccharides, and polyols), simplifying digestion.

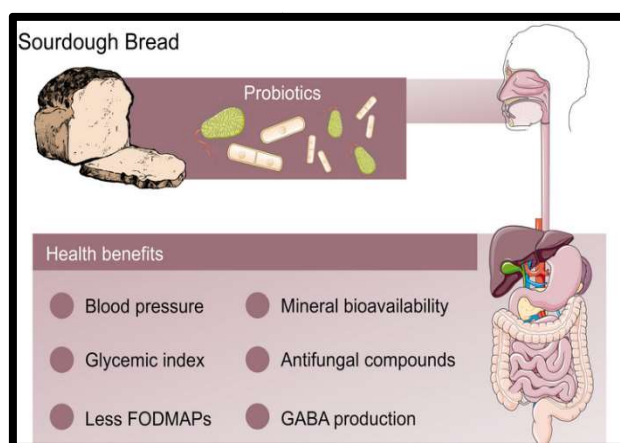


Fig 2 Impact Of Sourdough

Probiotics in the sourdough bread fermentation (Akamine, I. T., et al. 2023)

Conclusion

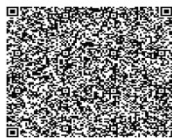
Sourdough bread represents the perfect blend of tradition, science, and nutrition, serving as a timeless example of how ancient methods can still hold significance today. Its distinctive fermentation process not only produces bread with unmatched flavor, texture, and aroma but also boosts its nutritional value, making it a favored option for health-conscious individuals. From its origins in ancient cultures to its revival in both global and Indian markets, sourdough illustrates the lasting relationship between food, culture, and wellness. As bakers keep pushing the boundaries by incorporating various ingredients and improving fermentation methods, sourdough remains a flexible and changing element within the culinary landscape. It showcases the capacity of natural processes to enhance health, sustainability, and the overall dining experience. Ultimately, sourdough transcends mere bread; it stands as a representation of heritage, skill, and the seamless connection between history and the future.

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From Spores to Clues: Fungi, a Potential Tool in Crime Investigations

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Introduction

Fungi are eukaryotic organisms that have unique cell structures and modes of nutrition that distinguish them from plants by lacking chlorophyll and being heterotrophic, deriving nutrients through absorption. Their cell walls are mainly composed of chitin, a robust and sturdy polymer that provides structural integrity. They play a crucial role in ecosystems as decomposers, pathogens, as well as symbionts.

Fungi exist in a variety of forms, including lichens, unicellular (yeasts), and multicellular organisms (moulds and mushrooms). These organisms thrive in sundry environments, often playing climacteric roles as decomposers, symbionts, or pathogens. They have complex life cycles, involving both asexual and sexual reproduction, which contributes to their adaptability and ecological success.

They have been understudied concerning their potential in forensic applications. These eukaryotic organisms, discerned by their adaptability and ecological versatility, have increasingly established their value as forensic tools. Forensic mycology is the application of fungal analysis in criminal investigations, which includes exploiting their unique biological traits to address pivotal questions in cases of clandestine burials, toxicological assessments, and environmental trace evidence. The dearth of published data in this field is explained by the scarcity of forensic investigators with training in it. Most justice professionals are unaware of the role that fungi play in these investigations.

Morphology and Structure of Fungi

The morphology of fungi includes distinctive structures that explicate their growth and reproduction:

Hyphae:

Hyphae are the building blocks of fungi. Fine, microscopic tubular filaments that spread out like underground roots, instead of absorbing water and nutrients like plant roots. They secrete enzymes to break down organic material. Hyphae grow at their tips, branching and intertwining to form vast networks. They can be:

- **Septate:** Compartmentalized by cross-walls (septa) with small pores for nutrients and organelles to pass through.
- **Coenocytic:** Continuous tubes without septa, containing multiple nuclei.

Hyphae provide fungi with an extraordinary ability to invade substrates, like soil, decaying wood, or even living organisms, making them formidable decomposers and sometimes pathogens.

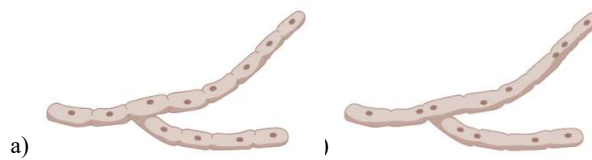


Fig 1: Hyphae; a) Septate b) Coenocytic

Source: Created with BioRender

Mycelium:

When many hyphae come together, they form a mycelium; a dense, interconnected network that's often hidden within the substrate (like a log or soil). It is the body of the fungus, tirelessly working behind the scenes. It's known as the engine of the fungal organism, responsible for absorbing nutrients from the environment.

The mycelium can spread over vast areas, adapting its structure to maximize resource acquisition. Some are so large they can cover several acres, like the famous *Armillaria spp.* (honey mushroom), one of the largest living organisms on Earth. Mycelium is typically hidden within the substrate and is responsible for nutrient acquisition and decomposition.

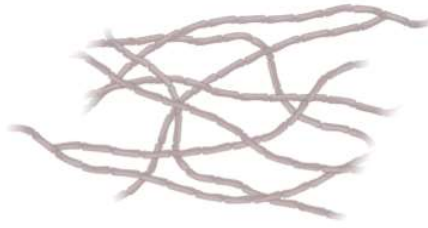


Fig 2: Mycelium

Source: Created with BioRender

Spores:

Spores are nature's dispersal and reproductive units that can be either produced sexually or asexually. They are highly resistant to hostile conditions. Spores are like the seeds of fungi but much simpler and more diverse. They are microscopic, resilient, and designed to travel. They can be:

- **Asexual:** Produced by mitosis, often forming in specialized structures like sporangia or directly on hyphae.
- **Sexual:** Created through meiosis, typically resulting in genetic variation. These are housed in structures like asci (in sac fungi) or basidia (in club fungi).

Spores are released into the environment to colonize new territories. They can travel through air, water, or via animals. When conditions are favourable (like moisture, warmth, and nutrients), a spore germinates, giving rise to new hyphae and starting the fungal lifecycle anew.

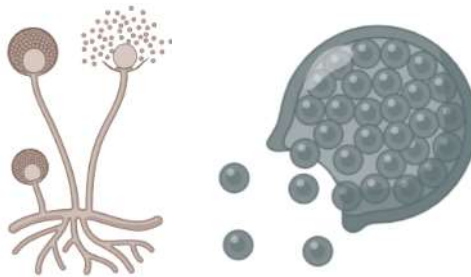


Fig 3: Spore release

Source: Created with BioRender

Fruiting Bodies:

They are macroscopic reproductive structures such as mushrooms, puffballs, and brackets. These structures produce and release spores and are often species-specific. It is the above-ground superstar designed to produce and release spores. They vary widely in size, shape, and colour, often optimized for spore dispersal. For instance:

- Mushrooms have gills or pores under their caps, where spores are formed and released.
- Puffballs release spores in a cloud when the outer skin is ruptured.
- Bracket fungi grow on trees, releasing spores from small pores underneath their shelf-like structures.

These structures are temporary, appearing only when conditions are right for reproduction. The rest of the fungus continues to live on as the hidden mycelium.



Fig 4: Fruiting bodies a) Mushrooms b) Puffball c) Bracket fungi

Source: a) Created with BioRender

b) <https://www.wildfooduk.com/mushroom-guide/common-puffball/>

c) <https://fineartamerica.com/featured/stacked-bracket-fungi-douglas-barnett.html>

Cellular Features:

Fungal cells are characterized by the presence of membrane-bound organelles such as:

- **Cell Walls:** Composed of chitin (not cellulose like plants), which gives strength and flexibility. This tough material is also found in insect exoskeletons.
- **Plasma Membrane:** Contains ergosterol, a unique molecule similar to cholesterol in animals, which is a target for antifungal drugs.
- **Nuclei:** Fungal cells can be multinucleate (coenocytic) or have just one nucleus per cell (monokaryotic). Sexual stages often involve a unique dikaryotic phase, where two nuclei coexist without fusing immediately.
- **Vacuoles and Vesicles:** Vital for storage and transport, especially for nutrients and enzymes.
- **Mitochondria:** Like other eukaryotes, fungi have mitochondria for energy production, supporting their rapid growth and metabolism.

Notably, the cell wall contains chitin, while ergosterol is found in the cell membrane, both of which serve as hallmarks that set fungi apart from other eukaryotic organisms.

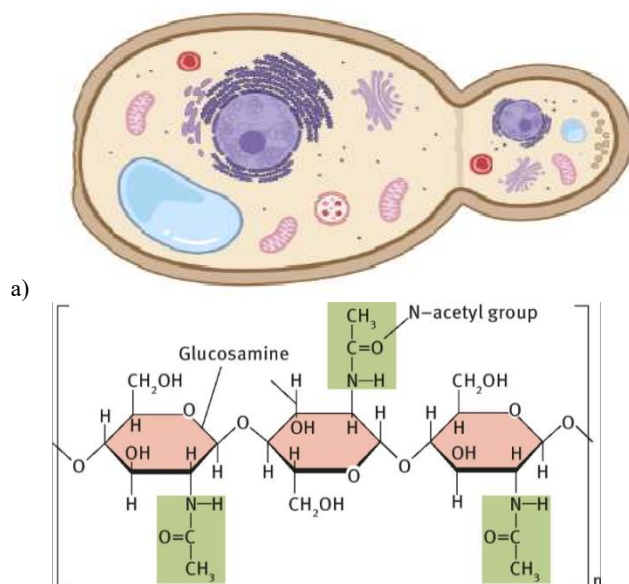


Fig 5: a) Fungal cell b) Chitin structure

Source:a) Created with BioRender b) <https://ibiologia.com/chitin/>

Potential of Fungi as a Forensic Tool

Fungi colonize cadavers in distinct successions, depending upon environmental factors like temperature, humidity, and availability of substrate. This colonization can provide a timeline for decomposition.

- **Locating Burial Sites:** Mycelial networks and fungal fruiting bodies may indicate the presence of decomposing organic matter underground, helping to locate clandestine graves.
- **Tracing Origins of Evidence:** Unique fungal profiles on items like shoes, clothes, or soil can link suspects to specific locations.
- **Identifying Environmental Conditions:** Fungal species are sensitive to environmental conditions, and their presence or absence can help reconstruct crime scenes. Many fungi grow only at a particular time of the year so the environment condition could be predicted accordingly.

The distinctive biological characteristics of fungi render them exceptionally suitable for forensic purposes. Notable features include:

- **Swift Growth and Versatility:** Fungi possess the ability to thrive on a variety of substrates, ranging from organic matter to inorganic surfaces, across a multitude of environmental conditions.
- **Substrate and Condition Specificity:** Certain fungal species are restricted to specific substrates or environmental conditions, which assists in determining the origin or age of forensic evidence.
- **Durability of Spores:** Fungal spores exhibit remarkable resilience to environmental challenges and can remain viable for extended periods, serving as dependable forensic indicators.
- **Varied DNA Profiles:** Progress in molecular biology facilitates the genetic identification of fungal species through methods such as polymerase chain reaction (PCR) and DNA sequencing, allowing for precise identification even from samples that are small or less in quantity.

For example, the growth of mould on a corpse can help estimate the PMI (post-mortem interval). In cases of clandestine burials, the presence of fungal fruiting bodies, such as mushrooms or moulds, frequently signifies areas that have been disturbed. Forensic investigators have also used fungal DNA to establish links between evidence and suspects, as fungal species often exhibit location-specific patterns.

Another critical domain involves investigations related to bioterrorism, where pathogenic fungi may provide evidence of biological attacks. Additionally, moulds like *Aspergillus* spp. and *Penicillium* spp. have been

examined to evaluate indoor air quality at crime scenes or to ascertain the timeline of contamination.

Phases of Decomposition of a Cadaver

The process of human cadaver decomposition is intricate and commences shortly after death. Initially, autolysis takes place, instigated by a rise in carbon dioxide levels within the cells, that results in increased acidity. This decline in pH leads to the rupture of lysosomes, releasing enzymes that promote the enzymatic digestion and disintegration of cellular structures. Following cell lysis, the outflow of nutrient-rich cellular fluids fosters putrefaction. The body's inherent microbiota, especially the flora from the intestines and lungs, contribute to the breakdown of soft tissues, disseminating throughout the body via the lymphatic and circulatory systems. The accumulation of gases produced as byproducts of microbial activity within body cavities leads to bloating. Decomposition occurs at a more rapid pace during the warmer months of spring and summer compared to the cooler months of autumn and winter.

After the active decay phase, the remaining skeleton and any low-nutritive connective tissues begin to desiccate, a process known as mummification, while bones gradually deteriorate through diagenesis (decay). After death, bones are vulnerable to colonization by bacteria and fungi. These microorganisms release enzymes that break down the organic components of bone, particularly collagen, leading to increased porosity and weakened structure. The degree of microbial degradation is determined by environmental factors such as soil pH, temperature, and moisture levels. The decomposition of cadavers in soil is notably enhanced by fungi, which serve as primary decomposers in nearly all terrestrial ecosystems. This process of decomposition significantly influences the populations of soil-dwelling organisms in the vicinity of the corpse, thereby altering both their biotic and abiotic environments. Substantial quantities of organic compounds are released into the surrounding area, with the composition of these compounds varying according to the stage of decomposition. While certain microbial species may utilize these compounds as substrates, others may experience inhibition due to changes in pH or other environmental factors. Consequently, the composition of the surrounding soil microbial community may undergo successive alterations as a result of the changes in substrate availability.^{a)}

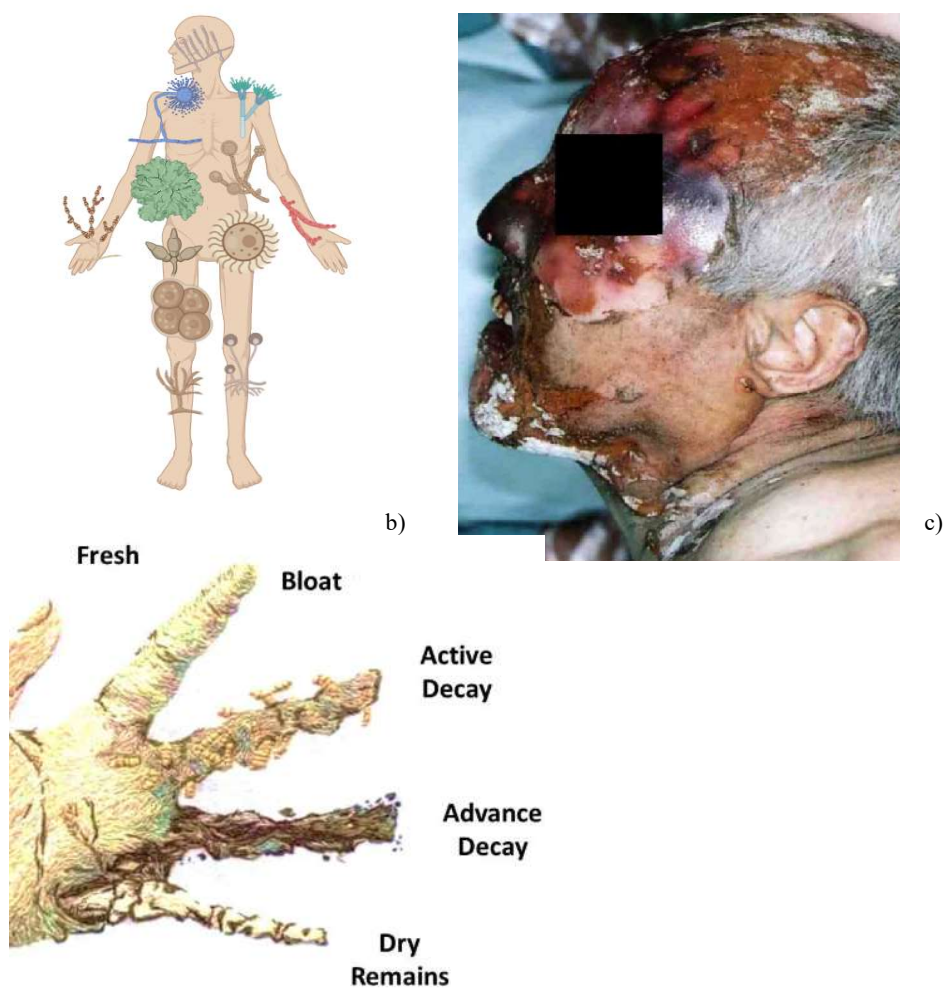


Fig 6: a) and b) Cadaver with various fungi, c) Phases of cadaver decomposition

Source: a) Created with BioRender b) Taken from Masahito Hitosugi et al., 2006 c) <https://forensicfield.blog/tag/dead-body-decomposition-stages/>

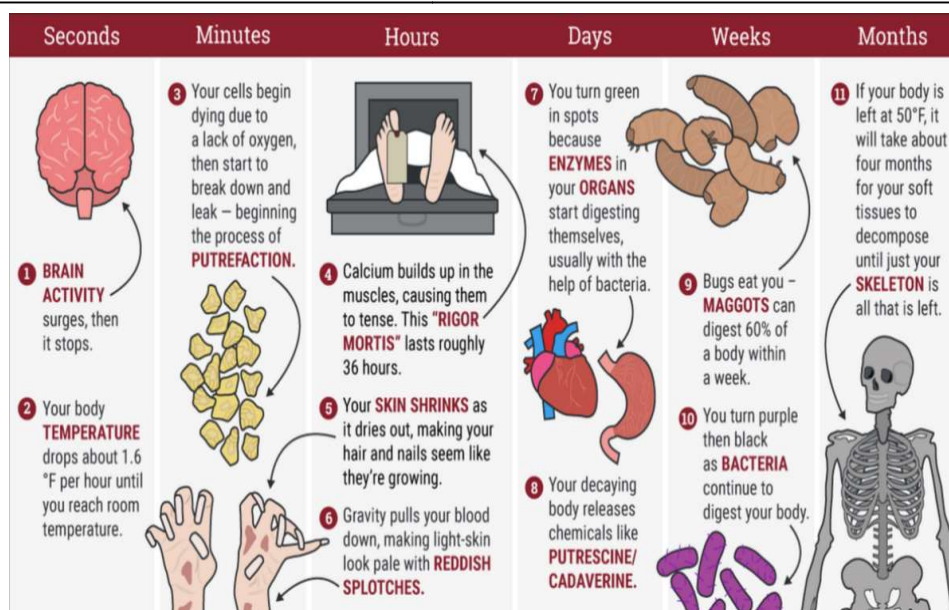


Fig 7: Changes observed in the body after death

Source: <https://www.businessinsider.com/how-human-bodies-decay-2015-10>

Trace Evidence

Fungal spores adhere readily to surfaces and can be transferred between objects, individuals, and environments, rendering them powerful trace evidence. To retrieve these spores, forensic investigators typically employ adhesive tape lifts or swabbing techniques, ensuring minimal contamination during collection. Samples are then subjected to microscopic examination, where spore morphology including size, shape, and ornamentation is analyzed using light or scanning electron microscopy (SEM). Comparative analysis of fungal assemblages from crime scenes, victims, and suspects can establish links with a high degree of specificity. Advanced techniques such as DNA barcoding are also employed to identify fungal species at a molecular level, especially when the morphological differentiation proves challenging. When habitat-specific or rare fungal spores are identified through these methods, it provides compelling evidence in forensic cases.

For instance, spores of *Periconiaspp.* and *Torula herbarum*, fungi associated with stinging nettles, were recovered from a crime scene as well as a suspect's vehicle, corroborating the connection. Similarly, the presence of fungal spores at a burial site and on a weapon reinforced the association between the suspect and the crime. The evidential value of fungi is further

amplified by their habitat specificity and rarity, which often distinguish them from more common palynomorphs (organic remains, including microfossils, and microscopic fragments of mega-organisms that are composed of acid-resistant organic material).

Estimating Post-Mortem Interval (PMI)

The post-mortem interval (PMI) denotes the time duration that has passed since an individual's death, serving as a vital element in forensic analysis. The presence of fungal growth on decomposing human remains offers a distinctive method, as certain species tend to colonize cadavers in predictable sequences influenced by environmental conditions such as temperature, humidity, and the nature of the substrate. Noteworthy species, including *Penicillium*spp. and *Aspergillus*spp., are recognized for their reliable colonization timelines under comparable circumstances.

The methodology for estimating PMI through fungi involves examining the growth stages and species diversity of the fungal communities that inhabit the cadaver. Initially, rapidly proliferating opportunistic fungi, such as *Penicillium*spp. and *Mucor*spp., are predominant, often appearing within days post-mortem. As time progresses, these are replaced by slower-growing and more specialized decomposers, including *Aspergillus*spp. or *Cladosporium*spp. The order and timing of these fungal successions can be correlated with environmental variables like temperature and humidity to refine the estimation of the time of death.

Moreover, fungal growth on bodies submerged in water has been instrumental in distinguishing various submersion periods. Aquatic fungi, such as certain *Fusarium*spp. and *Pythium*spp., colonize remains in water through distinct phases, influenced by factors such as water temperature and oxygen availability. The growth patterns of these fungi, along with those adapted to high humidity, provide further specificity, yielding insights into the length and conditions of submersion. Nevertheless, despite these advancements, the accuracy of PMI estimation through fungi is still constrained by the variability in fungal succession across different environmental settings. Controlled studies mimicking the specific conditions of death are critical to refine and standardize fungal-based estimation of PMI.

For instance, in a landmark case, the presence of fungal colonies on the skin of a cadaver suggested that death had occurred at least 18 days before the body was found, a timeline that was supported by the confession of the suspect.

Experiments were carried out at the University of Tennessee's Forensic Anthropology Centre. The primary experiment included sampling and analysis of the cadaveric soil at regular intervals whereas the secondary experiment included sampling and comparative analysis of the cadaveric soil of two cadavers that varied in weight. T-RFLP (Terminal restriction fragment length polymorphism) and PLFA (Phospholipid-derived fatty acid) analysis were done. The results of the primary experiment were visually proved and went through the stages of decomposition (fresh, bloat, active decay, post decay, and skeletonization) with evidence of a succession occurring and in the secondary experiment, some physical variations in the decomposition process were observed due to the two cadavers used in this experiment, differing in size. The first cadaver underwent an extremely swift process of decomposition, resulting in substantial skeletal exposure. This rapid decay was considered unusual and is likely associated with the cadaver's obesity. Conversely, the second cadaver exhibited a markedly slower decomposition rate, with little abdominal swelling and considerable fungal proliferation on the skin. Renal failure was identified as the cause of death for this individual, whose underweight condition is likely attributable to an extended illness prior to death.



Fig 8: DNA fragment analysis

Source: Created with BioRender

Determining the Time of Deposition

Fungi can also assist in establishing the time of deposition of remains or evidence through a systematic approach. This involves analyzing fungal growth patterns and their ecological context. Seasonal fungi, such as *Flammulinavelutipes*, which fruit during winter, serve as temporal indicators. Investigators study the developmental stages of fungal colonies, correlating them with known growth rates and environmental conditions such as

temperature and humidity. Such assessments provide precise timelines for the deposition of evidence or remains.

Lichens, which are symbiotic organisms composed of fungi and photosynthetic partners, offer another dimension to forensic analysis. Their extremely slow growth rates make them reliable indicators of long-term exposure. For instance, the colour change and diameter of lichen colonies on skeletal remains or associated objects can provide estimates of exposure periods. Additionally, lichen colonization can indicate environmental stability or disturbances, aiding in crime scene reconstruction. When combined with fungal analyses, lichens provide a comprehensive timeline and enhance the robustness of forensic interpretations.

Fungal Poisoning and Hallucinogens

Fungi are frequently implicated in poisoning incidents, whether through accidental ingestion or deliberate use. Poisonous fungi, such as *Amanita phalloides* (death cap), produce potent secondary metabolites like amanitins, which disrupt cellular processes and often result in fatal liver failure. These incidents typically occur when individuals consume toxic species misidentified as edible species.

The importance of identifying the fungi involved cannot be overstated. Accurate identification enables investigators to pinpoint the cause of death and establish whether the poisoning was accidental or intentional. This is achieved by examining fungal spores or fragments recovered from stomach contents, vomitus, or associated environmental samples. These biological remnants are analyzed using advanced microscopy techniques and chemical assays to detect the specific toxins.

In forensic contexts, the systematic approach to fungal identification involves a combination of methods tailored to maximize precision and reliability. Detailed morphological examination remains foundational, where microscopic features such as spore shape, size, and ornamentation are scrutinized using light or scanning electron microscopy. Biochemical analyses, including chromatography (e.g., gas chromatography-mass spectrometry or liquid chromatography), are employed to detect and quantify toxic compounds or unique secondary metabolites. Spectroscopic methods such as Fourier-transform infrared (FTIR) and nuclear magnetic resonance (NMR) spectroscopy add another layer of specificity by characterizing chemical structures.

Molecular techniques, such as DNA barcoding and next-generation sequencing (NGS), have revolutionized fungal identification, allowing species-level classification even from minimal or degraded samples. These advanced methods complement traditional approaches, enabling comprehensive profiling of fungal communities and precise identification of toxic or hallucinogenic species. The integration of these diverse methodologies ensures reliable identification, which is critical for both medical interventions and legal investigations.

Hallucinogenic fungi, particularly *Psilocybespp.*, pose additional forensic challenges due to their production of psychoactive compounds like psilocybin and psilocin. Psilocybin, a phosphorylated compound, is a prodrug that undergoes dephosphorylation in the body to form its active metabolite, psilocin. This conversion primarily occurs in the liver through the action of alkaline phosphatase and nonspecific phosphatases. Psilocin, the pharmacologically active compound, readily crosses the blood-brain barrier and interacts with serotonin (5-HT) receptors, particularly the 5-HT_{2A} receptor subtype. This interaction alters neurotransmitter signaling, leading to the characteristic effects of hallucinations, altered perception, and euphoria.

The metabolic pathway of psilocybin to psilocin is combined with its rapid absorption in the gastrointestinal tract and subsequent systemic distribution. These effects, while sought for recreational purposes, present legal and medical challenges, particularly in cases of misuse or overdose. These properties have made such fungi a focus of interest in both recreational use and legal enforcement.

Forensic investigations often require precise identification of hallucinogenic fungi, as their possession or distribution is illegal in many jurisdictions. This identification is typically achieved through microscopic examination of spore and tissue samples, revealing diagnostic features unique to the species. Chemical analyses, such as gas chromatography-mass spectrometry (GC-MS) or liquid chromatography-mass spectrometry (LC-MS), are employed to detect and quantify psilocybin and psilocin concentrations. These methods not only confirm the presence of psychoactive compounds but also provide evidence for legal proceedings and toxicological studies.

Understanding the pharmacological effects and distribution of these compounds is critical for both law enforcement and public health interventions, as improper identification can lead to legal or medical consequences.

Locating Buried Corpses

Fungi associated with disturbed soil environments, such as *Hebelomaradicosum*, have demonstrated significant potential in the systematic identification of clandestine burials. Known as “corpse-finder” fungi, these species thrive in nutrient-enriched soils above decomposing remains, where they utilize the decomposition products as a nutrient source. Notably, this species is known to appear 3–8 months after the application of urea or ammonium to soils and can continue to produce mushroom sporophores for 1–2 years. This ecological association makes them valuable indicators of burial sites.

The process begins with surveying the area for fungal growth, focusing on species that are known to proliferate in disturbed or enriched soils. Soil sampling is then conducted systematically from suspected sites, with samples examined for fungal spores and hyphae. Advanced methods, such as DNA sequencing and morphological analysis, are employed to identify specific fungi associated with human decomposition.

Changes in soil characteristics also play a pivotal role in detecting disturbances. Trampled or excavated areas often exhibit altered soil compaction, disrupted vegetation, and shifts in microbial and fungal communities. Observing fungal fruiting body orientation can reveal disturbances: for instance, logs or branches moved during burial may show reoriented fungal growth as the organisms adapt to changes in light and gravity. Additionally, the presence of fungi unique to decomposing organic matter, coupled with chemical soil analysis for elevated levels of nitrogen or phosphorus, provides further confirmation of burial sites.

Fungal growth patterns can also indicate environmental disturbances. For example, changes in the orientation of fungal fruiting bodies on displaced logs or branches have been used to refute alibis in forensic investigations. These subtle ecological clues often reveal tampering at crime scenes.

While the forensic application of such fungi is still emerging, their ability to complement traditional geophysical techniques, such as ground-penetrating radar or cadaver-detection dogs, enhances the precision and reliability of clandestine grave detection. It can also determine the possible path the offender has taken at the crime scene.

Biological Warfare and Toxins

The use of fungi in biological warfare stems from their ability to produce a wide range of mycotoxins. They are chemical compounds that can cause significant harm to living organisms. Fungi's ecological versatility and ease of cultivation make them particularly attractive as agents of biological weapons. Many fungal toxins are stable under environmental conditions and can be produced in large quantities with minimal resources. Some fungi naturally generate toxins that act quickly, causing acute effects, while others produce compounds with long-term impacts, such as carcinogens. This duality amplifies their potential misuse in warfare and terrorist activities.

Fungi's ability to serve as bioweapons lies in their adaptability. Through vat cultivation, certain moulds and mushrooms can yield high concentrations of toxins within controlled environments. Vat cultivation involves growing the fungi in large containers under carefully controlled conditions. These vats ensure the right temperature, humidity, and acidity for optimal fungal growth.

There is an urgent need for monitoring and regulating fungal strains in research and industrial applications. The risk is exemplified by historical cases: for instance, fungi were initially and wrongly implicated in the "Yellow Rain" phenomenon in Vietnam. However, a more critical incident occurred within the last 30 years when a Middle Eastern country was found attempting to procure toxin-producing strains of *Fusarium spp.* from microbial culture collections in North America and Europe.

A detailed review of toxin-producing fungi as potential biological weapons has identified the Fusarium T2 toxin as particularly dangerous. This toxin is a trichothecene mycotoxin, highly potent, capable of causing acute poisoning and severe health consequences, emphasizing the need for robust monitoring and preventive measures. Historical cases further underscore the necessity of interdisciplinary collaboration between mycologists, national security agencies, and microbial culture repositories to mitigate the risks associated with such biological threats.

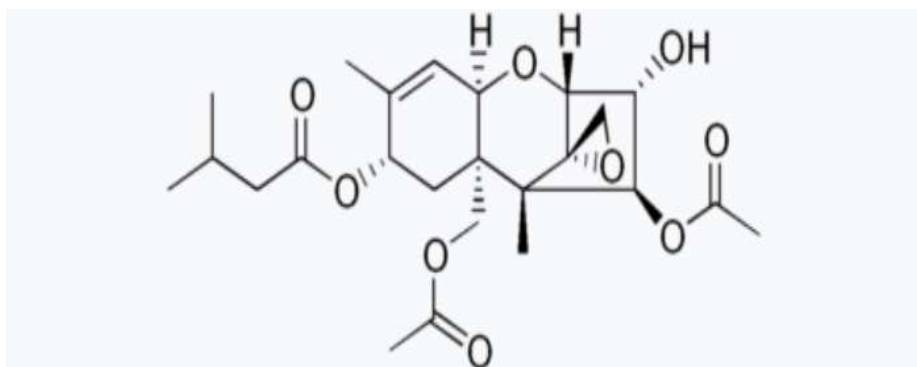


Fig 9: Structure of T2 toxin

Source:https://en.wikipedia.org/wiki/T-2_mycotoxin

Advancing Forensic Mycology

Forensic mycology's potential remains underexploited due to the limited awareness among investigators and a shortage of trained mycologists. Furthermore, replicating exact scenarios and environmental conditions in which the body was found at the crime scene to decode the hidden mysteries behind the crime is quite difficult. Expanding expertise in fungal taxonomy, coupled with investments in research and molecular identification technologies, is imperative for integrating mycology into mainstream forensic science.

Research into fungal ecology, growth dynamics, and species interactions under diverse environmental conditions will enhance the reliability of mycological evidence. Additionally, fostering collaborations between forensic scientists, mycologists, and law enforcement will facilitate the development of standardized methodologies and protocols.

Conclusion

Fungi offer unparalleled opportunities in forensic investigations such as providing novel approaches to trace evidence analysis, PMI estimation, toxicological evaluations, crime scene reconstruction, and much more. As the discipline of forensic mycology matures, it holds the potential to complement and enhance traditional forensic methods, offering investigators innovative tools for solving complex cases. By integrating fungal analyses into forensic protocols, law enforcement can leverage the latent capabilities of these versatile organisms.

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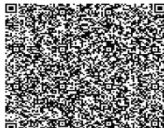
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Recent advances in Nanomaterials for Sustainable development

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Abstract

Nanomaterials are structured materials in nano dimension with unique properties have revolutionized diverse fields in the last decades. Development of nanomaterials for the sustainable growth of society has become a global challenge. One of the promises associated with nanomaterials and nanotechnology is to facilitate greener methods for sustainable development. Nanomaterials possess dimension between 1-100nm with different physical properties compared to bulk counterparts. The properties of these material finds promising for the sustainable development in the field of energy, water, medicine, food decontamination and agriculture. Nanotechnology has the potential to improve the environment through nanomaterials This book chapter summarizes the recent advances in nanomaterials and their role in sustainability. The first few sections of the chapter deal with the application of nanomaterials in renewable energy, environmental remediation, water purification and conservation. In addition, the book chapter includes application of nanomaterials for sustainable agriculture and therapeutics. It also includes biodegradable and eco-friendly nanomaterials for sustainable development. The last section summarizes on the health, environmental safety and future trends in nanomaterials for sustainable development.

Overview of Nanomaterials and their role in Sustainability

Over the past three decades, nanomaterials have revolutionized the field of material science, with the capability to characterize materials at the atomic scale. However, it quickly progressed as scientists were able to visualize matter at the nanometer scale which also enabled them to manipulate it. This breakthrough eventually led to the synthesis of nanomaterials with varied compositions, sizes, shapes and structures [1-4]. The development of

synthesized nanomaterials marks a significant advancement in material science by tuning their optical, magnetic, tribological and chemical properties. As a result, these materials have transitioned from the laboratory to numerous technologies and industries. Despite, the existence of nanomaterials that support sustainable technologies such as batteries and electrolyzers, the full potential of nanoscale materials in advancing sustainability has likely not been realized. This is especially true considering the vast and infinite potential for exploring the relationships between nanostructures, properties and composition. The increasing global demand for energy, coupled with growing environmental concerns about climate change have sparked major interest in recent years in transitioning to a low-carbon economy that relies on renewable sources such as wind and solar power. Energy existing in various forms can be harvested, converted, managed or stored. A primary focus of the scientific community is to improve the efficiency of energy harvesting and conversion, as well as enhancing the longevity of energy-related devices. In addition, it is crucial for us to prioritize the use of abundant and recyclable materials while minimizing the reliance on toxic materials. As our understanding of the nanoworld has progressed, new considerations were arisen on how to effectively utilize the properties of nanomaterials. This helps to fully optimize their potential in improving the sustainability of the planet. A key challenge is to develop sustainable methods for synthesizing nanomaterials, incorporating them into essential technologies that offer direct societal benefits, leveraging their properties to create new functionalities while minimizing their potential environmental impact during the synthesis, use and disposal. The advancement of material science over the last three decades has become one of the leading factors of the energy transition, enabling the development of a wide variety of new materials with different compositions, dimensions and geometries. Nanomaterials have quickly introduced new paradigms and phenomena, which are beginning to make a significant impact on the sustainable energy landscape.

Applications of Nanomaterials in Renewable Energy

The vast global energy consumption and substantial carbon dioxide emissions have opened a new frontier for research on sustainable and renewable energy. Today, one of the greatest challenges facing humanity is the transition to renewable energy and the creation of a sustainable, pollution-free society. Various forms of renewable energy such as geothermal, solar, wind, biofuel and hydropower have been explored and advanced by humans. This drive has led to the development of nanomaterials for renewable energy applications. Carbon nanomaterials are widely used in energy storage systems due to their excellent electrical and mechanical properties enabling them to

store charges effectively. These carbon nanomaterials have been employed as electrodes in fuel cells, solar cells, lithium –ion batteries and super capacitors[5-7]. Semiconductor nanomaterials offer several advantages due to their increased surface area, enhanced optical absorption, shorter charge migration distance, increased solubility and tunable electronic structure. In recent years, nanomaterials are used for developing hybrid nano-biocatalytic system by changing the feed materials and biocatalysts for generation of fuels[8]. Graphene based nanomaterials are widely used in electro catalysis, sensors and hydrogen production. Semiconductor nanomaterials are used for solar hydrogen production due to higher surface area, optical absorption, solubility, shorter charge migration, plasmonic resonance and tunable electronic structure [9,10]. TiO₂ nanorod arrays were used as working electrodes for photoelectrochemical cells for hydrogen production [11]. Borophene nanomaterials are quite different from 2D nanomaterials like graphene due to their vacancy defect and negative poisson's ratio. So, borophene nanomaterials are utilized in metal-ion batteries and supercapacitor applications [12]. Due to the higher surface area and shorter charge migration length, nanomaterials are widely used as electrode in Dye sensitized solar cells. Nanotubes are found to be better compared to nanorods and wires. This is because; the inner and outer walls of the tubes are readily available for absorption of dyes [13-24]. Bamboo based carbon nanocomposites reported by Kim et al showed higher efficiency than smooth walled tubes in DSSC applications [25].

Applications of Nanomaterials in Environmental Remediation

The rapid expansion of global population has led to an unprecedented demand for energy, food, chemicals propelling industrialization to new levels. However, this has resulted in production of enormous amount of waste water. Waste-water effluents from industries such as paper, pulp, oil, fabric, brewery, food and beverages present a severe threat to both public health and the environment. Improperly treated industrial effluents, when discharged into the ecosystem, causes significant harm to water quality. This not only disrupts the ecological equilibrium but also makes water contaminated and unsafe for everyday purposes having detrimental human health and environmental stability. Metal and Metal Oxide nanomaterials play a very important role in removing recalcitrant from the waste water. Their degree of removal can be improved through functionalization with organic groups that can selectively target pollutants present in air and water. Production of safe drinking water, water recovery, removal of metals, dyes, pesticides from the waste water can be achieved using nano membranes. Nano fibrous media can be used for

filtration applications due to the less weight, high permeability and small pore size. Good interconnectivity of pores and higher surface area of nanofibrous membranes helps in the removal of microparticles from waste water. Nano filtration process is commonly used for removing organic matter. Environmental pollution has become a worldwide menace that endangers the community and ecological sustainability, thereby escalating the air quality values provided by WHO causing chronic respiratory conditions, lung cancers, cardiac diseases and so on [26,27]. So, it is crucial to monitor the impact of pollutants on plants, animal, humans and aquatics. Owing to the interaction of light with nanostructured materials, nanoplasmonic optical sensors helps in detecting the pollutants with higher sensitivity. Heavy metals and Volatile Organic Compounds accumulated in the environment through food chain are found in air, water, soil, cosmetics and food causing acute or chronic illness [28, 29]. Liang Shang et al fabricated imprinted solgel coated on gold nanoisland sensor for detecting cis-jasmone [30]. This sensor will be used as a potential tool for sustainable agriculture [31, 32]. Chinh Van Tran et al synthesized Polyaniline oxide nanoparticles via hydrothermal method were studied for degradation of dyes under sunlight [33]. The development of these sustainable photo catalytic materials are useful for waste water treatment. It is always very much essential to support the United Nations Sustainable Goal for a healthier and sustainable development [34]. Nanofibers, nanotubes, nanoflowers, nanosheets, carbon and non-carbon based nanomaterials act as very good adsorbents due to their huge surface area. Fe_3O_4 /activated carbon nanoparticles have been used in removing methylene blue and brilliant green dyes. Chitosen sugar coated magnetitenanoparticles are used in removal of crocein orange G and acid green 25 dyes as they have very good adsorbing capacity [35, 36].

Applications of Nanomaterials for Water Purification and Conservation

Over 1.2 billion people around the globe do not have access to potable drinking water. A need for safe drinking water is a challenging problem due to the rapid increase in population. Most of the traditional techniques are costly and the ones which are affordable produce secondary waste. So, an eco-friendly and economical technologies are required to overcome water pollution and waste water treatment constraints. Introducing nanomaterials improved the efficiency for eliminating the recalcitrant's present in water. Nanomaterials are advantageous due to their large surface area, catalytic, optical, hydrophilic, hydrophobic and anti microbial properties. Therefore, the efficiency of water treatment process is increased by incorporating nanomaterials in the

membrane. Nanomaterials affect the pH and temperature as well which aids in betterment of the treatment process. Nanowires, nanotubes, nanofilms, nanoparticles, bio-based nanomaterials, quantum dots and colloids are employed to remove contaminants from the industrial pollutants, surface water, ground water and drinking water. Biobased nanomaterials such as SiO_2 , cellulose, graphene, chitosan, keratin and other metals can be used in desalination and waste water treatment. Iron nanomaterials have the capacity to remove heavy metals like arsenic from the waste water due to the magnetic nature. The divalent heavy metals such as copper, nickel and cobalt are removed by the polymer grafted Fe_2O_3 nanocomposite. Zinc oxide nanoparticles can be used as a substitute for UV disinfection as it has higher potential to kill microorganisms enhancing water quality. Magnetic nanoparticles, metal nanoparticles, quantum dots, dye doped nanoparticles are used in tracking of pathogens. Cellulose nanocrystals and Cellulose nanofibrils are used in the waste water and desalination treatment. Different bio-silica based nanomaterials synthesized from sugarcane baggage, rice straw, fungus, bacteria, bentonite clay and weeds [37-41] are used in different applications of water treatment. Synthesis of silica nanoparticles using fungi helped in producing vast quantities of enzyme in recent years. Silica and hybrid silica based nanomaterials are used for water filtration due to their stability, low cost and surface area [42]. Different bio-graphene based nanomaterials synthesized from leaf extracts of neem, algae, microalgae, mango leaves, red algae and orange peels have been used for its application in waste-water treatment [43-48]. Graphene quantum dots has received lot of attention among the research due to their physicochemical and photoluminescence properties. Chitosan/nitrogen doped graphene quantum dots degraded protein, bacteria, medicinal waste, dyes and fluoride more efficiently [49]. It was also observed from reusability investigation that percentage removal was maintained at acceptable standards and there was no much decline in the nanocatalyst. Recent studies on chitosan based adsorbents and their adsorption abilities towards heavy metal ions such as arsenic, lead, chromium, cadmium were discussed by Omer et al 2022 [50]. Chitosan based nanocomposite polymeric membranes helps in removal of organic pollutants from waste water. Surface modification of chitosan with graphene oxide nanocomposite makes it an effective intriguing adsorbant material for removal of arsenic in waste-water [51]. Nanomaterials such as zinc oxide, magnesium oxide, titanium oxide and ferric oxides act as very good photo catalysts as it has the capacity to adsorb contaminants from waste water due to their high surface area, pore size and shorter diffusion distance [52, 53].

Application of Nanomaterials for Sustainable Agriculture

By 2050, the global population will grow to 10 billion which likely result in 50% would increase in demand for food especially in low income countries as projected by Food and Agricultural Organization. Current agricultural systems are confronted with many challenges like rare diseases, increased temperature and salinity, imbalance in the nutrients and environmental pollution. Frana et al.[54] analyzed and found several were malnourished and the expected demand for food in developing countries will raise. So, there is a need for developing countries to enhance its agricultural productivity and efficiency by implementing sustainable food production. The present agricultural practices use massive resources for pesticides, fertilizers, water and energy [55]. It is not possible to achieve Sustainable Development Goals using the present farming systems. Also, sustainable agriculture is necessary to attain Zero Hunger SDG framed by United Nations .Therefore; nanomaterials designed for agriculture should adhere to biocompatibility, environment friendly and therapeutic compatibility. Eco-friendly synthesis of nanomaterials provides the potential required for the sustainable development of agriculture [56-58]. Traditional fertilizers and pesticides has been replaced by nanomaterials owing to the increased crop yield, chlorophyll content in leaves, longer root and shoots stress tolerance, disease detection in plants and soil remediation. Nanofertilizers, nanopesticides and nanosensors add to sustainable crop efficiency, quality and protection [59].Introducing carbon nanomaterials to plants influenced plant physiology in numerous ways. Carbon nanofertilizers has shown their development in germination and growth of seeds, enhancement of shoots, roots, chlorophyll, rate of photosynthesis, improved enzymatic activity and the overall productivity of crops. Carbon nanohorns are cone like structures with good micro porosity and higher surface area which make them act as adsorbants and carriers. Water soluble nanoparticles showed maximum growth on wheat plants as investigated by Saxena et.al 2014 [60].Mutiwall carbon nanotubes showed positive effect in many crop varieties such as rice, wheat, corn, peanut, garlic and corn [61-63]. Lateef et al, 2019 [64] and his coworkers synthesized Nanocomposite based biochar from corn waste and was able to find that these composites are suitable in providing nutrients to plants. In addition, nanocomposite biochar showed better water retention,absorption and slow release pattern of nutrients than corn biochar by providing nutrients for a longer time making them act as sustainable fertilizers. Yang-Yang Gao et al analyzed that the design and applications of fluorescent chemosensors helps in regulating plant health and environment [65]. Many research groups are still working on the interaction between plant-

nanomaterials and their levels of toxicity. Much more research is required to understand the tolerance level of different type of carbon nanomaterials in various crops for sustainable production.

Sustainable Nanomaterials for Therapeutic Applications

Inorganic medicinal nanoarchitectonics represent an innovative approach in health care due to its therapeutic applications. The unique properties of engineered inorganic nanomaterials such as stability, biocompatibility and multifunctionality interact with the biological systems at the molecular level. Hybrid systems based on medicinal nanoarchitectonics have been applied in photothermal therapy, photo-dynamic therapy, bio-imaging, drug delivery and so on. Inorganic nanomaterials are emerging as a tool for development of novel anti-cancer drug and vaccines. Lanthanide doped up-conversion nanoparticles are used as an alternate for bio-imaging applications [66]. Quantum dot nanocrystals are inorganic semiconductor materials from group 2 and 4 or group 3 and 5 elements with the size ranging from 2nm -8nm. The unique optoelectronic property of quantum dot nanocrystals shows its potential for detection of diseases in the early stage, targeted drug delivery and non-invasive cellular imaging and tissue engineering applications [67-69]. In recent times, Herceptin bound and Polyethylene glycol coated lecithin-insulin nanogel complex has been a suitable material for the treatment and diagnosis of breast cancer [70]. Similarly, iron oxide nanoparticles, silica nanoparticles, gold nanoparticles, lanthanide doped nanomaterials are finding its applications in cancer theranostics, MRI and CT imaging, target breast cancer cell identification, PET, optical imaging, vascular and tumor imaging and multi-modal theragnosis. Coating the nanoparticles with RBC or cancer cells evades the immune system and enhances therapeutic activity. Cell membrane coated nanoparticles are used as biomimetic vaccines for tumor therapy. Further research on biomimetic nanovaccines is increased due to their stronger immune response and they also act as biocarriers for antigens, adjuvants and active pharmaceutical ingredients [71]. Nanobiopolymers are used as targeted drug delivery agents for cancer therapeutics due to their biodegradability, renewability and sustainability [72]. Bioactive imidazole nanocarriers with drug loading using saponin and apigenin targets cancer cells more specifically compared to conventional chemotherapy. This break-through on cancer therapeutics helps to achieve high fractional cell killing and reduce the side-effects of the normal cells [73].

Biodegradable and Eco-friendly Nanomaterials for Sustainable Development

The regular discharge of effluents from industries contains organic pollutants, dyes, metalloids and heavy metals. Any substance that is generated during production or application can cause a threat to human life. Chitosan nanomaterials are used to treat waste water effluents from pharmaceutical industries due to their biocompatibility, biodegradability, durability, high surface area and low toxicity [74]. Bio based nanomaterials has become an alternate to fossil fuels because of their availability, low cost and sustainability. Silver nanoparticles, zinc based nanoparticles, copper nanoparticles, Chitosan nanoparticles, sulphur nanoparticles synthesized using eco-friendly method was studied for plant disease management applications. This helps in increasing farm inputs and to produce quality products [75]. The vegetable and fruit peel wastes derived enzyme powered TiO_2 and ZnO nanomaterials are considered as promising materials for cost-effective dye degradation and anti-cancer studies [76]. 2D transition metal borides are utilized as catalysts to convert organic compounds into less toxic substances and for electro-reduction applications. It has also been used as catalysts for production of clean energy, batteries, super capacitors and information energy devices. These transition metal borides are also used in medical applications such as biosensing, tumor therapy and neuro degenerative diseases. In addition, it aids in understanding the interaction of nanosheets with the biological system known as nanometabolomics [77]. So, the transition metal borides act as sustainable and cost effective materials for various applications. Researchers are currently focused on bacterial nanocellulose based materials in different types of wound healings and dressings due to their high crystallinity, mechanical strength, water adsorption capacity, porosity, purity, flexibility and sustainability [78]. Biodegradable nanomaterials such as starch nanoparticles have gained interest among researchers due to their non-toxicity, availability, hydrophilicity and sustainability; it can be used as nanofertilizers, nanopesticides and in plant-genetic engineering applications. Green approach for the synthesis of manganese molybdate nanoparticles using *Camellia sinensis* extract enabled these materials as high performance super capacitors making them a promising candidate for energy storage applications in future [79].

Health and Environmental Safety of Nanomaterials

Environmental impact is commonly understood as the outcome of exposure and toxicity. A nanomaterial present in small amounts is unlikely to pose significant environmental risks. However, as nanomaterials gain wider acceptance, the potential for exposure rises, and even substances with relatively low toxicity may have unforeseen effects. The production of nanomaterials increases daily in the global market. So, these nanomaterials possibly will be released to air, water and soil. Rain water may also contain nanomaterials, which disturbs the whole ecosystem. Nanomaterials are classified under the list of emerging pollutants. But some reports on TiO_2 and silver nanoparticles, carbon nanotubes and fullerenes in water treatment plants have reached alarming values [80]. Nanomaterials can permeate soil when fertilizers are used contributing to the problem of pollution. Since, nanomaterials have unique properties, the recyclability and reusability depends on the type of nanomaterial and its application. Safe and effective methods for disposal of nanomaterials should be incorporated without affecting human health and environment. Nanomaterials naturally decompose and they degrade into less harmful chemical over time. These procedures help in reducing the damage to the environment [81, 82]. Though there are theoretical knowledge, frame works and legislation, the consumers must be aware of the benefits and risks associated with nanomaterials. In 2020, the vaccines against SARS-COV-2 Pfizer and Moderna used nanoliposomes as carriers'. It is required to talk about the regulations related to SDG such as good health and well being, dirt-free water and hygiene, inexpensive and clean energy, climate, aquatic life, life on land. The development of these regulations is considered as strategy towards achieving sustainable development goals in all countries. The purpose of the nanomaterials regulations in using nanotechnological products must guarantee the safety of the public and quality of the environment. Owing to the positive and negative impact of nanomaterials, it is crucial to address the risks associated with nanomaterials as it is always very important to guarantee the safety and sustainability while promoting science and technology to save our planet.

Conclusions and Future Trends in Nanomaterials for Sustainable Development

Research on nanomaterials provide incredible opportunities to advance social well being in areas such as energy storage, catalysis, water purification, environmental remediation, therapeutics and agriculture to enhance global sustainability. The major challenges for sustainable development includes

applications of nanofungicides, nanoherbicides and nanofertilizers for promoting sustainable agriculture; use of nanofilter membranes for sustainable management of safe and clean water; using nanomaterials to remove green house gases for combating against climate change, use of nanoarchitectonics, nanobiopolymers, nano-imaging and nanosensing for therapeutic applications and for food packaging and preservation. The development of nanotechnology with Information Communication and Computation technologies (ICCT) will provide automated products and services for individuals in all member countries by 2030, thereby paving the way to achieve the Sustainable Development Goals. These SDG's can be achieved by creating awareness among the youths and expand the funding related to Sustainable Development Goals. So, Green Nanotechnology aids in promoting the use of renewable energy and smart materials, developing sustainable environment by minimizing pollutions caused by soil, air and water, providing clean drinking water using renewable energy, in finding simple and effective drugs for curing all types of diseases, supporting sustainable industrialization for sustainable economic growth, controlling green gas emissions and production of toxic materials to save our planet mother earth. If nanotechnology can be replaced by Green nanotechnology, it is environmentally safe and effective to achieve one of the sustainable development goals by 2030.

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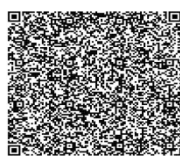
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Real-World Data Analytics for Outcome-Based Healthcare

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1. Introduction

Real-world data (RWD) refers to data collected from various sources outside traditional clinical trial settings, including electronic health records (EHRs), claims and billing data, patient registries, wearables, and mobile health applications. Unlike the controlled environment of randomized controlled trials (RCTs), RWD captures the dynamic and diverse nature of healthcare delivery and patient behavior in real-world contexts. This data provides valuable insights into the actual effectiveness, safety, and utility of healthcare interventions, complementing the findings from RCTs. For instance, RWD helps assess how treatments perform in broader patient populations, such as those with comorbidities or those excluded from clinical trials, providing a more comprehensive understanding of healthcare outcomes (Sherman et al., 2016; Makady et al., 2017).

The importance of RWD has grown with advancements in data collection technologies and the increasing emphasis on evidence-based and outcome-focused healthcare. Regulatory agencies like the U.S. Food and Drug Administration (FDA) and the European Medicines Agency (EMA) rely on real-world evidence (RWE) generated from RWD for post-marketing surveillance, label expansions, and the approval of new indications. Additionally, RWD supports the transition from volume-based to value-based healthcare models by tracking key outcomes such as hospital readmissions, medication adherence, and quality-of-life improvements. While the potential of RWD is vast, challenges such as data quality, interoperability, and privacy concerns must be addressed to fully harness its benefits. With advancements in data analytics and machine learning, RWD is poised to play a transformative role in clinical practice, public health, and healthcare policy-making (FDA, 2021; Sivarajah et al., 2017; Hassoun et al., 2020).

2. Applications of Real-World Data in Pharmacy Practice

Real-world data (RWD) has become a cornerstone for advancing pharmacy practice by providing actionable insights into patient care, medication use, and health outcomes. Unlike controlled clinical trial data, RWD reflects the complexities of everyday healthcare delivery, making it invaluable for addressing challenges in pharmacy practice. The use of RWD enables pharmacists to better understand patient behaviors, medication efficacy, and the broader implications of healthcare interventions (Makady et al., 2017).

One key application of RWD in pharmacy practice is monitoring **medication adherence** and its impact on patient outcomes. Data collected from electronic health records (EHRs), prescription refill records, and digital adherence tools can help identify patterns of non-adherence. Pharmacists can use these insights to design targeted interventions, such as counseling or reminders, to improve adherence rates. For instance, studies leveraging RWD have shown that timely pharmacist interventions can significantly reduce hospital readmissions among patients with chronic conditions such as diabetes and hypertension (Brown et al., 2019).

RWD also plays a pivotal role in **pharmacovigilance**, the science of detecting and preventing adverse drug reactions (ADRs). Traditional methods of ADR reporting often suffer from underreporting and delayed detection. By mining RWD from diverse sources like EHRs, claims databases, and patient registries, pharmacists can identify safety signals earlier and assess the real-world safety profiles of medications. This proactive approach enhances patient safety and supports regulatory decision-making (Sherman et al., 2016).

In the realm of **personalized medicine**, RWD enables pharmacists to tailor therapies based on individual patient characteristics. By analyzing data such as genetic information, comorbidities, and medication history, pharmacists can recommend treatment regimens that are more likely to succeed for specific patients. For example, RWD has been used to optimize anticoagulant therapy in patients with atrial fibrillation by accounting for factors like age, renal function, and prior bleeding events (Smit et al., 2018).

Another important application is in **health economics and outcomes research (HEOR)**. Pharmacists and researchers use RWD to conduct cost-effectiveness analyses, budget impact assessments, and resource utilization studies. These analyses help healthcare systems and payers make informed decisions about the inclusion of medications in formularies and the allocation

of resources. For example, RWD has been instrumental in evaluating the economic impact of newer biologics and specialty drugs in real-world settings (Martin et al., 2020).

Lastly, RWD enhances the role of pharmacists in **population health management**. By analyzing data at the community or population level, pharmacists can identify trends in medication use, emerging health issues, and disparities in access to care. These insights allow pharmacists to participate in designing and implementing public health interventions, such as vaccination campaigns or chronic disease management programs, to improve overall health outcomes (Sivarajah et al., 2017).

3. Predictive Analytics in Outcome-Based Healthcare

Predictive analytics leverages real-world data (RWD) and advanced algorithms to forecast patient outcomes and optimize care delivery. By analyzing historical data and identifying patterns, predictive models help healthcare providers anticipate clinical events, reduce risks, and improve both individual and population health outcomes. This aligns with the goals of outcome-based healthcare, where care delivery is tied to measurable health improvements (Obermeyer & Emanuel, 2016).

A key application of predictive analytics is risk stratification, which categorizes patients based on their likelihood of experiencing adverse events. For example, predictive models identify patients at high risk of hospital readmissions by analyzing factors like comorbidities and medication adherence. Personalized interventions, such as follow-up care or medication adjustments, can then prevent readmissions and improve outcomes (van Walraven et al., 2010). In chronic disease management, predictive models forecast events like hyperglycaemia or hypoglycaemia in diabetes, allowing proactive interventions. Similarly, cardiovascular care benefits from algorithms that predict strokes or myocardial infarctions based on patient-specific data (Goldstein et al., 2017).

In pharmacovigilance, predictive analytics enhances medication safety by identifying potential adverse drug reactions (ADRs) before they occur. By integrating EHRs, prescription data, and demographics, predictive models flag high-risk patients, allowing proactive adjustments to medication regimens. This improves safety and reduces complications (Bate et al., 2018). Another critical application is in personalized treatment plans. For instance, predictive models in oncology assess tumor genetics and prior therapy responses to tailor treatments, maximizing efficacy and minimizing side effects (Topol, 2019).

Despite its promise, predictive analytics in healthcare faces challenges like data quality, model interpretability, and ethical concerns. Addressing biases, ensuring transparency, and safeguarding data privacy are crucial for building accurate and fair models. As healthcare systems increasingly adopt predictive analytics, overcoming these challenges will be essential for maximizing its potential in driving outcome-based healthcare (Rajkomar et al., 2018).

4. Real-World Evidence (RWE) for Regulatory and Payer Decisions

Real-world evidence (RWE) is derived from the analysis of real-world data (RWD) and plays a critical role in informing regulatory and payer decisions in modern healthcare. By offering insights into how drugs, devices, and interventions perform in actual practice, RWE complements the findings of randomized controlled trials (RCTs), which often exclude diverse patient populations and real-world variables. Regulators and payers increasingly recognize the value of RWE in ensuring safety, efficacy, and cost-effectiveness, driving its integration into decision-making processes (Sherman et al., 2016).

RWE in Regulatory Approvals

Regulatory agencies like the U.S. Food and Drug Administration (FDA) and the European Medicines Agency (EMA) leverage RWE to make informed decisions throughout the lifecycle of medical products. For example, RWE is extensively used in **post-marketing surveillance** to monitor the long-term safety and effectiveness of approved drugs. By analyzing adverse event reports, electronic health records, and patient registries, regulators can detect rare adverse reactions and take timely action, such as updating warning labels or issuing safety alerts (FDA, 2021).

RWE also supports **label expansions** for existing drugs. For instance, the FDA approved palbociclib for certain breast cancer indications based on RWE demonstrating its effectiveness in broader patient populations not included in initial RCTs (Corrigan-Curay et al., 2018). This approach accelerates access to effective treatments while ensuring regulatory oversight.

RWE in Payer Decision-Making

Payers, including insurance companies and government healthcare programs, utilize RWE to evaluate the **value and cost-effectiveness** of medical interventions. Outcome-based reimbursement models rely heavily on RWE to

link payments to measurable improvements in patient outcomes. For instance, in outcome-based contracts, pharmaceutical companies agree to refund payers if a drug fails to deliver the expected clinical benefits, as evidenced by RWD (Sullivan et al., 2017).

RWE in Health Technology Assessments

Health technology assessment (HTA) agencies, such as the National Institute for Health and Care Excellence (NICE) in the UK, use RWE to supplement RCT findings during their evaluations. RWE provides a broader perspective on real-world effectiveness, adherence, and patient-reported outcomes, ensuring that recommendations reflect practical, everyday healthcare scenarios (Makady et al., 2017). This evidence strengthens the credibility of HTA decisions and fosters the adoption of innovative therapies.

5. Measuring Clinical and Economic Outcomes with Real-World Data (RWD)

Real-world data (RWD) provides a comprehensive lens for evaluating both clinical and economic outcomes in healthcare. Unlike randomized controlled trials (RCTs), which operate in controlled settings, RWD reflects the complexities of routine clinical practice, capturing diverse patient populations, treatment pathways, and healthcare resource utilization. This makes RWD an essential tool for assessing the real-world impact of healthcare interventions on patient outcomes and economic efficiency (Sherman et al., 2016).

Clinical Outcome Measurement

RWD is instrumental in evaluating the effectiveness and safety of interventions in real-world settings. Clinical outcomes measured using RWD include improvements in disease symptoms, prevention of disease progression, reduction in hospitalizations, and quality of life improvements. For example, data from electronic health records (EHRs) can track disease remission rates in patients receiving biologic therapies for rheumatoid arthritis, providing insights into the therapy's long-term effectiveness. Similarly, patient-reported outcomes collected through digital health tools can quantify changes in pain levels, mobility, or mental health, which are crucial for evaluating treatment success (Wang et al., 2021).

Adverse drug reactions (ADRs) are another critical clinical outcome assessed through RWD. By analyzing claims databases and pharmacovigilance systems, researchers can identify safety signals associated with specific drugs. This proactive approach enables healthcare providers to mitigate risks and

regulators to update drug safety profiles, thereby enhancing patient care and safety (Bate et al., 2018).

Economic Outcome Measurement

RWD is equally valuable in measuring economic outcomes, particularly in the context of value-based healthcare. Economic outcomes typically assessed include healthcare resource utilization (e.g., hospital admissions, emergency department visits), medication costs, and overall treatment costs. For instance, claims data can reveal the impact of a new drug on reducing hospital readmissions, thereby demonstrating its cost-effectiveness. Budget impact analyses, often conducted using RWD, help healthcare systems understand the financial implications of adopting new therapies or interventions (Sullivan et al., 2017).

RWD also supports cost-effectiveness analyses (CEA) by linking clinical outcomes with economic metrics. For example, the effectiveness of anticoagulants in preventing strokes can be translated into cost savings by analyzing the reduction in stroke-related hospitalizations. These insights are critical for decision-makers in determining the allocation of limited healthcare resources and prioritizing high-value interventions (Garrison et al., 2017).

Integration of Clinical and Economic Outcomes

RWD enables the integration of clinical and economic outcomes into a unified framework, providing a holistic view of a treatment's value. For instance, evaluating both the survival benefits and cost savings of a new oncology drug can help payers and policymakers make informed decisions about its inclusion in formularies. This approach is central to health technology assessments (HTAs), where RWD complements clinical trial data to assess the real-world value of interventions (Makady et al., 2017).

6. Data Integration and Interoperability in Healthcare Analytics

Data integration and interoperability are pivotal in healthcare analytics, enabling seamless access to and use of data across diverse systems and stakeholders. With the growing reliance on real-world data (RWD) for clinical, operational, and financial decision-making, the ability to unify data from disparate sources is essential. By breaking down silos and ensuring interoperability, healthcare organizations can derive comprehensive insights that drive better patient outcomes, enhance operational efficiency, and support value-based care models (Benson & Grieve, 2016).

Challenges in Healthcare Data Integration

Healthcare data often exists in fragmented forms, such as electronic health records (EHRs), laboratory systems, imaging systems, pharmacy records, and claims data. Each system may use different formats, terminologies, and standards, making integration complex. For instance, one hospital may record a patient's diagnosis using ICD-10 codes, while another uses SNOMED CT, leading to inconsistencies that complicate data aggregation and analysis (Kulikowski et al., 2021). Additionally, legacy systems with limited interoperability capabilities pose a significant challenge, as they lack the infrastructure to share or integrate data effectively.

Standards for Interoperability

Adopting standard data formats and protocols is critical for enabling interoperability in healthcare analytics. Commonly used standards include:

- 1.**HL7 (Health Level 7):** Facilitates the exchange of clinical and administrative data.
- 2.**FHIR (Fast Healthcare Interoperability Resources):** A modern standard developed by HL7 to improve the exchange of healthcare information using RESTful APIs.
- 3.**DICOM (Digital Imaging and Communications in Medicine):** Ensures the integration of medical imaging systems.
- 4.**LOINC (Logical Observation Identifiers Names and Codes):** Standardizes laboratory and clinical observation data.

These standards provide a common framework for sharing data across systems and organizations, reducing errors and improving the reliability of analytics (Bender & Sartipi, 2013).

Role of Data Integration in Healthcare Analytics

Data integration combines information from multiple sources into a unified repository, such as a data warehouse or data lake, enabling comprehensive analysis. This integration is crucial for:

- **Population Health Management:** Combining EHRs, claims data, and social determinants of health allows providers to identify high-risk populations and design targeted interventions.

- **Clinical Decision Support:** Integrated data provides clinicians with a holistic view of a patient's health, supporting evidence-based decision-making.
- **Operational Efficiency:** Aggregating data from scheduling, billing, and resource management systems helps optimize hospital operations and reduce costs.

Advances in Interoperability: APIs and Cloud Platforms

The rise of cloud computing and application programming interfaces (APIs) has revolutionized data integration and interoperability in healthcare. Cloud-based platforms enable real-time data sharing and storage, while APIs facilitate communication between disparate systems. For example, using FHIR-based APIs, a hospital's EHR can seamlessly exchange data with a wearable device, providing clinicians with up-to-date patient metrics such as heart rate or glucose levels (Shah et al., 2020).

Security and Privacy Considerations

Interoperability efforts must address data security and patient privacy concerns. Ensuring compliance with regulations like HIPAA (Health Insurance Portability and Accountability Act) in the U.S. and GDPR (General Data Protection Regulation) in Europe is critical. Data encryption, access controls, and audit trails are essential components of secure interoperability solutions (Raghupathi&Raghupathi, 2014).

7. Ethical and Privacy Concerns in Real-World Data (RWD) Analytics

The rise of real-world data (RWD) analytics has transformed healthcare by enabling evidence-based decision-making, improving patient outcomes, and reducing costs. However, the use of RWD also raises critical ethical and privacy concerns. These concerns revolve around ensuring the responsible collection, analysis, and use of patient data while protecting individual rights and maintaining public trust. Addressing these challenges is essential for leveraging RWD without compromising ethical standards or data privacy (Rumbold&Pierscione, 2017).

Patient Consent and Data Ownership

One of the primary ethical concerns in RWD analytics is obtaining informed consent from patients. Many RWD sources, such as electronic health records (EHRs), claims databases, and wearable devices, collect data passively,

often without explicit patient awareness or consent. This raises questions about the extent to which patients should have control over their data and whether they should be compensated for its use. Respecting patient autonomy requires transparent data collection practices and clear communication about how data will be used (Kim et al., 2021).

Data Privacy and Security

Privacy breaches are a significant concern in RWD analytics, given the sensitive nature of healthcare data. Even de-identified or anonymized data carries risks of re-identification when combined with other datasets. High-profile data breaches in healthcare have highlighted vulnerabilities in data storage and sharing systems, undermining public confidence. Implementing robust data security measures, such as encryption, access controls, and regular audits, is critical to preventing unauthorized access and maintaining privacy (Marr, 2020).

Bias and Equity in RWD Analytics

Ethical concerns also arise from biases in RWD, which can perpetuate health inequities. RWD often reflects systemic disparities in healthcare access, treatment, and outcomes, leading to biased analytics and decision-making. For example, underrepresented populations may not be adequately represented in RWD sources, resulting in algorithms that fail to address their specific needs. Addressing this issue requires proactive efforts to ensure data diversity and equity in RWD analytics (Rajkomar et al., 2018).

Balancing Public Benefit and Individual Rights

RWD analytics offers significant potential for public health benefits, such as improving disease surveillance, accelerating drug development, and optimizing healthcare delivery. However, the pursuit of these benefits must be balanced against individual rights to privacy and autonomy. Ethical frameworks should prioritize patient welfare while ensuring that data is used responsibly for societal gain. Initiatives like dynamic consent, where patients can modify their data-sharing preferences over time, offer a promising solution (Grady et al., 2017).

8. Outcome-Based Healthcare in Public Health and Population Health Management

Outcome-based healthcare is a paradigm shift that focuses on achieving measurable improvements in health outcomes rather than the volume of services provided. This approach has significant implications for public health

and population health management, emphasizing patient-centered care, value-driven resource allocation, and accountability. By leveraging real-world data (RWD) and advanced analytics, outcome-based healthcare seeks to address the needs of populations while promoting equity and sustainability in healthcare systems (Porter, 2010).

Enhancing Public Health through Outcome-Based Models

Public health initiatives aim to improve community health by addressing social factors, reducing diseases, and promoting prevention. Outcome-based healthcare supports these efforts by setting measurable goals like higher vaccination rates, fewer disease cases, or longer life expectancy. For example, diabetes prevention campaigns measure success through fewer new diagnoses and better HbA1c levels in the population (Koh et al., 2018).

Real-world data (RWD) is vital for tracking public health outcomes. Data from electronic health records (EHRs), claims, and wearable devices provide real-time insights into population health trends. This helps public health authorities allocate resources effectively and take timely actions, such as managing outbreaks or preventing chronic diseases (Hsu et al., 2020).

Population Health Management and Risk Stratification

Population health management focuses on improving health outcomes for specific groups, such as individuals with chronic conditions or those at high risk of adverse health events. Outcome-based models in population health rely on stratifying populations based on risk profiles derived from RWD. For example, predictive analytics can identify patients at high risk of hospital readmissions, enabling targeted interventions such as medication management or post-discharge follow-ups (Fisher et al., 2018).

Promoting Health Equity in Outcome-Based Healthcare

One of the core principles of outcome-based healthcare is ensuring equitable access to care and addressing disparities in health outcomes. Population health initiatives supported by RWD can identify gaps in healthcare access and outcomes among underserved communities. For example, geographic mapping of healthcare utilization data may reveal regions with limited access to preventive services, prompting targeted outreach efforts (Artiga & Hinton, 2018).

Applications in Chronic Disease Management

Chronic diseases such as diabetes, cardiovascular disease, and asthma impose significant burdens on healthcare systems. Outcome-based healthcare models in population health management focus on improving outcomes for patients with chronic conditions while reducing costs. Metrics such as reductions in hospitalizations, improvements in medication adherence, and patient-reported quality-of-life scores are commonly used to measure success (Berwick et al., 2008).

9.Challenges in RWD Analytics

1. Data Quality and Completeness

Ensuring data quality and completeness is one of the biggest challenges in RWD analytics. Data is often collected from various sources, like electronic health records (EHRs) and claims databases, which may use different formats and have inconsistencies. Missing or inaccurate data can affect the reliability of analysis, making data cleaning and standardization critical steps.

2. Data Integration and Interoperability

Data from different systems often exist in silos, making it difficult to integrate. Lack of common standards for data exchange, such as HL7 FHIR, creates barriers to interoperability. This fragmentation limits the ability to analyze healthcare data comprehensively.

3. Ethical and Privacy Concerns

Handling sensitive patient information raises ethical and privacy issues. Even anonymized data carries risks of re-identification when combined with other datasets. Complying with privacy laws like GDPR and HIPAA is crucial to protecting patient data and balancing its use with individual rights.

4. Bias and Generalizability

RWD often reflects biases in healthcare systems, such as disparities in care access. This can lead to skewed analytics and inequitable decisions. Furthermore, data from specific populations may not represent broader groups, limiting the generalizability of findings.

5. Analytical Complexity

The complexity and volume of RWD require advanced tools and expertise to analyze. Managing unstructured data, such as clinical notes or imaging, is challenging. Transparency and interpretability of models are also essential for trust among stakeholders.

10. Future Directions in RWD Analytics

1. Advanced Analytics and AI Integration

AI and machine learning will play a significant role in analyzing large datasets and uncovering patterns. These technologies can improve predictions and enable personalized medicine. Explainable AI (XAI) will help make complex models more transparent and trustworthy.

2. Blockchain for Secure Data Sharing

Blockchain can address data security and interoperability issues by providing a decentralized and tamper-proof platform for sharing data. Smart contracts can automate agreements, improving collaboration while maintaining privacy.

3. Standardization and Global Collaboration

Efforts to standardize data formats, like HL7 FHIR, and global collaborations, such as the OHDSI initiative, will help address interoperability challenges. These initiatives will enable seamless data sharing and analysis across organizations.

4. Dynamic Consent Models

Dynamic consent frameworks allow patients to control how their data is used, fostering trust and ethical use of RWD. Patients can update their consent preferences, making data-sharing more flexible and inclusive.

5. Real-Time Analytics and Decision Support

Real-time data processing will provide immediate insights for clinical decision-making. Integration with clinical decision support systems (CDSS) and wearable devices will further enhance data collection and analytics for better outcomes.

Conclusion

Real-world data (RWD) analytics is transforming the healthcare landscape by providing actionable insights into patient outcomes, treatment effectiveness, and system efficiency. Despite its immense potential, challenges such as data quality, interoperability, privacy concerns, and analytical complexity must be addressed to ensure that RWD analytics delivers its intended benefits. Establishing robust data governance frameworks, adopting standardized formats, and employing advanced analytics techniques such as artificial intelligence (AI) and machine learning are crucial steps toward overcoming these barriers. Furthermore, integrating ethical considerations, such as dynamic consent and equitable data representation, will build public trust and promote the responsible use of RWD.

The future of RWD analytics lies in fostering global collaboration, leveraging emerging technologies like blockchain for secure data sharing, and embracing real-time analytics to support clinical decision-making. As healthcare systems increasingly adopt outcome-based models, RWD analytics will play a central role in shaping evidence-based policies, improving patient care, and reducing healthcare disparities. By addressing the current challenges and capitalizing on future opportunities, RWD analytics has the potential to revolutionize healthcare delivery and create a more sustainable, equitable, and patient-centered system.


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Assessment of compost maturity and seed germination index

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Abstract

In this experiment seeds of three different plant species were used to assess the compost maturity with an objective to find an alternative to the conventionally used cress seeds (*Lepidium sativum*). Two immature food waste composts and one mature MSW compost was used to validate the claims. The garden cress is commonly used for SGI test. Among the seeds tested cucumber (*Raphanus sativus*) has the potential to be used as an alternative seed in place of garden cress to conduct SGI test.

Introduction

Plant growth and environmental pollution cleanup are two major uses for compost. Assessing the compost's quality is crucial, and a potent method for determining its toxicity is the seed germination test. Although the test is now frequently used, its development and application are limited by the fact that test results vary depending on the methodologies and seed species used. The challenge can be solved in part by standardizing procedures and modeling seeds. Additionally, the inaccuracy brought on by the methods of analysis and judgment for interpreting the data can be minimized, according to the probabilistic theory of seed germination.

Composting is an excellent method for managing solid waste since it converts organic solid wastes biologically into stabilized organic matter (OM). In the present era, research on many indicators associated with the composting process has grown in importance and scope. As a result, assessing the different stability and maturity indices of the composting process are of great importance and focused. By adopting suitable stability and maturity indices, the termination of composting process can be decided which bears significant monetary benefits.

Mature compost is defined by its high porosity, low moisture content, and plenty of microorganisms. It is a stable aerobic degradation product that is produced during the composting of solid waste. In order to boost free porosity, control moisture content, and inoculate microorganisms, mature compost has therefore always been utilized as a conditioning synergist in composting feedstock (Iqbal et al., 2010). Mature compost has been demonstrated in studies to enhance maturity. According to Yang and Zhang (2022), adding mature compost to green waste composting raised the final compost's total nitrogen content by 31.3% and seed germination index (GI) by 45%.

Approximately one-third (1.3 billion tons) of the food produced worldwide is lost to trash each year. Globally, this led to severe issues with the economy, society, and ecology (Browne and Murphy, 2013). It is imperative to turn food waste into compost because of the detrimental effects that such massive output of food waste has on the ecosystem. Globally, managing organic solid waste is a significant concern that, if handled improperly, may harm human health and the environment. The process of turning organic waste into compost stable substance that resembles humus involves controlled aerobic biological decomposition. The process is fundamentally the same as spontaneous decomposition, but in order to maximize microbial growth, organic waste is mixed with various chemicals to speed up and improve the process.

The potential benefits of composting manure and other organic waste include improved manure handling. Composting is easily adapted to agricultural operations because farmers generally produce suitable amounts and types of waste for composition, have adequate land, will benefit from the application of compost to the soil, and have the necessary equipment already available.

Fresh food waste has a high moisture content and little physical structure; therefore, it is critical to combine it with a bulking agent to create structure and absorb part of the excess moisture. A high C:N ratio bulking material, such as sawdust or dried leaves is to be mixed with the fresh food waste to adjust its physical environment suitable for composting. Food waste is quite vulnerable to the formation of odors, particularly ammonia, and huge amounts of leachate. Any woodworking process produces a large amount of sawdust. Sawdust may be recycled and composted with ease as an alternative to being thrown in the garbage. In addition to giving you a great soil conditioner, this will help the environment. Since sawdust is a dark, high-carbon composting material, it must be combined with "green," high-nitrogen elements.

Materials and methods:

Food waste and treatment design

Food waste was gathered from the Male dorm at Annamalai University. It had dhal, vegetables and cooked rice. The raw materials size had been decreased to 0.5 cm³. The ratio of food waste to sawdust was kept constant at 1:1 on dry weight basis. The moisture content of the food waste and sawdust mix was between 55% and 60%. A 20-liter homemade composting reactor was used to compost approximately 9 kg of food waste and 3½ kg of sawdust mixture for each mixing ratio. The aim of this study is to find alternative seed for *Lepidum sativum* to use in seed germination index analysis as a marker to analyze compost maturity. To obtain different stage of composting mass, the food waste mixed with sawdust was composted. The samples were carefully mixed after being weighed using a balance and added to the reactor. After completely mixing and sprinkling the samples with tap water, the containers were securely closed. The samples were well mixed to allow for enough aeration, which in turn promoted the growth of microorganisms. When necessary, water was sprayed onto the samples to maintain their moisture content.

Temperature of the composting mass is a clear indication of the microbial activity which indirectly indicates the active state of the composting mass. The temperature was monitored everyday using thermometer inserted in to the middle of the composting reactor. On day 7 and 18 the samples were collected from the composting reactor and the extracts of these samples were used in seed germination test.

Collection of seeds

Seeds of cress was obtained from a local supermarket in Nagercoil while the seeds of radish and cucumber were brought from Vegetable seed store. The picture of the seeds used in the study are presented in Figure 1.

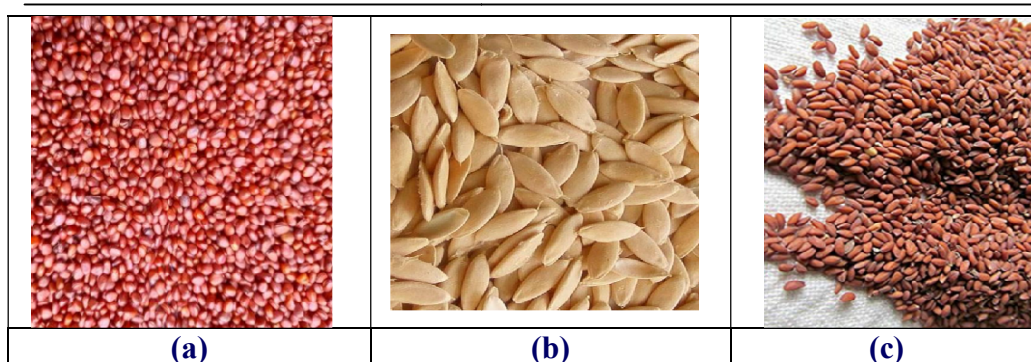


Figure 1. Seeds used in the study: (a) Garden cress seed (*Lepidium sativum*), (b) cucumber (*Cucumis sativus*) and (c) radish (*Raphanus sativus*).

Seed Germination Index

The compost obtained from specific days was used to analyze the seed germination index. About 20 g of compost sample was placed in a conical flask and 100 ml of distilled water added, 1:5 ratio on wet weight basis, and the flask was placed in an orbital shaker. After 30 minutes of shaking, the material was centrifuged for 10 minutes at 3000 rpm and filtered subsequently using Whatman No. 1 filter paper to obtain the supernatant. For each species, ten seeds of uniform size were placed in a glass petri dish and 7 ml of extract was added to the petridish. These Petri dishes were then incubated for 72 hours (3 days) at 25 °C in the dark. For every sample, three replicates were examined.

After 72 hours, the number of seeds germinated was counted, and the length of the radicle produced by each seed was measured to calculate the SGI. The seed germination index (SGI) was calculated using the following formula:

$$SGI (\%) = \frac{\left(\frac{\text{Number of seeds germinated in compost samples}}{\text{Number of seeds germinated in water control}} \times \frac{\text{Root length of seedling in compost samples}}{\text{Root length of seedling in water control}} \right)}{1} \times 100$$

Result and discussion:

Physiochemical properties of compost:

The food waste mixed with sawdust was composted only for a period of 18 days and two immature samples as indicated by a high temperature was

collected for analysis. The temperature profile of the composting mass is presented in Figure 2. A temperature of more than 45°C was observed for a period of 18 days indicating that the microbial activity was high and the compost cannot be a mature one. Temperature is an important indicator of composting it affects microorganism activity, organic matter degradation and composting efficiency (Kong et al.,2022).

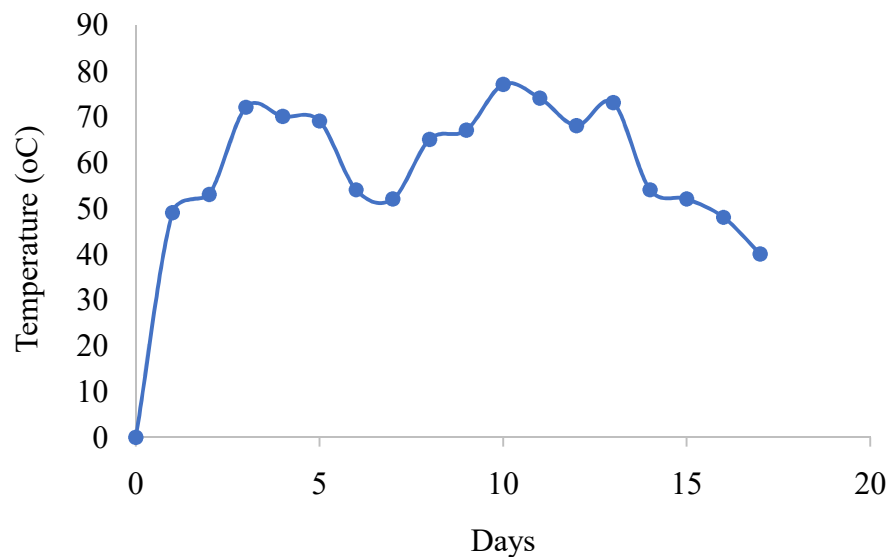


Figure 2. Changes in temperature of food waste: sawdust composting mass

Due to the time constraint, the composting experiment was not continued until maturity, i.e., the temperature of the composting mass reaches the ambient temperature. Therefore, a matured compost was obtained from a Micro-composting center (MCC) in Gandhi Nager, Tirunelveli for the analysis.

Seed Germination Index Result

The SGI results obtained for the three samples of different age are presented in Table 1 while the relevant figures are presented in Figures 3 -6.

Table 1. Seed germination index values obtained for different age of the composting mass.

Plant Species	Sample age	SGI (Mean±SD)
<i>Lepidium sativum</i>	Mature Compost	7.41 ± 9.33
	18 th Day	35.47 ± 55.87
	7 th Day	43.19±11.89
<i>Raphanus sativus</i>	Mature compost	279.45±147.33
	18 th Day	29.48±33.29
	7 th Day	38.76±15.06
<i>Cucumis sativus</i>	Mature compost	84.05±13.56
	18 th Day	98.12±17.81
	7 th Day	7.43±0.51

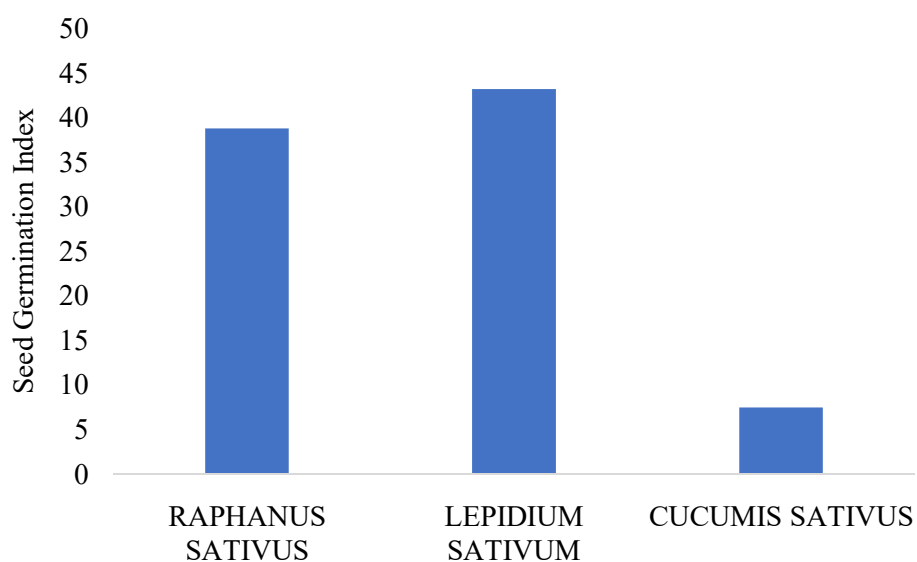


Figure 3. SGI values obtained for Day 7 food waste compost samples.

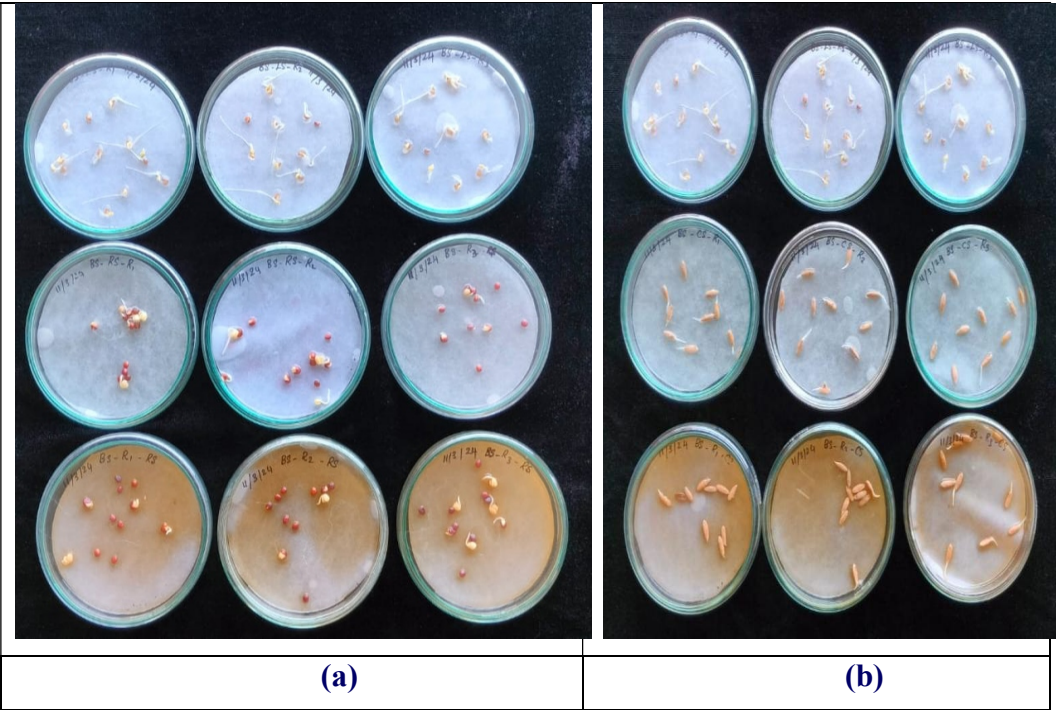


Figure 4. SGI test for Day 7 samples. a) *Lepidium sativum* and *Raphanus sativus*; b) *Lepidium sativum* and *Cucumis sativus*.

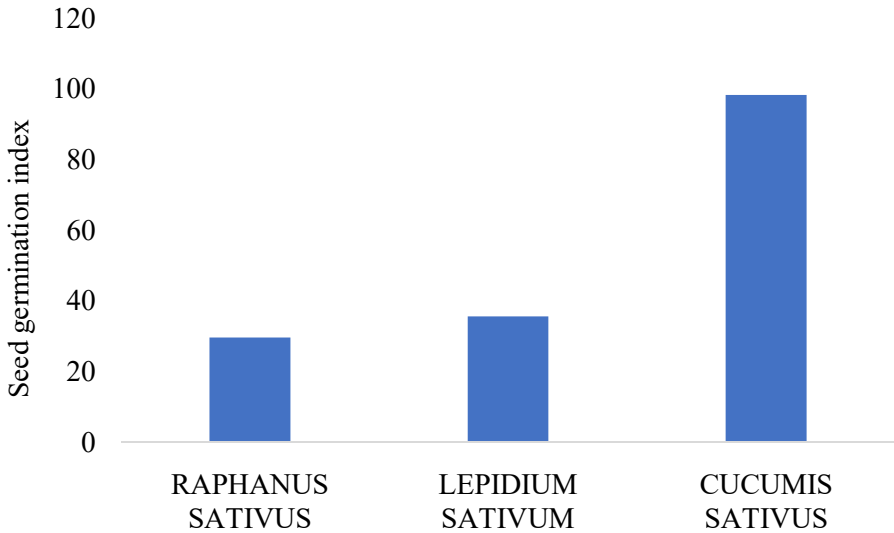


Figure 5. SGI values obtained for Day 18 food waste compost samples.

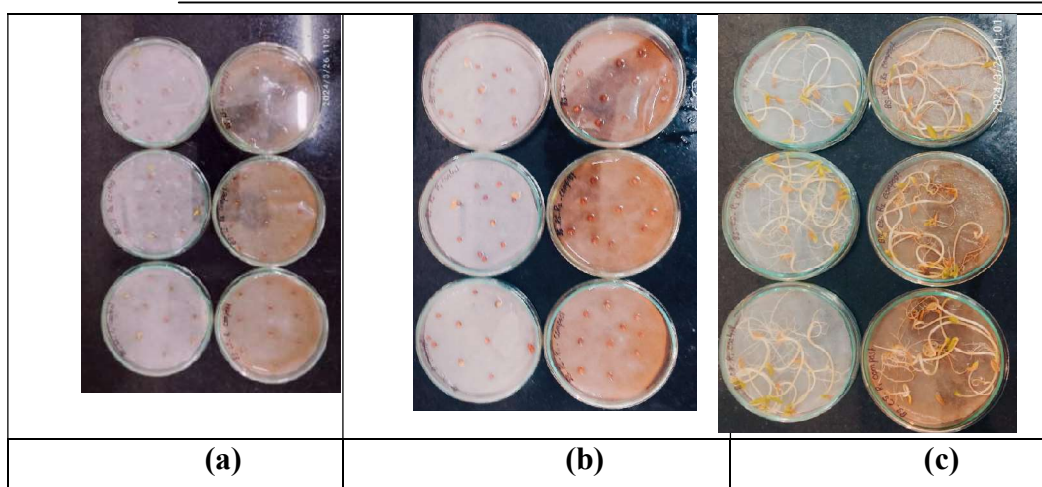


Figure 6. SGI test for Day 18 samples. a) *Lepidium sativum*, b) *Raphanus sativus*; c) *Cucumis sativus*.

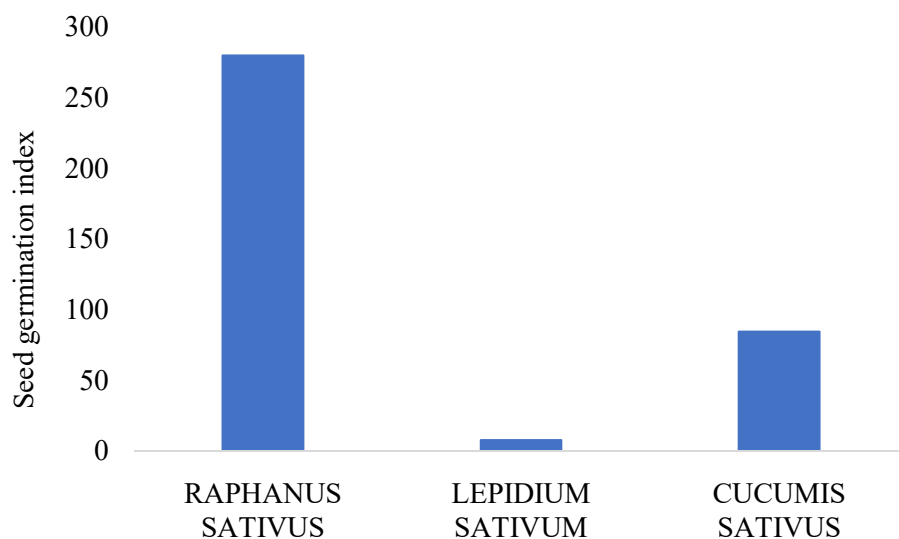


Figure 7. SGI values obtained for Day 18 municipal solid waste compost sample collected from MCC.

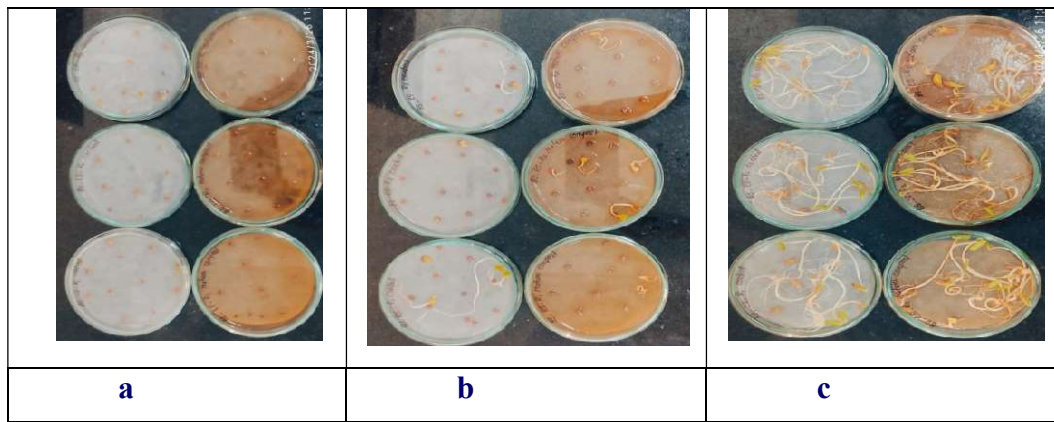


Figure 8. SGI test for mature MSW sample. a) *Lepidium sativum*, b) *Raphanus sativus*; c) *Cucumis sativus*.

Experiments involved the comparison of three plant species used to determine compost maturity/phytotoxicity. In the experiment two types of compost is used one mature MSW compost and two immature compost FW compost using 3 different types of seeds: cress (*Lepidium sativum*) radish (*Raphanus sativus*) and cucumber (*Cucumis sativus*).

The mature compost and 18th day cucumber has greater than 100% and 7 day all seeds are lower than 100%. GI 33.99% for *Lepidium sativum*, 126.9% for *Raphanus sativus* and 63.08% for *Cucumis sativus*. An SGI value of >80% indicates maturity. Results indicate that the germination index increased above 100% in the radish seed was indication of the elimination of phytotoxicity beyond that day and used to the control of compost.

GI value recorded formature compost indicates sufficient maturity. In the overall observation GI radish and cucumber suited for assessing compost maturity and better correlation compared to the cress seed.

Mostly cress seed is used to test the maturity of the compost. In the present study not only cress seed is used in control the test result but also radish and cucumber seeds were used.

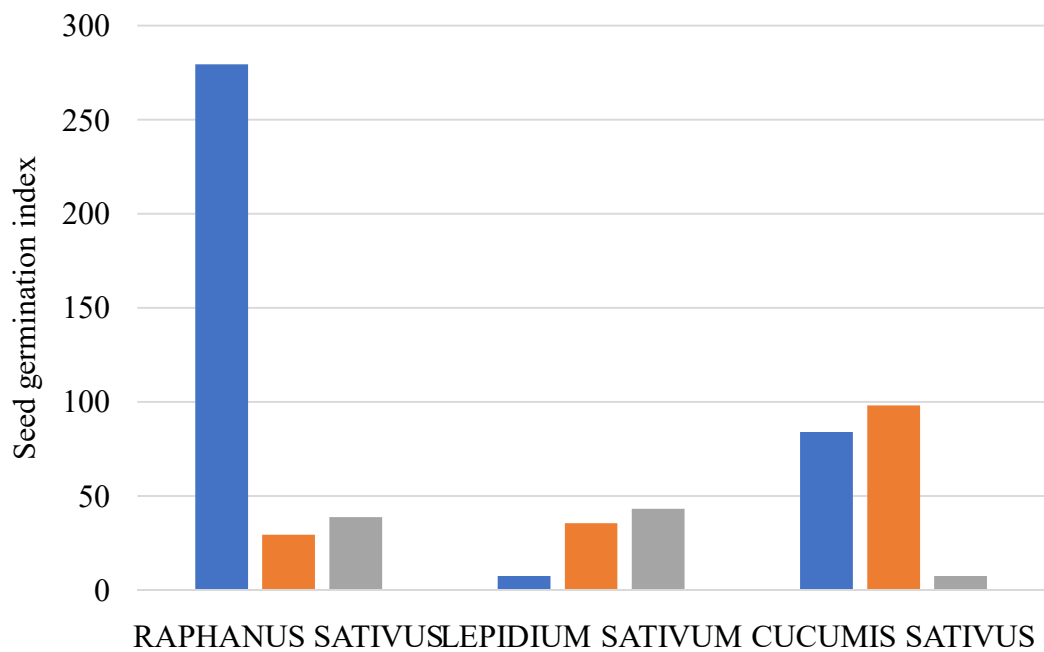


Figure 9. Overall observation of Seed Germination Index in mature and immature compost.

The blue colour is mature compost Raphanus sativus, orange colour is day 18 compost, 25 grey colour is day 7 compost extract. As the same as blue colour is mature compost Lepidium sativum orange colour is day 18 compost and grey colour is day 7 compost extract. Blue colour is mature compost, Cucumis sativus orange colour is day 18 compost and grey colour is day 7 compost extract.

Conclusion

Seeds of different species show distinct sensitivities to the biological toxicity of compost. The suitability of seeds of three species to assess the compost maturity was screened using the seed germination test. Then a toxicity experiment revealed that the GI of radish was suitable for evaluation of compost maturity and better correlation with characteristics of compost compared to the cress seed so the vegetable radish has the potential to be used as a seed for Seed germination index in place of cress seeds.

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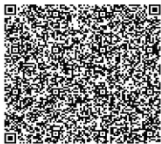
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Palmyra Tree -An Source of Nutritious Food

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Abstract

Originating from the tropical areas of India and Southeast Asia, the palmyra tree (*Borassus flabellifer*) exemplifies the intricate relationship between humans and their environment. This species is well-adapted to survive in challenging conditions, including drought and saline soils, due to its towering height, sturdy trunk, and distinctive fan-shaped foliage. Beyond its impressive physical characteristics, the palmyra tree serves as a vital source of nutrition, offering various edible components such as fruit pulp, sap, seeds, sprouts, and leaves, all rich in essential vitamins, minerals, fiber, and protein. These nutritional elements underscore the tree's significance as a fundamental food source for local populations, contributing to improved digestive health, enhanced immune function, and overall well-being. The Palmyra tree is useful for more than just food. Its sap can be turned into palm sugar, jaggery, and toddy. The leaves and fibers are made into mats, ropes, and baskets. These uses help the local economy and support sustainable practices in rural communities. Additionally, the tree has medicinal properties that are important in traditional healing methods like Ayurveda. Different parts of the tree are used to treat various health problems, including digestive issues and skin conditions.

The Palmyra tree holds profound cultural significance, representing themes of resilience and endurance. It is intricately integrated into local traditions and ceremonies, thereby strengthening the community's bond with its environment. The diverse uses of the tree underscore its vital role in sustaining ecological balance and bolstering local economies, which highlights the importance of conservation and sustainable practices. In essence, the Palmyra tree transcends its classification as a simple plant; it embodies a living testament to the complex interplay between humanity and nature. Its contributions to nutrition, medicine, economy, and culture position it as a crucial component of sustainable living, urging us to protect and honor such natural resources. By recognizing the multifaceted value of the Palmyra tree, we emphasize the urgent need to preserve biodiversity and cultivate a harmonious relationship with our surrounding.

Key words: Palmyra tree, Nutritional benefits, Medicinal properties, Sustainable practices

Introduction

The Palmyra palm, known scientifically as *Borassus flabellifer*, is a tall species indigenous to the tropical regions of India and Southeast Asia. Its striking presence in the landscape is characterized by its considerable height, robust trunk, and fan-shaped leaves. The tree's resilience and ecological significance are evident in its capacity to thrive in challenging environmental conditions, including drought and saline soils. Furthermore, the Palmyra palm plays a crucial role in the diets of local communities, as its various edible parts offer a wide array of nutrients, flavors, and textures, thus serving as a vital nutritional resource. Beyond its role as a food source, the Palmyra tree symbolizes a profound connection between humans and their environment, encapsulating elements of cultural heritage, culinary practices, and ecological stewardship.

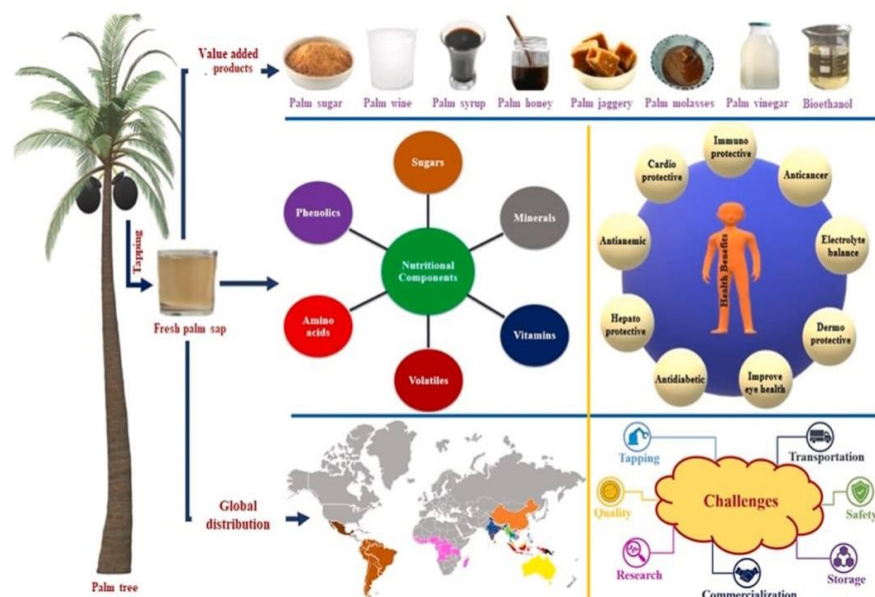


Fig no:01 Palmyra tree benefits
(Source:Chayanika Sarma et al 2022)

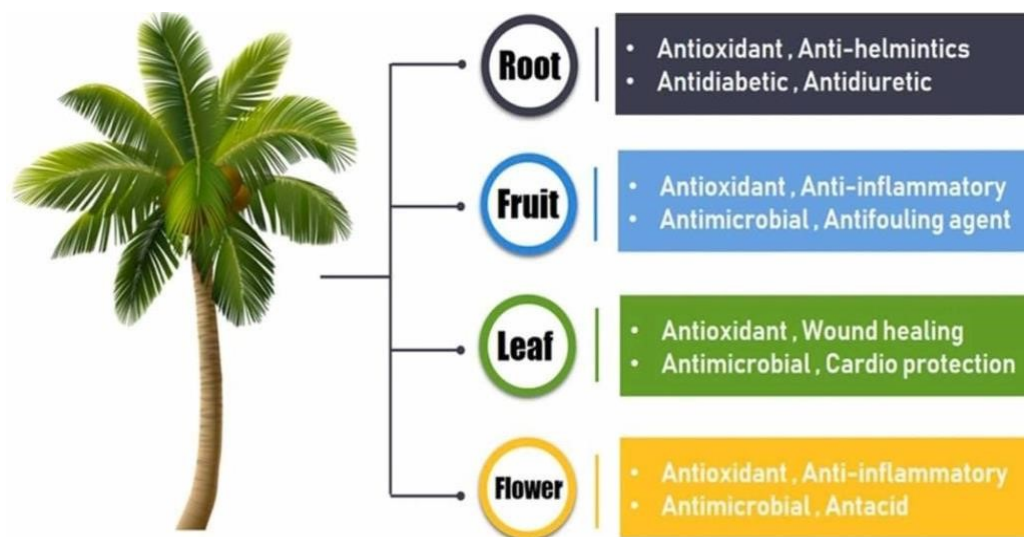


Fig No: 2 Nutritional highlights of palmyra tree
(Source:A.R. Basava Prasad et al 2023) **Vitamins and Minerals**

The Palmyra tree (*Borassus flabellifer*) serves as a significant natural source of essential vitamins and minerals, contributing notably to the enhancement of overall health and wellbeing. This tree is particularly abundant in vitamin C, which is vital for strengthening immune responses, protecting against oxidative stress, and aiding in collagen synthesis necessary for maintaining healthy skin and facilitating tissue repair. Furthermore, the vitamin B complex found in Palmyra products supports various metabolic processes, energy production, and the maintenance of optimal neurological health.

Regarding mineral content, Palmyra products are particularly rich in iron, an element critical for red blood cell production and for reducing the likelihood of anemia. Potassium, another vital mineral, is essential for fluid balance regulation, nerve function support, and the maintenance of healthy blood pressure. Additionally, calcium plays a crucial role in improving bone density, supporting dental health, and ensuring overall skeletal integrity. In summary, the vitamins and minerals present in Palmyra products constitute a valuable dietary component for enhancing immune function, metabolic efficiency, and cardiovascular health.

Protein content

Protein is an essential macronutrient vital for cellular repair, muscle development, and enzyme synthesis. Although Palmyra products are not typically known for their high protein content, they still provide a moderate yet valuable contribution to overall dietary protein levels. The seeds and tubers of the Palmyra plant are abundant in essential amino acids that support metabolic functions and encourage cellular regeneration.

Additionally, the protein derived from Palmyra products is noted for its excellent digestibility and bioavailability, making it a beneficial source of plant-based protein for individuals seeking alternative protein sources. Consequently, incorporating Palmyra into daily nutrition can support muscle preservation, improve tissue repair, and help maintain enzymatic balance.

Fiber Content

The palmyra tree is important because it has a lot of dietary fibre, which is key for good metabolism and digestive health. Products from the palmyra tree include both soluble and insoluble fibres. These fibres help the body absorb nutrients better, keep bowel movements regular, and ensure smooth digestion. Soluble fibre slows sugar absorption, helping to keep blood sugar

levels steady, while insoluble fibre prevents constipation and supports regularity.

Moreover, palmyra dietary fibre can lower cholesterol, reducing the risk of heart diseases. It also helps people feel full, which can prevent overeating and assist with weight control, making it a vital part of a healthy diet.

Palmyra tree as a source of food

Edible Components of the Palmyra Tree

The Palmyra tree presents a variety of edible components, each characterized by distinct flavors, textures, and nutritional advantages. The young shoots, collected from the top of the tree, provide a gentle crunch and a mild sweetness, whereas the juicy fruit pulp delivers a refreshing tartness that evokes a sense of tropical delight. Furthermore, the seeds, which are protected by fibrous coverings, represent a concentrated source of energy and vital nutrients, encapsulating the essence of natural health and sustenance.

Palmyra tree products

The Palmyra tree produces more than just raw produce; it also yields an abundance of handcrafted goods that are the epitome of creativity, tradition, and skill. Made from the tree's sap, palm sugar has an earthy scent and sweetness akin to caramel that enhances desserts, drinks, and savoury foods. The traditional sweetener palm jaggery, made from boiled sap, has a complex flavour profile and a sweetness that is rich in minerals. Its unique character enhances food preparations. Moreover, toddy or palm wine is highly esteemed in the community and is used as a social mixer and ceremonial libation in addition to having probiotics and antioxidants that may have health advantage.

Toddy

Fresh palm juice is clear, very sweet, and not yet fermented. The main ingredient in fresh toddy juice is sucrose, which makes up about 12-15% of its weight. It also has a small amount of reducing sugars. Natural yeasts in the sap start to ferment glucose, creating alcohol. Acetobacter helps by turning the alcohol into acetic acid, or vinegar. Because of the low vinegar content, it's best to drink fresh juice within a day of tapping; after that, it becomes more sour. The sap ferments quickly after collection due to the natural yeasts. The result is palm wine, which has about 4% alcohol. If the sap is left in the sun, it can turn into palm wine in about two hours. Toddy typically has an alcohol content of around 5%.

Toddy refers to the way wild yeasts and bacteria ferment the sugar sap. This process happens naturally without control. Many different yeast and bacterial strains are involved in fermentation. Neera has around 75 different LAB strains. In controlled lab settings, palm sap fermented with pure yeast cultures can reach an alcohol content of 7.8%. The sugars in fermented toddy mainly include sucrose, glucose, and fructose, which gradually convert to ethyl alcohol during fermentation.



Jaggery

Jaggery, also known as Gur, is a key ingredient in Indian cooking. Its sweet flavor, combined with earthy notes, makes it a delicious treat. While it is enjoyed worldwide, it holds a special place in India, where many desserts rely on Gur for sweetness. Jaggery is also important in various ancient Hindu ceremonies and in Ayurveda, the traditional Indian medicine. Both sugarcane and palm jaggery provide quick energy when eaten. Compared to regular sugar, jaggery is a healthier choice.



Nungu

Nungu, commonly referred to as palm fruit, possesses anti-inflammatory and antioxidant properties, primarily attributed to its high concentration of crude flavonoids. The fruit's pulp is known to aid in the healing of skin irritations and is utilized for alleviating symptoms of nausea, vomiting, and parasitic infections. Furthermore, it serves as an expectorant and contributes positively to liver health. A thin application of sugar palm fruit jelly on affected skin can provide relief from the itchiness associated with

prickly heat. Rich in essential vitamins and minerals, sugar palm fruit is particularly advantageous for individuals adhering to specific dietary restrictions or managing diabetes, as it contains vitamins B and C, along with vital minerals such as iron, zinc, potassium, calcium, and phosphorus. During warmer months, the consumption of palm fruit aids in maintaining hydration and replenishing lost nutrients, thereby preventing discomfort such as painful urination and fatigue. It is also effective in addressing digestive concerns and can function as a laxative. Sugar palm fruit is beneficial for those on weight-loss regimens and plays a crucial role in preventing malnutrition in both children and adults. The Nungu season typically spans from May to August. The fruit develops in clusters, measuring approximately 4 to 7 inches in diameter, encased in a black shell. Inside, it contains three sweet, jelly-like seed pockets enveloped by a thin, yellowishbrown skin. The white, fleshy interior is filled with a watery fluid, while the fibrous outer layer can be consumed raw, boiled, or roasted, offering a jelly-like texture.



Palmyra sprouts

Palmyra palms, scientifically known as *Borassus flabellifer*, are the plants on which palmyra sprouts, also known as palmyra tubers, develop. A crunchy kernel that tastes like a sweeter water chestnut is revealed when the hard shell of the germinated seed is opened. These sprouts are delicious and have a lot of health advantages. Only during specific seasons are they accessible. Palmyra sprouts reduce body temperature and help avoid constipation. They contain a lot of fiber, which might help you lose weight. They can also improve immunity and ease stomach problems. Palmyra sprouts are good for women's health and bone strength. Omega-3 fatty acids, which can help lower elevated cholesterol, are abundant in them. In addition to strengthening the immune system, these sprouts may lower the risk of cancer and heart disease.

According to the analysis, the roots have a very low fat content, 8.54% protein, 23.53% carbs, and 7.29% crude fiber. These roots are high in calories. The fleshy roots and seedlings are both eaten, and they are an essential source of sustenance for persons with little money. In loose, sandy soil, 100–150 seeds are sown in 3–4 layers per 0.8 square meters, with the potential to produce at least 100–150 seedlings, and occasionally much more. The long, club-shaped, starchy portions of the seedlings, which are collected between two and four months of age, can be crushed into flour, baked, roasted, fried, or boiled. The seedlings are sun-dried and cooked to preserve them for subsequent use. When they are around four months old, they begin to devour the meaty roots. They are low in proteins and lipids and heavy in starch. The root is used to cure gonorrhea and is well-known for its cooling, restorative, diuretic, and anti-parasitic qualities.



Neera or Palm sap

Neera is the liquid collected from the flowers of the Palmyra palm. It is often referred to as sweet toddy because it has no alcohol content. The sap naturally ferments due to the yeast present in it, starting soon after collection. This fermentation can lead to a low alcohol level, typically under four percent, but the sap must be consumed within 24 hours for best quality. To prolong its freshness, the sap is first filtered to remove impurities. It is then clarified using a special microfiltration system and pasteurized by heating it to 90-95°C for 3-7 minutes. After this process, the sap is packaged and kept in a refrigerator, which can extend its shelf life to about 10 weeks. Common packaging materials for Neera include High-Density Polyethylene (HDPE) pouches, Polyethylene Terephthalate (PET) bottles, and glass bottles.

Palm candy

Neera, cleaned of impurities, is boiled in a metal pot with a small amount of superphosphate (Golly et al., 2017). Once it reaches a steady boiling point, the liquid is cooled down. After filtering out sediments, it is heated to 110°C for 2 hours until it becomes thick like honey (Ullah et al., 2018) or boiled at 180°C until it reaches 67 OB (Mani et al., 2018). The liquid is then

cooled and poured into a crystallizer. Sugar crystals begin to form after 45-60 days. The final product is taken out and rinsed with water. Lastly, it is sun-dried at a temperature of 40-80°C for about 2 hours.

Health benefits of palmyra product consumption

Boosting Immune System

The nutritional abundance of the Palmyra tree, which is rich in immune-stimulating vitamins and minerals, strengthens the body's defences against environmental stressors and pathogenic diseases. Specifically, vitamin C is essential for encouraging tissue regeneration and repair, strengthening resistance to respiratory infections and supporting immunological function.

Regulating Digestion

The diet high in fibre from the Palmyra tree supports metabolic balance and digestive health, which in turn promotes gastrointestinal regularity and nutrient absorption. Insoluble fibres provide stool more volume and relieve constipation by encouraging bowel regularity, while soluble fibres create a gel-like matrix in the digestive tract that slows down the absorption of carbohydrates and cholesterol. Furthermore, fibre provides a varied microbial habitat that is vital for immunological response and metabolic equilibrium by acting as a prebiotic substrate for good gut flora.

Providing Energy

The products of the Palmyra tree are a concentrated supply of carbohydrates that provide the body with long-lasting energy and vitality which makes them perfect fuel sources for physical activity and metabolic needs. Particularly palm jaggery and palm sugar provide a slow-release form of glucose that keeps blood sugar levels stable and energy levels steady all day. In addition, the tree's seeds which are high in fibre and healthy fats make a satisfying snack that promotes sustained endurance and lasting satiety.

Promotes Weight Management And Metabolism

Products made from palmyra are renowned for having a low glycaemic index and naturally lower sugar content than other tropical fruits, which makes them a great option for people who want to control their blood sugar levels. Blood glucose spikes are avoided because the natural sugars, such as sucrose, are absorbed gradually into the bloodstream. For people who already have diabetes or are at risk of getting it, this trait is very beneficial. Furthermore, Palmyra fruit's high fibre content encourages fullness, which lowers overeating and helps with weight management. By regulating blood sugar and cholesterol

levels, the tree's sap is also known to promote metabolic health and promote cardiovascular wellbeing in general.

Skin Health

The palmyra tree's components, especially its sap and leaves, are valued for their positive benefits on skin health. Because of its many antioxidants and anti-inflammatory qualities, the sap is commonly used as a tonic to preserve skin brightness and reduce the appearance of wrinkles. These antioxidants are essential for scavenging dangerous free radicals and minimising aging-related skin damage. Because of their antibacterial and therapeutic properties, the fruit and leaves are also used to treat eczema, acne, and other skin disorders.

The fruit provides moisture and sustenance, and the leaves, when applied as a paste, can relieve sensitive skin.

Health Of The Liver and Detoxification

Products made from palmyra, especially the sap, are known to have detoxifying qualities. By increasing urination, the sap acts as a natural diuretic, aiding in the removal of waste and extra pollutants. This process aids in the liver and kidneys' purification, increasing their capacity to filter out dangerous chemicals. Furthermore, by strengthening the body's detoxification mechanisms and protecting the liver from oxidative damage—a major cause of liver disorders and cirrhosis—Palmyra products are thought to support liver health.

Rejuvenation and Vitality

Palmyra sap is highly valued in traditional medicine due to its refreshing and energizing attributes. In Ayurveda, it is classified as a Rasayana, or rejuvenating tonic, believed to enhance vitality, energy, and lifespan. This sap is commonly utilized to alleviate fatigue, boost mental clarity, and restore physical strength. Its nourishing and revitalizing effects make it especially beneficial for individuals recovering from illness or experiencing prolonged stress. Furthermore, it is thought to possess adaptogenic properties that assist the body in coping with both mental and physical stressors.

Stress Reduction and Mental Health

Stress reduction and improved mental health are linked to using items made from the palmyra tree, especially the sap. The sap's mild sedative and relaxing properties aid in promoting mood, lowering anxiety, and relaxing the nervous system. Palmyra is regarded in Ayurveda as a natural stress reliever

because of its cooling qualities, which calm the body and mind and reduce tension, irritation, and mental exhaustion. It is also thought that the relaxing properties of palmyra tree roots can help with insomnia and promote better sleep.

Blood Sugar and Cholesterol Control

The fruit and sap of the Palmyra tree are highly effective in managing blood sugar levels, making them advantageous for individuals with diabetes or those aiming to prevent the condition. Packed with dietary fiber, the fruit slows sugar absorption into the bloodstream, minimizing the chances of sudden spikes in blood sugar. Furthermore, Palmyra-derived products have a low glycemic index, making them ideal for diabetic-friendly diets. In addition to these benefits, Palmyra may aid in lowering cholesterol levels, promoting cardiovascular health. Regular inclusion of Palmyra products in one's diet can improve metabolic function and reduce the risk of heart disease.

Palmyra tree in traditional medicine

Healing Properties

Embedded within the annals of traditional medicine, the Palmyra tree assumes the role of a medicinal marvel, revered for its therapeutic properties and healing virtues. Its fruits, leaves, sap, and roots harbour a wealth of bioactive compounds, ranging from antioxidants and phytochemicals to antimicrobial agents and anti-inflammatory substance each contributing to its multifaceted pharmacological profile.

Medicinal Uses

The Palmyra tree's medicinal prowess finds expression in a myriad of traditional remedies and formulations aimed at alleviating a spectrum of ailments and afflictions. From respiratory disorders and digestive complaints to dermatological conditions and metabolic disorders its botanical bounty serves as a natural pharmacy offering respite and rejuvenation to those in need.

Ayurvedic Applications

In the ancient tradition of Ayurveda, the Palmyra tree occupies a revered position as a botanical ally in the quest for holistic wellness and vitality. Endowed with a spectrum of therapeutic virtues, it finds resonance in Ayurvedic formulations designed to balance doshas, pacify ailments, and restore harmony to body, mind, and spirit. Whether employed as a rejuvenating tonic, a digestive aid, or a skin tonic, the Palmyra tree's botanical bounty embodies the essence of Ayurvedic wisdom and natural healing.

Conclusion

The Palmyra tree symbolizes the deep connection between humanity and the natural environment, serving as a remarkable source of nutrition, culinary experiences, and medicinal benefits. Its impressive stature and botanical richness evoke a sense of wonder and reverence, highlighting nature's boundless generosity and resilience. Known scientifically as *Borassus flabellifer*, the Palmyra tree exemplifies the complex interplay between people and their surroundings. Its towering form is not only visually striking but also essential for communities that rely on its diverse products for their sustenance and economic well-being. The tree's nutrient-dense sap can be processed into sweet syrup or fermented into drinks, while its fibrous trunk and leaves are utilized in construction and crafts, making it integral to the ecological and economic landscape of its habitat.

Moreover, the therapeutic properties of the Palmyra tree enhance its significance. In traditional medicine, various components of the tree are utilized for their health benefits, including digestive support and treatment for skin ailments. This multifaceted utility underscores the importance of preserving species that offer both nutritional and medicinal value, reinforcing our responsibility to protect these vital natural assets.


In addition to its practical uses, the Palmyra tree holds substantial cultural significance in many communities. It is often celebrated in local traditions and rituals, symbolizing strength and endurance. This cultural relevance fosters a deep connection between people and the environment, reminding us of our role as stewards of nature. Given the urgent global challenges such as climate change and biodiversity loss, embracing the lessons imparted by trees like the Palmyra can inspire sustainable practices that honor both our cultural legacy and the well-being of future generations.

In summary, viewing the Palmyra tree as more than just a resource, but rather as a vital component of our ecosystem, encourages a holistic perspective on conservation efforts. Recognizing its multifaceted roles—ranging from nutritional and economic to medicinal and cultural—enhances our commitment to nurturing a balanced relationship with the natural world. The Palmyra tree is not simply a plant; it serves as a profound symbol of the interdependence of all living beings.

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Microbial Ecology: Microorganism In Natural Environment

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Microorganism in natural environment

In the Natural Environment microorganism play crucial roles in maintaining ecology balance and contributing to essential processes. Microbial ecology in natural environment refers to the interaction between microorganisms bacteria, fungi, and viruses are crucial functioning of natural environment. These organisms are involved in key ecological processes that sustain life on earth.

Key Aspects of Microbial Ecology in Natural Environment

1. Microbial Diversity

The diversity of Microbial species in different ecosystems is vast, and their functions contribute significantly to ecosystems processes like soil fertility, plant growth, and water purification.

2. Nutrient Cycling

Microorganisms help in decomposing organic matter and recycling nutrients like carbon, nitrogen, sulfur, and phosphorus which are essential for life.

3. Adaptation to Environment Condition

Microbes are highly adaptable and can thrive in diverse environment, from extreme heat to cold, acidic to alkaline and low oxygen to oxygen rich condition.

4. Symbiotic Relationship

Microbes engage in various symbiotic relationships, including Mutualism, Commensalism, and Parasitism, with Plants, Animals, and other microorganisms.

5. Ecological Niches

Microbes occupy specific niches within ecosystems, including soil, water, air, and the human body, where they contribute to the overall health and balance of the earth.

6. Microbes and Climate change

Examining the impact of Microbial activities on green house gas emissions (e.g., CH_4 , CO_2 , and N_2O).

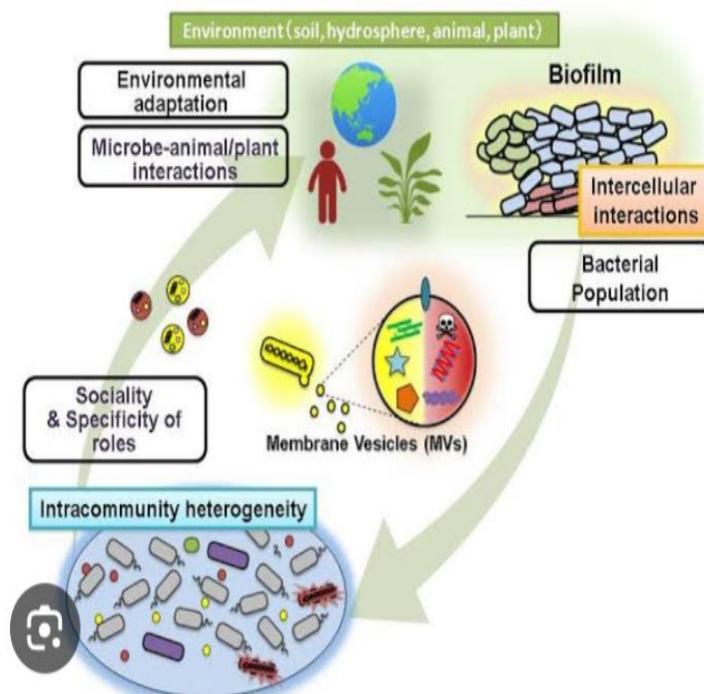


Figure 1: Microbial ecology and Role of microorganisms in ecosystems

Methods Used in Microbial Ecology Research

- Culture-independent techniques: since many microbes cannot be cultured in the lab, techniques like metagenomics, 16S rDNA sequencing, and amplicon sequencing, are used to analyse microbial communities in sites.
- Stable isotope probing (SIP): To track the metabolic activities of specific microbial groups.
- Microcosms and field experiments: To study microbial interactions and community responses to environment changes.
- Environment DNA (eDNA) Sampling: To capture genetic material from microbes in environment samples like soil, water, or air.

Microbial ecology in natural environment:

1. Taxonomic diversity

Microbial population in natural environment exhibit vast taxonomic diversity. The diversity can be classified into different functional groups like primary producers (photosynthetic microbial), decomposers and nitrogen fixers.

Different microbial species contribute to various functions within ecosystems, such as decomposing, nitrogen fixation, and sulfur cycling.

Extreme Environment

Microbes can thrive in extreme conditions where other life forms might not survive. These include.

- Thermophiles: Heat-loving microbes found in hot springs and hydrothermal vents.
- Halophiles: Salt-loving microorganism found in salt lakes or saline environment.
- Acidophiles and Alkalinephiles: Microbial that live in environment with extreme pH levels (acidic or basic).
- Psychrophiles: Cold-loving microbes found in polar regions and deep-sea environment.

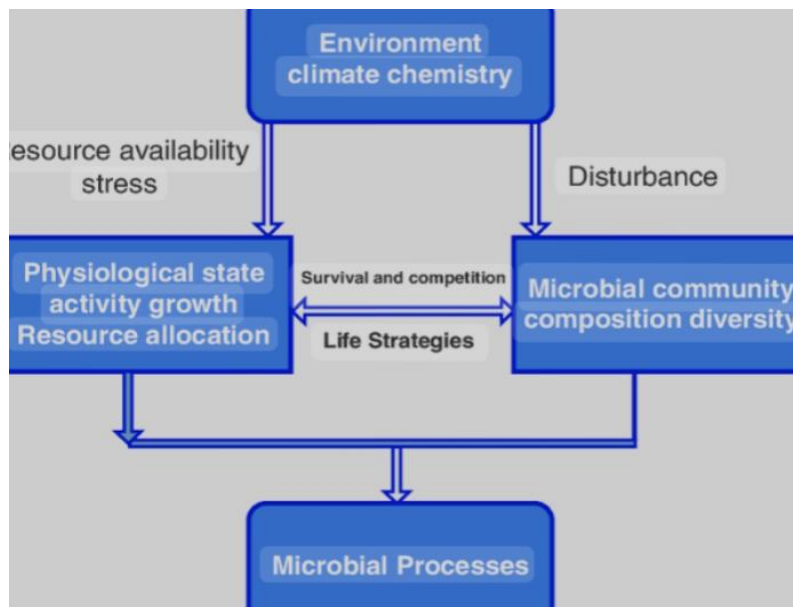


Figure 2: Microbial diversity

Types of Microorganisms in Natural Environment

Bacteria: Bacteria are among the most abundant and diseases microorganism in nature.

- Example: *Nitrosomonas*(nitrification), *Rhizobium*(nitrogen fixation), *Lactobacillus* (fermentation).

Archea: Although similar to bacteria, archea are genetically distinct and often thrive in extreme environment (extremophiles) ,Many archea are involved in methane production (methogens) or sulfur cycling.

- Example: *Sulfolobus*(acidic hot springs), *Methono bacterium*(methane production).

Fungi: Fungi include yeast, molds, and mushrooms.

- Example: *Aspergillus* (decomposition), *Pencillium* (antibiotic production), *Glomus*(mycorrhizal fungi).

Algae and Caynobacteria (Blue-Green algae): Algae including both macroalgae (e.g. sea weeds) and microalgae(e.g., diatoms, green algae) are key photosynthetic organisms in aquatic environment.

- Example: *Chorella*(freshwater), *Anabaena*(cayanobacteria), and *Porphyra*(red algae).

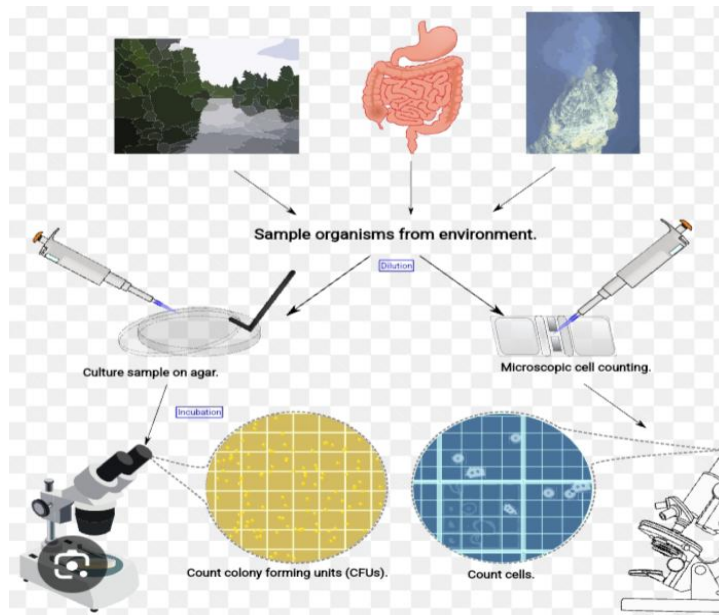


Figure 3: Microbial Ecology

Challenges in Microbial Ecology Research

- **Microbial Complexity:** Microbial communities are incredibly diverse, and analysing their interactions in complex, seat-world environment is difficult.
- **Environment Variability:** Natural Environment are highly dynamics, which can make it challenging to isolate specific microbial roles and behaviours.
- **Technological Limitations:** Despite advances in sequencing and bioinformatics, understanding the full functioning potential of Microbial communities is still a work in progress.

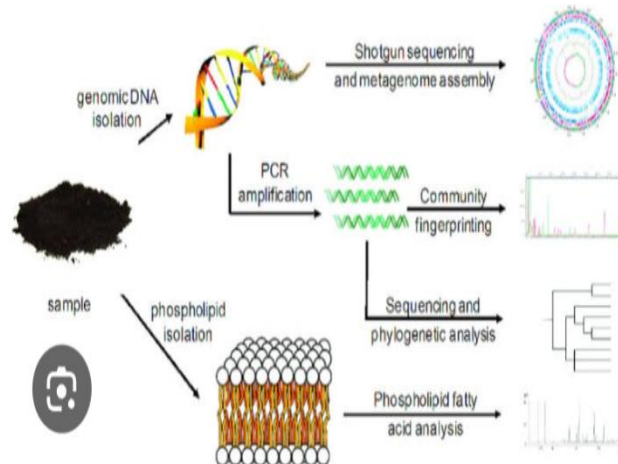


Figure 4: Microbial Ecology approaches used to examine both cultivatable.

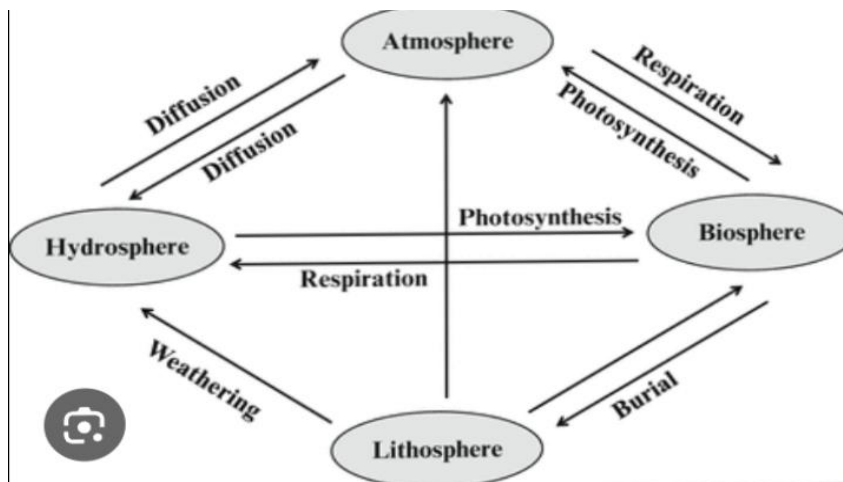


Figure 5: Microbes and environment.

Methods and Methodology

Studying microorganism in natural environment involves various methodologies and techniques designed to understand their diversity function, interacts and roles in ecological processes. The methods span from traditional cultivation based techniques to advanced molecular and genomics approaches. Below are key methodological and methods commonly used in Microbial researches in natural environment.

1. Culture-Based Methods

a) Isolation and cultivation

- Purpose: To isolate and identify individual microbial species from natural environment.
- Method: Sample (e.g. soil, water, air) are collected from the environment and plated onto selective media that factor the growth of specific microbial groups (bacteria, fungi, etc.) the microbes are then cultured and identified using various biochemical or morphological characteristics.
- Challenges: Many microorganism in natural environment are unculturable under standard laboratory conditions, leading to underrepresentation of Microbial diversity.

b) Enrichment Culture

- Purpose: To increase the concentration of a specific microorganism of interest by providing growth conditions favourable to its growth.
- Method: Samples are inoculated into selective media or incubated under specific environmental conditions (e.g. temperature, PH) that favors the growth of the desired microorganism. Enrichment Culture is particularly useful for isolating rare or slow-growing Microorganism.
- Challenges: It may not represent the full diversity of microorganisms present in the environment and the microbes might still not grow due to environmental constraints.

3. Molecular Techniques

c) DNA/RNA Extraction

- Purpose: To obtain genetic material from environment sample to analyse microbial diversity and activity.

- Method: DNA or RNA is extracted from environment sample (soil, water, etc.,) using specialized kits or protocols. The extracted DNA is then used for downstream application such as PCR amplification or metagenomics sequencing.
- Challenges: Environment samples often contain inhibitors that can interfere with DNA/RNA extraction and extraction method must be optimized for different sample type.

d) Polymerase Chain Reaction(PCR)

- Purpose: To amplify specific DNA region to identify microorganism or quantify specific gene.
- Method: PCR is used to amplify a target gene often the 16S rRNA gene (for bacteria) or ITS region (for fungi) which is used to identify microorganism present in a sample. Primers specific to these regions are desired and PCR amplification is performed.
- Challenges: PCR amplification can sometimes lead to biases, as certain Microorganism may not amplify efficiently. Additionally PCR is limited to detection of known sequences. For example, while among the moths, only caterpillars cause the damage.


Conclusion

In the context of research and applied fields microbial ecology helps in areas like medicine (understanding infection and diseases), Agricultural (enhancing soil fertility or controlling pests), and Environment Sciences (Bioremediation or climate change).

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Medical science and its foundations

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Introduction

Medical science is the field of study and practice concerned with maintenance health, diagnosis diseases, and treating illnesses in humans and animals.it is a multidisciplinary science that combines biology, chemistry, physics and social sciences to understand the mechanism of life and the factors affecting health and diseases .the foundations of medical science is built upon centuries of knowledge, discovery, and advancement that have shaped modern medicine leading to improved public health and life expectancy worldwide.

Historical Development

The roots of medical science date back to ancient civilizations,with early practice emerging from place like ancient Egypt , Mesopotamia, Greece,and India. Each of these societies contributed to the understanding of anatomy, surgery, pharmacology,and hygiene, which were rudimentary but foundations.

Ancient Egypt:

The Egyptian are credited with early practice in medicine, especially in the field of surgery and anatomy. Notable figures include Imhotep,who is often regarded as the first physician.

Ancient Greece:

The Greek physician Hippocrates (406-370BCE) is often considered the “Fatherof Medicine “. He introduced the idea that disease were caused by natural factors rather than divine punishment.theHippocratic oath, with forms the ethical foundations for medical practices,is still used in modified forms today.

Mediaeval and Renaissance periods :

Medical knowledge stagnated in Europe during the early middle Ages but was preserved and expanded upon by scholarly World, such as Avicenna, whose work “The canon of medical” was used in Europe for centuries. The Andreas vesalius providing.

Modern Era:

The 17th and 18th centuries saw the birth of modern scientific method, and with it, a rapid acceleration in medical knowledge. The development of microbiology, germ theory by Louis Pasteur and Robert Koch, and advance revolutionized medicine

2. Core Foundations of medical science

1. Biology and physiology:

Understanding how the body works is the foundation of medical science. physiology studies the normal function of the body and its system, including circulation, respiration, digestive, and nervous coordination

2. Anatomy:

Anatomy is the study of the structure of organisms and their parts. Medical practitioner need an intimate knowledge of human anatomy to perform surgeries, diagnosis diseases, and understanding how illness affect the body.

3. Bio chemistry:

The study of chemical processes within and relating to living organism is crucial to understand how diseases develop at the molecular level and how various treatment can alter biochemical pathways to restore health.

4. Pharmacology:

This area focuses on the study of drugs and their effect on the human body. knowledge of pharmacology is essential for prescribing medications and understanding their side effects, interaction, and efficacy.

5. Pathology:

Pathology is the study of diseases, including their cause, development, and effect on the body. it cover the study of abnormal tissue changes, including the identification of microorganisms, cancerous growth, and genetic disorders.

6. Microbiology :

This branch examines microorganisms that cause diseases. Medical microbiology aids in diagnosing infection and understanding how these pathogens invade and affect the human body.

7. Immunology:

Immunology is the study of the immune system and how the body defends itself against infections and affects the human body.

8. Epidemiology:

This is the study of disease patterns in a population. It is crucial for understanding how diseases spread, how they can be prevented, and how public health policies can be formulated to improve community health.

9. Medical Ethics and Legal Aspects:

Medical science is not only about the technical knowledge but also about how to apply that knowledge responsibly. Medical Ethics focuses on moral issues like patient autonomy, consent, confidentiality, and equity in healthcare.

The Modern Medical Approach

Modern medicine integrates advanced technologies and research both in diagnosis and treatment. Key components include:

Diagnostic Technology:

Medical imaging techniques (X-ray, MRI, CT scan, ultrasound) allow physicians to look inside the body and assess disease conditions without invasive procedures.

Genetics:

Advances in genetic research and molecular biology are paving the way for personalized medicine, where treatment can be tailored to the genetic makeup of an individual.

Surgical Innovation:

Minimally invasive surgery, robotics, and advanced anesthesia techniques have revolutionized patient care and recovery.

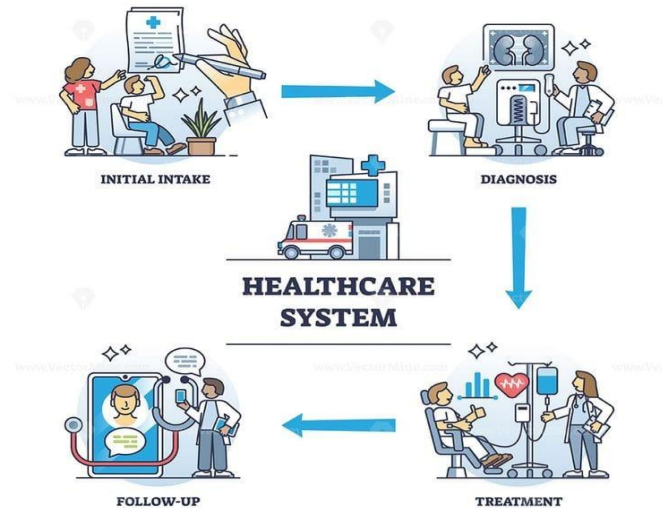
Pharmacogenomics:

This area study how an individual 's genetic makeup affects their respons to drugs,enabling more precise prescribing and fewer side effects.

Aspect	Description
Medical Science	A field of study focused on the diagnosis, treatment, prevented, and management of diseases and medical conditions
Core Principle	Based on understanding biological functions, diseases mechanism and treatment methods.
Foundations Discipline	1.Anatomy: Study of body structures. 2.physiilogy: study of functions. 3.Biochemistry: study of chemical processes in the body. 4.pathology: study of diseases and their causes.
Key Branches	1.pharmacology:Study of drugs and their effect. 2.Medical Technology:use of tools for diagnostic and treatment. 3.Epidemiology: study of diseases patterns in populations.
Medical Ethics	Focus on principle like patient autonomy, benefices and justice in medical practices.
Clinical medicine	Direct care for patients including various specialist such as surgery,pediatrics,and obstetrics.
Research and Innovation	Ongoing clinical and laboratory research driving evidence-based practices and technological advancements.

Emerging Research Concept in Life Sciences

Historical Foundations	Ancient contribution from Greek Roman, and Islamic scholars; significant advance during the Renaissance.
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Pathology

Path

-pertaining to a morbid process (diseases)
*(From “pathos”=travail or sorrow (diseases)

ology

-the scientific of...

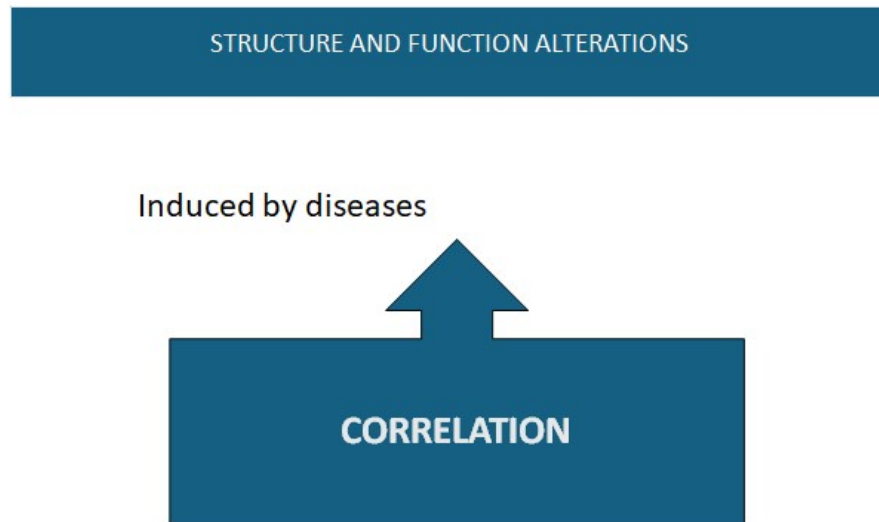
Pathology Objectives:

1.To introduced a NEW VOCABULARY of terms, definition, and diseases processes

2.To introduced the BASIC RESPONSE of the human organisms to injury

- Correlation with clinical pictures.

Pathology objective (2).



From the basis by which attending

Signs and symptoms are produced

Levels of study of pathology

- Molecular
- Chemical
- Ultrastructural
- Cellular
- Tissue
- Organ

Differential diagnosis

Vascular -sudden onset

Inflammatory -cardinal sings /symptoms

Neoplastic -MA

Drugs -history

- Infection -cardinal sings and symptoms
- Congenital -From Birth
- Autoimmune -systemic
- Traumatic -history

- Endocrine/metabolic -sn/sx.

Pathomorphology:

PATHO...the diseases

MORPH...The shape (structure)

LOGY...The study of

i.e.how a diseases process alters the “shape”(structure)of cells, tissues and organs.....or.....

The “Damage”caused by the diseases process.

2.The Role of Technology in medicine:

Medical technology has drastically transformed healthcare, enabling more accurate diagnoses, effective treatment, and better patients outcomes.here are some key examples:

Imaging Technologies :

MRI,CT scan, X-ray,and ultrasound allow non -inasive observation of the body's internal structure,helping diagnose condition such as fracture,tumors,and internal bleeding

Genetic Technologies:

Advance in genomics, like DNA sequencing,have led to personalized medicine, where treatment are tailored to an individual genetic makeup, improving efficiency and reducing side effects.gene editing technologies,likeCRISPR , hold promise for treating genetic disorder.

Robotics and Minimally Invasive Surgery:

Robotics and enhanced surgical instruments allow for less invasive,more precise surgeries,often reducing recovery time and risk of infection.

Artificial Intelligence and machine learning:

AI is revolutionizing medicine by analysing large datasets for patterns and prediction diseases progression.Altools can also assist doctors in interpreting medical images or assessing patients data.

3.medical Research and Innovations:

Medical science is heavily driven by research that allows us to understand the mechanism behind diseases, discovery new treatment, and

develop innovative Medical technologies.there are two primary types of research.

Basic Research:

This research seeks to expand our understanding of fundamental biological o processes .for exaples , scientific studying cellular mechanisms of aging or immune response could lead to new therapeutic strategies.

Clinical Research:

In clinical trials,new drugs or treatment protocols are tested on human subjects to assess their safety and efficacy.these trials follow a rigorous, ethical framework to ensure patients safety.its through clinical research that we gain approval for medication vaccines,and surgical techniques.


Conclusion

The foundation of medical science rests on centuries of accumulated knowledge and continuous advancement in multiple fields.by integrating biology,technology,and Innovation, modern medical science strives to improve health outcomes and enhance quality of life.the future of medical science holds great promise,with on going research likely to uncover new

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Crop Production: Techniques and Trends

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Introduction

Crop production denotes the fundamental feature of agriculture, focused on the cultivation of plants for food, fiber, medicinal purposes and raw materials. Crop production depends on the accessibility of arable land and is affected by yields, macroeconomic uncertainty, as well as consumption patterns; It also has a great occurrence on agricultural commodities prices. Crop production becomes more complex with a proliferation of available cultivars, greater pest pressures and governmental regulations, fewer agricultural chemicals to control pests, more emphasis on appearance of harvested crops, more competition from domestic and foreign sources, and lower prices for many crops. The significance of crop production is related to harvested areas, produced. As the global population continue to grow and challenges like climate challenges, resource limitations, and environmental concern intensify, the require for improved crop production techniques, trends and technologies has become more pressing.

Techniques involved in crop production :

Crop production involves a variety of techniques aimed at increasing yield, quality and sustainability. These techniques vary depending on the type of crop, climate conditions and available resources. There are several steps involved in techniques of crop production.

1.Land preparation :

Land preparation is a crucial first step in crop production that directly impacts crop establishment, growth, and overall yield. Land preparation covers tillage or minimum the damage.



Figure 1:land preparation

Tillage which reduce soil disturbance through to a totally 'puddled' soil which actually destroys soil structure. The purpose of land preparation is to provide the necessary soil conditions which will enhance the successful formation of the young offshoots or the tissue culture plants received from the nursery. It can be done of using manual labor, machinery (e. g., bulldozers), or herbicides depending on the scale of the operation.

2.selection of seeds :

Healthy and good quality seed are the roots of a healthy crop. The seeds that are used to cultivate new crops have to be selected very carefully and high level quality. The selected seed should not be infected. Farmers should also check the germination period of the selected seed, nutrients required for the growth, and other benefits which can result in a good yield.

3.Sowing and planting :

Sowing and planting are critical step in crop production, determining the foundation for healthy crop development and optimal yields. After the preparation of soil the previously selected seeds are scattered in the field. Traditionally, sowing is done manually by hand and in some places, seed drilling machine are used. The process involve selecting the right techniques, timing and conditions to ensure successful crop establishment. There are different types of sowing such as hand sowing, seed drill, open -field planting and pre-treatment of seed and soil before sowing.

4.Irrigation management :

Irrigation management is the important method and techniques used to apply water to crops in areas where rainfall is insufficient or inconsistent. Proper irrigation management is critical for optimizing crop yield, conserving water, preserving soil health. With climate change, droughts, and increasing

competition for water resources, effective irrigation management helps mitigate these challenges, ensuring food security and enhancing agricultural productivity.

Methods of Irrigation :

Irrigation can be carried out in two methods.

1.Traditional methods -In this method, irrigation is done manually. Here, a farmer pulls out water from wells or canals by himself or using cattle and carries to farming fields. This method can vary in different regions.

2.Modern methods -The modern method compensates the disadvantages of traditional methods and thus helps in the proper way of water usage.

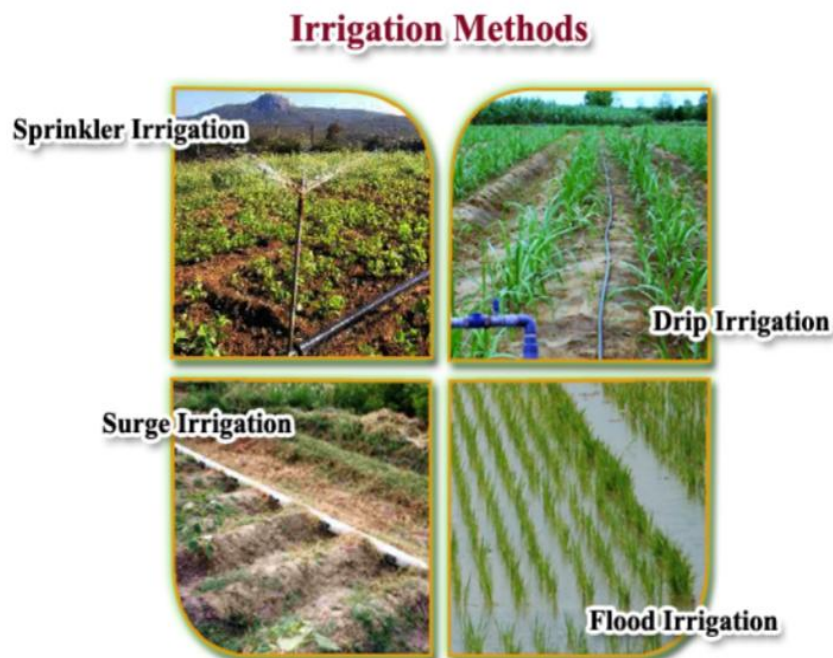


Figure 2 :Methods of irrigation

5.Fertilization management :

Fertilization management in crop production is a practice that ensures crops receive the appropriate nutrients in the right amounts, at the right time, and in the right form. These nutrients are crucial for various physiological processes like photosynthesis, root development, and fruiting. In this method the goal is to ensure there are sufficient levels of nutrients based on crop

requirements, peak harvest times, and the soil's natural ability to provide nutrients. Proper nutrient supply helps crops grow to their full potential. Fertilizers improve the water holding capacity of the plants and increase root depth. Proper fertilizer management involves the correct selection, application, and timing of fertilizers to achieve maximum efficiency and minimize environmental impact.

6.Crop protection :

Crop protection is the general method or the practice of protecting crop yields from different agents including pests, weeds, plant diseases, and other organisms that cause damage to agricultural crops. It encompasses a wide range of techniques, products, and technologies, including

1.Chemical control :

Herbicides : Substances that kill or inhibit growth of unwanted plants(weeds).

Insecticide : Substances that kill arthropod pests, i. e. insects and mites.

Fungicides : Substances that destroy or prevent the growth of pathogenic fungi.

2.Biological control :

The use of natural predators, parasites, or pathogens to control pests and disease. Biological control is particularly useful where chemical pesticides are not suitable or are impractical in environmentally sensitive areas, or on low-unit-value crops.

3.Cultural practices:

Crop rotation, intercropping, and selecting resistant crop varieties are strategies that help manage pest and disease outbreaks and improve soil health.

4.Physical control :

Physical control is the practice of mechanical or hand controls where the pest is actually attacked and destroyed. These controls are used mostly in weed control. Methods such as mulching, netting, or traps to physically prevent pests from reaching crops or to reduce pest population.

7.Monitoring crop growth :

Crop monitoring in agriculture is transforming how farmers manage their fields, offering real-time insights into crop health and growth. The growth of crops is influenced by various factors such as weather, soil conditions, pests,

disease, and management practices. Crop growth monitoring involves the systematic observation and measurement of various parameters related to plant development including size, health, biomass, and environmental conditions.

In today's agricultural landscape, a wide array of tools and technologies, from visual inspection to advanced satellite imaging and remote sensing, is used to monitor and analyse crop growth.

8. Harvesting :

Harvesting of crops involves the process of gathering mature crops from the fields. It is a critical step in agriculture that ensures the crops are collected at their peak quality and stored properly to prevent spoilage and loss. The process can be done by either harvesting machines or by skilled workers. The timing and method of harvesting play a vital role in determining the quality, quantity, and marketability of the final product. In today's times, harvesting machines such as combine harvesters are used. Successful harvesting not only involves collecting the crop but also incorporates post-harvest handling practices such as sorting, cleaning, drying, and storing, all of which play an essential role in preserving the product's quality and preventing losses.



Figure 3 : Harvesting

9. Processing and Marketing :

The processing and marketing of crops are the final step in crop production that take place after harvesting, contributing to the value of the produce and ensuring that it reaches the final consumer in the desired form.

✓ Processing – crop processing involves converting raw agricultural products into consumable or usable forms. It is designed for producers who

supply produce exclusively for further processing in the supply chain-from freezing, juicing, roasting , milling, precooking, and more to ingredients for animal feed.

✓ Marketing - Marketing involves the activities and processes through which processed or raw crops are made available to consumers or industrial buyers. Good marketing takes planning, selling discipline, access to good market information and a good understanding of pricing and delivery alternatives.

Trends in crop production :

Crop production trends have been shaped by a variety of factors over time, including technological advancements, climate change, changing consumer preferences, and shift in agricultural policies. As the global population continues to rise, the demand for food has increased, driving innovations in farming practices, crop varieties, and resource management. Emerging agricultural trends mark a shift towards skilled cultivating and productive use of time and assets while lessening crop misfortunes.

Some of the trends in crop production:

1.Artificial Intelligence (AI) :

AI in crop production encompasses a variety of functions, from precision farming to predictive analytics. Artificial Intelligence technologies, such as machine learning algorithms and data analytics, enable precision farming by analysing vast amount of data from sensors, satellites, and drones. Simulated intelligence offers prescient experience for gauging climate information, crop yield, and costs, subsequently helping farmers to pursue informed choices. AI – driven autonomous machines, such as tractors, harvesters, and planters, are increasingly used in agriculture. These robots can plant, water, monitor, and harvest crops with minimal human intervention. AI powered robotics improve efficiency, reduce labour costs, and minimize human errors in crop production.



Figure 4 : Artificial Intelligence in crop production

2.Precision Agriculture :

Precision agriculture is an innovative farming management concepts that uses technology and data – driven insights to optimize crop production, increase efficiency, and reduce environmental impacts. It is a technique where farmers utilize careful measures of information, like water, pesticides, and composts, to upgrade the quality and efficiency of yield.

Benefits of precision Agriculture :

- ✓ Increased yield : By optimizing resource use and minimizing waste, precision agriculture leads to higher crop yield and more efficient production.
- ✓ Resource Efficiency :With more precise application of water, fertilizer, and pesticides, farmers can reduce input costs and minimize environmental impact .
- ✓ Cost savings : By minimizing input waste and maximizing crop yields, precision agriculture helps farmers lower production costs.
- ✓ Environmental protection : By reducing the overuse of chemicals and fertilizers, precision agriculture minimizes the risk of water pollution and soil degradation.
- ✓ Sustainability : Precision farming practices contribute to sustainable agriculture by reducing over -farming, preserving soil health, and conserving water.

3.Regenerative Agriculture :

Regenerative Agriculture (RA) is “a synergistic approach on various agricultural systems that verily inculcates the techniques to satisfy the global

agricultural demands as well as ensures the environmental wellbeing of all the resources it uses rather than destroying or depleting them”. This approach emphasizes soil regeneration, carbon sequestration, and biodiversity. It incorporates farming practices that seeks to improve biodiversity by reducing carbon footprint .Techniques include no –till farming, cover cropping, and agroforestry, aiming to restore soil health and increase resilience to climate change.



Figure 5: Regenerative Agriculture

4.Genetically Modified(GM) and Hybrid Crops:

GM crops are organisms whose genetic material has been altered in a laboratory setting using techniques like recombinant DNA technology. This allows the insertion, deletion, or modification of specific genes to achieve desired traits, which may not be possible through traditional breeding methods.

Examples : Bt cotton (resistant to pests), Roundup ready soybeans (resistant to herbicides), Golden rice (enhanced with vitamin A), and drought – resistant corn.

Benefits :

- ✓ Increased resistance to pests and diseases.
- ✓ Improved crop yields.
- ✓ Reduced need for chemical pesticides and herbicides.
- ✓ Enhance nutrition content .

Hybrid crops are typically produced by cross – pollinating two different varieties or species of plants. The resulting offspring (F1 generation)

exhibit hybrid vigor or heterosis, meaning they grow more vigorously, are larger, and often more productive than either parent.

Examples : Hybrid corn, hybrid tomatoes, hybrid wheat, and hybrid rice.

Benefits :

- ✓ Increased yield and productivity.
- ✓ Improved resistance to disease, pests, and environmental stress.
- ✓ Enhanced uniformity in crops.

Both GM and hybrid crops play vital roles in modern agriculture, enhancing food security by improving crop productivity and resilience. However, the choice between the two methods depends on factors like desired traits, environmental impact, regulatory consideration, and market demand.

5.Drones :

Drones equipped with cameras and sensors can fly over fields, capturing high – resolution images and videos. This allows farmers to monitor the overall health of crops, detect pest infestations, or identify areas with diseases, all in real – time. Drones have found diverse applications on farms of all scales, from field scouting to enhancing security measures. The data they collect isn't just numbers; it's the backbone of 'precision agriculture,' guiding farmers towards informed decisions that can boost yields by up to 5%, a game-changer in an industry known for tight profit margins.



Figure 6 : Application of drones in crop production

Expanding ranch efficiency while saving expenses is testing. In any case, drones, otherwise called automated airborne vehicles, assist farmers with

conquering this issue in a successful manner. Drones gather crude information which converts into valuable data for ranch observing. Drones outfitted with cameras work with ethereal imaging and studying of all over extended fields.

Table 1 : Production of important crops in three largest producing states in 2023-2024

Crops/Groups of Crops	States	Production	Per cent Share of Production to All India	Cumulative per cent Share of Production
(1)	(2)	(3)	(4)	(5)
I. Foodgrains				
Rice	Telangana	16.63	12.17	12.17
	Uttar Pradesh	15.72	11.50	23.67
	West Bengal	15.12	11.06	34.73
Wheat	Uttar Pradesh	35.43	31.38	31.38
	Madhya Pradesh	21.28	18.84	50.22
	Punjab	17.78	15.75	65.97
Maize	Karnataka	5.49	15.39	15.39
	Bihar	4.61	12.93	28.32
	Madhya Pradesh	4.33	12.14	40.46
Total Nutri/ Coarse Cereals	Rajasthan	8.03	14.66	14.66
	Karnataka	7.61	13.90	28.56
	Madhya Pradesh	5.49	10.02	38.58
Tur	Karnataka	1.02	30.13	30.13
	Maharashtra	0.86	25.41	55.54
	Uttar Pradesh	0.38	11.23	66.77
Gram	Madhya Pradesh	3.19	27.52	27.52
	Maharashtra	2.86	24.72	52.24
	Rajasthan	2.23	19.31	71.55
Total Pulses	Madhya Pradesh	6.18	25.23	25.23
	Maharashtra	4.00	16.33	41.56
	Rajasthan	3.63	14.83	56.39
Total Foodgrains	Uttar Pradesh	59.29	18.03	18.03
	Madhya Pradesh	39.84	12.12	30.15
	Punjab	32.59	9.91	40.06
II. Oilseeds				
Groundnut	Gujarat	4.64	45.12	45.12
	Rajasthan	2.02	19.59	64.71
	Madhya Pradesh	0.99	9.63	74.34
Rapeseed & Mustard	Rajasthan	5.98	45.40	45.40
	Uttar Pradesh	1.87	14.24	59.64
	Madhya Pradesh	1.75	13.28	72.92
Soyabean	Madhya Pradesh	5.47	41.92	41.92
	Maharashtra	5.23	40.09	82.01
	Rajasthan	1.17	8.96	90.97
Sunflower	Karnataka	0.07	40.72	40.72
	Haryana	0.03	17.37	58.09
	Odisha	0.02	13.77	71.86
Total Oilseeds	Rajasthan	9.57	24.17	24.17
	Madhya Pradesh	8.37	21.15	45.32
	Gujarat	7.19	18.15	63.47
Sugarcane	Uttar Pradesh	205.56	46.45	46.45
	Maharashtra	112.09	25.33	71.78
	Karnataka	41.81	9.45	81.23
Cotton@	Gujarat	9.06	27.86	27.86
	Maharashtra	8.05	24.74	52.60
	Telangana	5.08	15.62	68.22
Jute & Mesta\$	West Bengal	7.87	80.97	80.97
	Bihar	0.99	10.15	91.12
	Assam	0.68	7.05	98.17


Conclusion

Crop production techniques are involving to meet the challenges of increasing food demand, climate change, and sustainability. Key trends include the use of precision agriculture technologies for efficiency, the adoption of sustainable farming practices to minimize environmental impact, the development of climate – resilient crops, and the rise of organic farming. Additionally data – driven approaches are enabling more informed decision to optimize yields. The future of crop production will rely on integrating innovation, sustainability, and technology to ensure food security and environmental health.

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Artificial Intelligence In Sustainable Agriculture

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Introduction

Artificial Intelligence (AI) referred to the capability of machines or computers to carry out tasks that typically needed human intelligence, including learning, problem-solving, and decision-making. In 1956 "Artificial intelligence" (AI) was termed by John McCarthy, who is a computer scientist at a workshop in Dartmouth College. Agriculture involves the cultivation of soil, the growing of crops, and the raising of animals for food, fiber, medicinal plants, and other essential products for human life. The term "agriculture" was introduced by the Roman scholar Marcus Terentius Varro in the 1st century BCE. The term agriculture was derived from the Latin words "ager" (field) and "cultura" (cultivation). Artificial Intelligence (AI) in agriculture uses the technology to improve farming by optimizing crop yields, monitoring soil health, detecting pests, and automating tasks like irrigation and harvesting. It uses machine learning, computer vision, and sensors to analyse data, increasing productivity, reducing waste, and promoting sustainability. Examples include AI-powered drones and autonomous machinery. This chapter reviews the applications, advantages, disadvantages and development of artificial intelligence in agriculture.

Applications of Artificial Intelligence In Agriculture

Artificial intelligence (AI) enhances the efficiency, productivity and sustainability of agriculture. Here are some main applications of Artificial intelligence (AI) in agriculture:

1. Precision farming:

Artificial intelligence (AI) helps farmers monitor the use of various resources like water, fertilizer, weather, temperature, and pesticides to optimize planting, irrigation, and harvesting. AI-powered tools, including drones and sensors, help farmers by providing real-time data."



Figure 1 : Precision farming using AI

2. Plant Monitoring :

Traditional approaches to monitoring plant health and crop conditions are both labor-intensive and time-consuming. Implementing AI provides a more efficient way to detect and diagnose potential plant health issues or soil nutrient deficiencies. By leveraging deep learning, AI-driven applications are being created to analyze patterns in plant health, offering valuable insights into soil conditions, pest issues, and plant diseases. For example, Ramos-Giraldo *et al.*, in their study "Drought Stress Detection Using Low-Cost Computer Vision System and Machine Learning Techniques," developed an automated, low-cost drought detection system using computer vision and machine learning algorithms to document the drought response in corn and soybean field crop.

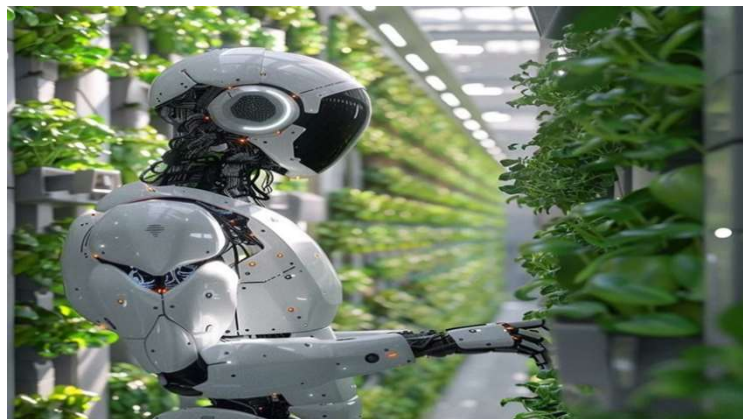


Figure 2 : plant monitoring by robots

3.Predictive analysis:

Predictive analytics and remote sensing are increasingly used agriculture to forecast crop production, yield, and harvest outcomes under different conditions. AI-driven data analytics helps farmers protect natural resources like land, air, and water while reducing input needs. Hatfield et al. developed tools using remote sensing, neural networks, and machine learning to identify variable areas in fields and devise adaptive strategies to improve profitability and reduce environmental impact through more efficient use of resources. AI is transforming agriculture, enabling more sustainable food production with fewer resources. Technologies such as the Internet of Things (IoT), Big Data Analytics, Cloud Computing, Mobile Computing, and AI support the development of smart agriculture. IoT connects intelligent devices through networks, using hardware (smart sensors) and software (AI algorithms) to monitor and control agricultural processes in real time. These technologies help with reasoning, tracing, management, and performance within the agricultural supply chain, contributing to improved agricultural practices (Channe *et al.*, 2016).



Figure 3 : predictive analytics in agriculture

4. Data Science :

Farms generate vast amounts of data daily, which can be analysed using AI technologies to inform better farming decisions regarding weather conditions, temperature, water usage, and soil quality. AI systems must be capable of filtering large volumes of data while remaining responsive to important events, requiring expert knowledge to ensure that only the most relevant data is processed, which improves both speed and accuracy. The development of agricultural expert systems necessitates collaboration among agricultural specialists and farmers. By merging AI and big data, data analytics in agriculture can lead to increased production and cost reductions, offering precise recommendations for practices like irrigation, fertilization, crop protection, and harvesting. AI technologies enable farmers to make informed decisions while using fewer natural resources. The true value of data collection lies in its application, helping eliminate guesswork and refine farming methods. Peters et al. demonstrated how AI recommender systems (RS) combined with machine learning (ML) can leverage large data sets to solve agricultural challenges, enhance the efficiency of research, and improve the accuracy of food production estimates. This approach helps predict agroecosystem dynamics under changing environmental conditions, making the most of both scientific expertise and real-time data.

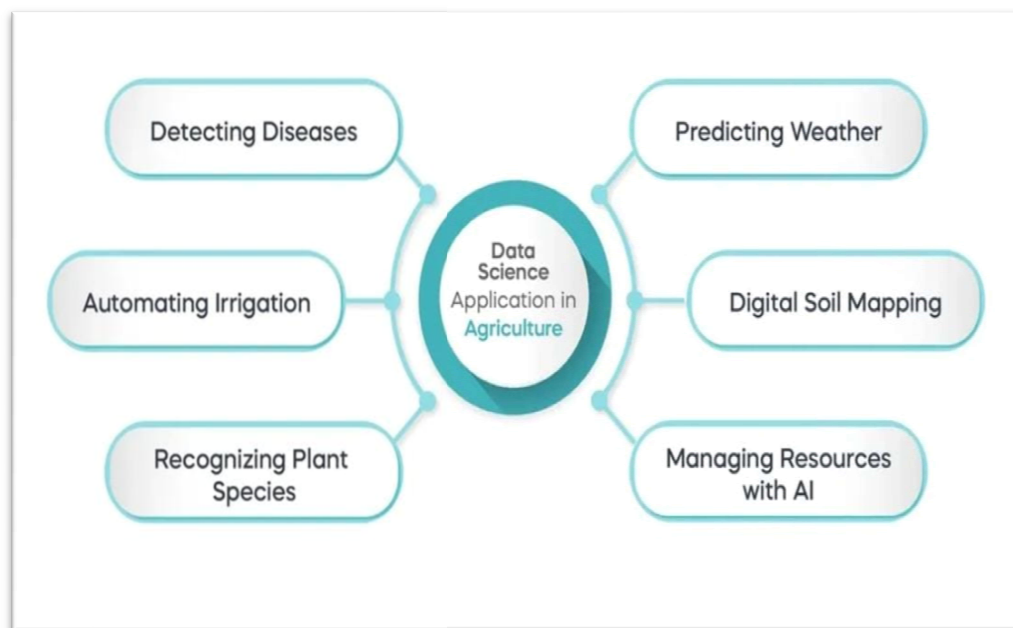


Figure 4 : data science application in agriculture

5. Genetic improvement of crops:

AI is revolutionizing crop genetic improvement by accelerating the process of identifying beneficial traits in plants. Machine learning models analyse vast amounts of genomic, phenotypic, and environmental data to predict which genetic variations will enhance crop yield, resistance to diseases, and tolerance to stressors like drought. This allows for more precise breeding, reducing the time and resources needed to develop improved crop varieties. AI-powered tools also enable a better understanding of plant genetics, speeding up the development of crops that can meet the growing global demand for food in a sustainable manner.

6. Disease detection:

Effective disease control is crucial for optimal agricultural yield, as plant and animal diseases significantly impact crop production. Factors such as genetics, weather, and soil conditions contribute to the spread of diseases, making management challenging, especially in large-scale farming. An integrated disease control approach that combines physical, chemical, and biological methods is essential, but it's time-consuming and costly. To address this, AI-based systems are used for disease detection and management. These systems utilize fuzzy logic, rule-based inference engines, and image recognition technologies to identify diseases and suggest treatments, enhancing early detection and providing timely interventions. AI tools, including mobile apps, can support farmers in managing crop diseases and pests more effectively, leveraging big data for site-specific decisions and sustainable practices.



Figure 5: Disease detection in crops

7. Weed management :

AI is revolutionizing weed management in agriculture by using computer vision, machine learning, and robotics. Key applications include:

- Weed Detection: AI systems identify weeds in real-time using cameras and sensors, improving accuracy in distinguishing them from crops.
- Precision Herbicide Application: AI enables targeted herbicide spraying, reducing chemical use and environmental impact.
- Weed Monitoring: Satellite and drone images, processed by AI, help map and predict weed growth, guiding proactive management.
- Robotic Weeding: Autonomous robots use AI to mechanically remove weeds without harming crops.
- Data-Driven Decisions: AI platforms analyse data to recommend efficient weed control strategies.
- AI enhances efficiency, sustainability, and cost-effectiveness in weed management, leading to healthier crops and higher yields.



Figure 6 : Herbicide spraying for crops.

Advantages of AI in Agriculture

1. Improved Crop Yields

AI systems can monitor crops through sensors, satellites, or drones to track the health and growth of plants in real time. They can analyse soil conditions, weather patterns, and growth stages.

Merits: By offering guidance on the optimal timing for watering, fertilizing, and harvesting, AI enables farmers to maximize crop yields. Additionally, it helps avoid overwatering and over-fertilization, which can harm crops and result in wasted resources.

2. Advanced Farming Techniques

AI-driven technologies like self-operating tractors, unmanned aerial vehicles (drones), and sensors can make choices based on data evaluation. For instance, soil sensors can monitor moisture content, and AI can direct irrigation systems to supply water solely to the regions that require it.

Merits : This helps conserve resources like water and fertilizers, reducing costs and minimizing environmental impact, while also improving crop productivity.

3. Early – stage illness and Pest Identification

Drones or distant sensors can survey fields and capture images that AI examines for indications of pests, diseases, or nutrient deficiencies. Machine learning algorithms can recognize trends in crop health from these images and data.

Merits : By spotting issues early, farmers can take focused measures (such as applying targeted pesticides or modifying irrigation) to avert widespread harm, thereby reducing crop loss.

4. Sustainable Resource Management

AI utilizes weather predictions, soil monitors, and past data to forecast the optimal quantity of resources (such as water, nutrients, etc.) required for crops. It can also recommend the most suitable planting and harvesting timelines based on climate patterns.

Merits : This guarantees that resources are utilized effectively, resulting in higher yields, less waste, and more eco-friendly farming practices. For example, AI can assist in lowering water usage by calculating precisely when and how much water to apply to crops.

5. Automated Farming Equipment

AI-driven autonomous agricultural machinery (such as driverless tractors, harvesters, and planters) can carry out repetitive tasks like sowing, weeding, and harvesting.

Merits : This reduces the need for human labour, which is particularly beneficial in regions where labour is limited or costly. These machines also operate with accuracy, minimizing mistakes and enhancing productivity.

6. Analytics - Based Decision Making

AI platforms analyse vast quantities of agricultural data (such as climate conditions, soil quality, crop performance, etc.) and offer farmers practical recommendations.

Merits : With analytical insights, farmers can make well-informed choices regarding irrigation, fertilization, crop rotation, and other factors. This can result in enhanced productivity, lower expenses, and increased profitability.

7. Supply Chain Optimization

AI can forecast demand trends, enhance stock management, and evaluate transportation routes to guarantee that crops are delivered to markets in the most effective manner.

Merits : This minimizes food waste and logistical expenses, ensuring that fresh produce reaches consumers more quickly and efficiently. AI can predict demand patterns, optimize inventory management, and analyse transportation routes to ensure that crops are delivered to markets in the most efficient way.

8. Labour Shortage Solution

In many regions, agricultural labour is in short supply. AI-powered machines (such as robotic harvesters) can automate labour-intensive tasks, providing an alternative to human labour.

Merits: Farmers can continue operations without relying on seasonal workers or manual labour, and this ensures that productivity doesn't suffer due to labour shortages.

9. Sustainability

AI can help optimize farming practices that conserve resources like water, reduce the use of chemicals, and minimize soil erosion. For instance, AI can calculate the most sustainable crop rotation plans.

Merits : This promotes more environmentally friendly farming practices that help preserve natural resources, reduce greenhouse gas emissions, and protect biodiversity.

Disadvantages of AI in Agriculture

1.High Initial Investment

The expense of adopting AI technologies, such as acquiring drones, automated machinery, sensors, and software, can be substantial.

Demerits : For small-scale farmers or those in developing regions, this can present a significant obstacle, as the upfront costs may be unaffordable.

2. Technical Knowledge Requirement

Operating AI-based tools often demands specialized skills in areas like data analysis, machine learning, or operating advanced equipment.

Demerits : Farmers may need to undergo training to effectively utilize AI technologies. In areas with limited access to education or technical resources, this can create a knowledge gap and hinder widespread use.

3. Reliance on Technology

AI systems are heavily reliant on technology such as sensors, internet connectivity, and machine learning algorithms to function properly.

Demerits : If there is a malfunction in the technology (e.g., a sensor fails or a machine breaks down), it could disrupt the entire farming process. This may lead to delays, crop losses, or added costs to repair the technology.

4. Data Privacy Concerns

AI systems depend on large amounts of data, such as weather trends, crop health, and market conditions. This data is often collected from farmers' fields and may be shared with third parties (such as tech companies or researchers).

Demerits: This raises concerns about data ownership, usage, and potential exploitation. Some farmers may be hesitant to share their data, fearing it could be sold or misused.

5. Job Losses

As AI and automation take over more tasks in agriculture (such as harvesting, weeding, and planting), there may be a decreased need for human labour in certain roles.

Demerits : This could result in job losses, particularly in rural areas where agriculture is a primary source of employment. Workers dependent on farming jobs may face challenges in adapting to new careers or industries.

6. Unequal Access

Large agribusinesses and wealthy farmers can afford to invest in AI technologies, smaller farmers or those in underdeveloped regions may not have the financial resources to do so.

Demerits : This creates a digital divide where larger farms benefit from cutting-edge technologies, while small-scale farmers are left behind, exacerbating inequalities in the agricultural sector.

7. Maintenance and Technical Issues

AI systems, particularly complex machinery like autonomous tractors and drones, require regular upkeep, software upgrades, and troubleshooting.

Demerits : This can be difficult or costly for farmers, especially if they lack access to technical support. Malfunctions can also lead to downtime, affecting overall productivity.

8. Information Overload

AI generates vast amounts of data, including sensor readings, crop imagery, weather data, and more. While this data is valuable, it can be overwhelming to process and act upon.

Demerits : Farmers may find it challenging to interpret and utilize this data without proper tools or expertise. This could result in confusion or poor decision-making, undermining the intended benefits of AI.

9. Environmental Impact of Technology

While AI can aid in promoting sustainable farming practices, the production and disposal of AI-powered machinery (such as drones, sensors, and automated tractors) can generate electronic waste and carbon emissions.

Demerits : If not managed properly, the environmental footprint of AI technology could counterbalance some of the ecological advantages it aims to offer agriculture.

10. Bias in Algorithms

AI models are trained using data, and if the data is biased or incomplete (for instance, not reflecting certain environmental factors), the AI's recommendations might be inaccurate.

Demerits : This could result in faulty advice, such as incorrect irrigation schedules or ineffective pest control methods, potentially harming crop yields.

Development of agricultural robots

The development of robots in agriculture, particularly for tasks traditionally performed by humans, such as pesticide spraying, fruit picking, and weed management. Agricultural robots have been researched since the 1980s, with Japan developing early robots like those for pesticide spraying. The main challenge in agricultural robotics is navigating real-world environments and maintaining precision. Early robots, such as AURORA, aimed to reduce the risk of human exposure to harmful chemicals in greenhouses. Examples include robots using GPS and machine vision for tasks like apple picking, and systems for mapping weed populations using embedded controllers. Challenges in weed management involve distinguishing crops from weeds, requiring image recognition and machine learning. Other innovations include the development of flexible grippers for fruit picking to minimize damage, and an advanced vision system for fruit recognition using OHTA colour space and Otsu threshold algorithms to ensure high accuracy in identifying ripe fruit.


Conclusion

Artificial intelligence (AI) is transforming agriculture by improving efficiency, productivity, and sustainability through technologies like machine learning, drones, and sensors for precision farming, pest control, and crop management. Over the past 34 years, AI has evolved from rule-based expert systems to advanced models like neural networks, fuzzy systems, and hybrid approaches. These innovations enable real-time, automated systems for better resource optimization and yield improvement. While AI holds significant potential to address food security and sustainable farming, challenges such as implementation costs, data privacy, and the need for skilled labour remain. Continued research aims to make agriculture more cost-effective and precise

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The Role of Medicinal Plants In Modern Pharmacology

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Introduction

Medicinal plants have served as the foundation of medical treatments for thousands of years, providing a wealth of bioactive compounds that continue to be instrumental in healthcare. Despite advances in synthetic chemistry, these natural resources remain invaluable for drug discovery and the development of modern pharmaceuticals. This chapter explores the historical significance, scientific advancements, and current applications of medicinal plants in pharmacology.

Features of Medical Plants:

- ❖ **Historical Significance:** Highlights the ancient use of medicinal plants across various civilizations, such as Mesopotamia, Egypt, China, and India, laying the foundation for modern drug discovery.
- ❖ **Phytochemistry:** Provides an in-depth discussion of bioactive compounds like alkaloids, flavonoids, terpenoids, glycosides, and polyphenols, and their biological activities (e.g., anti-inflammatory, antioxidant, antimicrobial, anticancer).
- ❖ **Drug Discovery:** Explores key examples of modern drugs derived from medicinal plants, such as aspirin, artemisinin, vincristine, and vinblastine, demonstrating their critical role in pharmacology.
- ❖ **Integration into Modern Medicine:** Examines how medicinal plants are utilized today, including phytomedicine, adjuncts in drug therapy, and nutraceuticals.
- ❖ **Challenges and Sustainability:** Identifies major challenges like sustainability, standardization, regulatory hurdles, and drug interactions, emphasizing the need for ethical and scientific approaches.

- ❖ **Technological Advancements:** Discusses tools like high-performance liquid chromatography, mass spectrometry, and AI-driven analysis in the identification and utilization of plant-based compounds.
- ❖ **Future Directions:** Outlines research priorities, including genomic profiling, synthetic biology, and computational methods, to maximize the potential of medicinal plants in modern pharmacology.
- ❖ **Global Health Impact:** Underlines the contribution of medicinal plants in addressing pressing health challenges and their potential for inspiring new therapeutic innovations.

Techniques involved in medicinal plants in modern pharmacology:

Advancements in science and technology have led to the development of sophisticated techniques for studying medicinal plants and their bioactive compounds. These include:

1. Extraction Techniques

- **Solvent Extraction:** Uses solvents like ethanol or methanol to extract bioactive compounds.
- **Supercritical Fluid Extraction:** Employs supercritical CO₂ for efficient and eco-friendly extraction.
- **Microwave-Assisted Extraction (MAE):** Enhances yield and reduces extraction time using microwave energy.

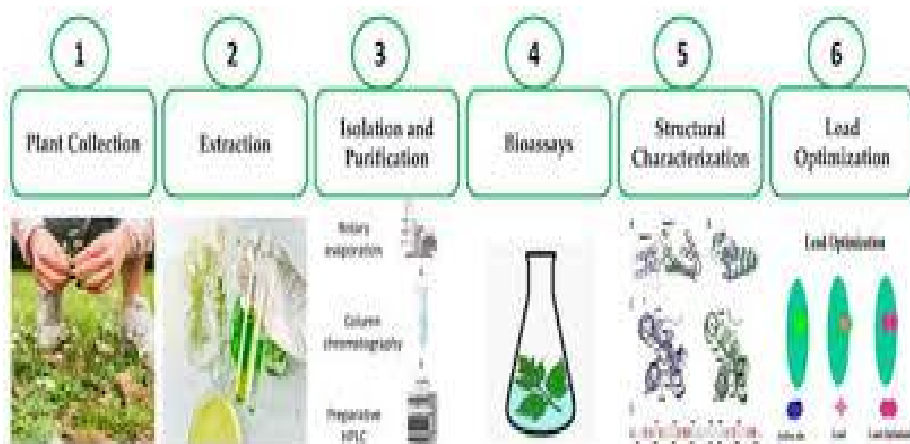


Figure 1: Techniques of medical plants.

2. Phytochemical Screening

- **Chromatographic Methods:** High-performance liquid chromatography (HPLC) and gas chromatography (GC) for separating and analyzing compounds.
- **Spectroscopy Techniques:** UV-Vis, infrared (IR), and nuclear magnetic resonance (NMR) spectroscopy for structural elucidation.

3. Bioassays

- **In Vitro Assays:** Evaluating bioactivity (e.g., antioxidant, anti-inflammatory) in controlled lab environments.
- **In Vivo Studies:** Testing efficacy and safety in animal models.
- **Cell Culture Techniques:** Investigating molecular mechanisms at the cellular level.

4. High-Throughput Screening (HTS)

Automated techniques to screen large libraries of plant extracts for potential therapeutic agents.

5. Genomic and Metabolomic Approaches

- **Genomic Analysis:** Identifying genes responsible for biosynthesis of bioactive compounds.
- **Metabolomics:** Profiling metabolites using mass spectrometry and NMR to understand plant metabolism.

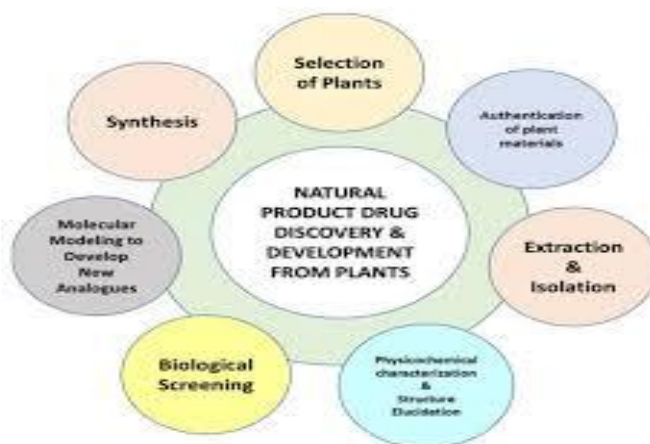


Figure 2:Flow chart for medical plants and products.

6. Synthetic Biology

Engineering microbes or plants to produce high yields of medicinal compounds by mimicking natural biosynthetic pathways.

7. Computational Tools

- **Molecular Docking:** Predicting interactions between plant-derived compounds and target proteins.
- **AI and Machine Learning:** Analyzing large datasets to identify novel bioactive molecules.

8. Nanotechnology

- **Nanoencapsulation:** Improving the delivery and bioavailability of plant-based compounds.
- **Nanoparticles:** Enhancing stability and targeting of phytochemicals in therapeutic applications.

Role in modern drug discovery:

Medicinal plants have contributed significantly to the discovery of modern drugs. Notable examples include:

- **Aspirin:** Derived from salicin in willow bark.
- **Artemisinin:** Isolated from *Artemisia annua*, a key treatment for malaria.
- **Vincristine and Vinblastine:** Extracted from the Madagascar periwinkle, used in chemotherapy.

High-throughput screening of plant extracts, coupled with bioinformatics, has accelerated the identification of novel therapeutic agents.



Figure 3: Role of modern drug discovery.

Integration into Modern Medicine

Phytomedicine and Herbal Drugs

Standardized extracts of medicinal plants are now used as phytomedicines. Examples include:

- **Milk thistle extract:** Used for liver disorders.
- **St. John's wort:** For mild depression and anxiety.

Adjuncts in Drug Therapy

Phytochemicals often serve as adjuncts to enhance the efficacy or reduce the side effects of synthetic drugs. For instance, ginger extracts can mitigate chemotherapy-induced nausea.

Nutraceuticals:

Medicinal plants are also utilized in nutraceuticals, which bridge the gap between nutrition and medicine. Omega-3 fatty acids, flavonoids, and polyphenols derived from plants are incorporated into dietary supplements for disease prevention.

Recent Updates in The Role of Medicinal Plants in Modern Pharmacology:

1. AI Integration in Phytochemistry

- Artificial intelligence and machine learning models are now being widely used to predict bioactivity, analyze molecular docking, and identify novel plant-based compounds, significantly reducing the time required for drug discovery.

2. CRISPR in Medicinal Plant Research

- CRISPR-Cas9 technology is enabling genetic editing of medicinal plants to enhance the production of specific bioactive compounds or to introduce traits that improve sustainability.

3. Metabolomics Advances

- Enhanced metabolomic tools, including ultra-high-resolution mass spectrometry, are providing deeper insights into the metabolic pathways of medicinal plants, aiding in the discovery of new therapeutic agents.

4. Plant Tissue Culture

- Innovations in plant tissue culture and bioreactors are improving the yield and purity of active compounds, making large-scale production more feasible and environmentally sustainable.

5. Green Chemistry Approaches

- Eco-friendly extraction and synthesis methods, such as ionic liquids and supercritical fluid technology, are gaining traction to ensure sustainability in medicinal plant utilization.

6. Microbiome Research

- Studies on the interaction between medicinal plants and soil or human microbiomes are shedding light on how plant metabolites influence health and drug efficacy.

7.COVID-19 Research

- Medicinal plants such as *Nigella sativa* (black cumin) and *Withania somnifera* (ashwagandha) have been explored for their potential antiviral and immunomodulatory effects during the COVID-19 pandemic.

8. Integration with Digital Health

- Wearable AI devices and digital health platforms are beginning to incorporate phytomedicines into holistic health monitoring and management systems.

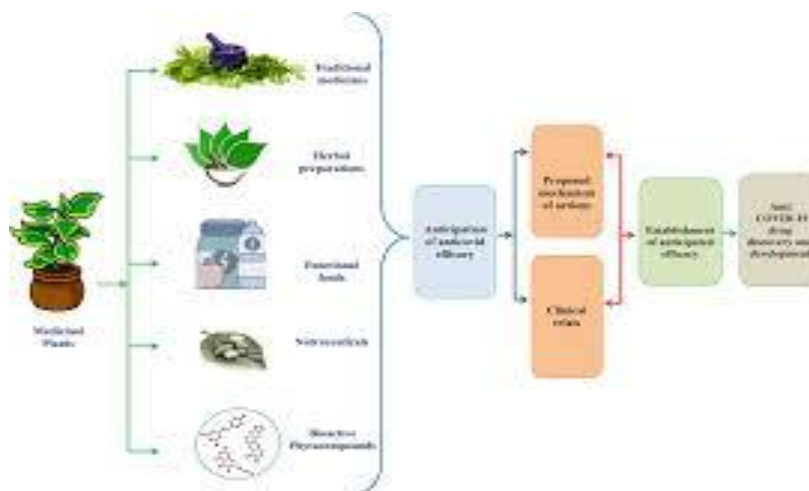


Figure 4: recent updates of medical plants

9. Blockchain in Supply Chain

- Blockchain technology is being implemented to ensure the traceability and quality control of medicinal plant products, addressing challenges in standardization and authenticity.

Biotechnological Importance of Medicinal Plants in Modern Pharmacology

Biotechnology has revolutionized the utilization of medicinal plants by enhancing their therapeutic potential and sustainability. Key contributions include:

1. Genetic Engineering

- ✓ **CRISPR-Cas9 Technology:** Enables precise editing of plant genomes to enhance the production of bioactive compounds or introduce desirable traits.
- ✓ **Transgenic Plants:** Engineered to produce pharmaceutical proteins, such as vaccines and therapeutic enzymes.

2. Plant Tissue Culture

- ✓ Callus Culture: Used for mass propagation of medicinal plants and the production of secondary metabolites.
- ✓ Hairy Root Cultures: Induced by *Agrobacterium rhizogenes* to enhance metabolite production in plants like ginseng and *Catharanthus*.
- ✓ Micropropagation: Ensures large-scale production of disease-free, genetically identical plant material.

3. Metabolic Engineering

- ✓ Pathway Manipulation: Enhancing or redirecting metabolic pathways to increase yields of valuable compounds like artemisinin or taxol.
- ✓ Synthetic Biosynthesis: Utilizing microorganisms like yeast or bacteria to synthesize plant-derived compounds through engineered pathways.

4. Bioreactor Technology

- ✓ Large-Scale Cultivation: Production of secondary metabolites in controlled environments, reducing reliance on wild plant populations.
- ✓ Elicitor Technology: Use of biotic or abiotic elicitors to stimulate the production of bioactive compounds in cultured cells.

5. Pharming

- ✓ Plant-Based Biopharmaceuticals: Developing plants as biofactories for producing therapeutic proteins, monoclonal antibodies, and vaccines. For example, tobacco plants have been engineered to produce antibodies for Ebola.

6. Omics Technologies

- ✓ Genomics: Sequencing medicinal plant genomes to identify genes involved in the biosynthesis of pharmacologically active compounds.
- ✓ Transcriptomics: Studying gene expression to understand the regulation of metabolite production.
- ✓ Proteomics and Metabolomics: Profiling proteins and metabolites to discover new therapeutic agents and understand plant physiology.

7. Sustainable Cultivation Practices

- ✓ Biotechnological Interventions: Developing stress-tolerant plant varieties to ensure stable supply under changing climatic conditions.

- ✓ Conservation Techniques: Cryopreservation and seed banking for the preservation of rare or endangered medicinal plants.

8. Nanobiotechnology

- ✓ Nanoparticle Synthesis: Using plant extracts for the green synthesis of nanoparticles with antimicrobial or anticancer properties.
- ✓ Targeted Drug Delivery: Employing plant-based compounds in nanocarriers to improve therapeutic efficacy.

Biotechnology not only enhances the therapeutic potential of medicinal plants but also ensures their sustainable utilization in modern pharmacology, bridging traditional knowledge with advanced science.

Future of medicinal plants in modern pharmacology:

The future of medicinal plants in modern pharmacology is poised to be transformative, driven by advancements in science and technology. Key trends and possibilities include:

1. Personalized Medicine

- ❖ Integration of phytomedicines into personalized treatment regimens based on an individual's genetic, metabolic, and microbiome profile.
- ❖ Development of targeted therapies using plant-derived compounds tailored to specific diseases or patient groups.

2. Synthetic Biology Innovations

- ❖ Leveraging synthetic biology to design and engineer plants or microorganisms for the sustainable and scalable production of complex medicinal compounds.
- ❖ Creating synthetic pathways to replicate plant-based bioactive molecules with enhanced efficacy.

3. Digital and AI Integration

- ❖ Use of artificial intelligence and big data analytics to identify new bioactive compounds and predict their therapeutic potential.
- ❖ Development of digital tools for monitoring the efficacy and safety of phytomedicines in real-time.

4. Global Collaboration

- ❖ Increased international cooperation to share knowledge, genetic resources, and biotechnological innovations for medicinal plant research.
- ❖ Harmonization of regulatory frameworks to facilitate global trade and acceptance of phytomedicines.

5. Eco-Friendly Practices

- ❖ Adoption of green chemistry techniques for sustainable extraction and production of plant-based compounds.
- ❖ Conservation of biodiversity through responsible sourcing, cultivation, and habitat restoration initiatives.

6. Integration with Nanomedicine

- ❖ Enhanced bioavailability and targeted delivery of phytochemicals using nanotechnology.
- ❖ Development of nanoparticle-based formulations to improve stability and reduce side effects.

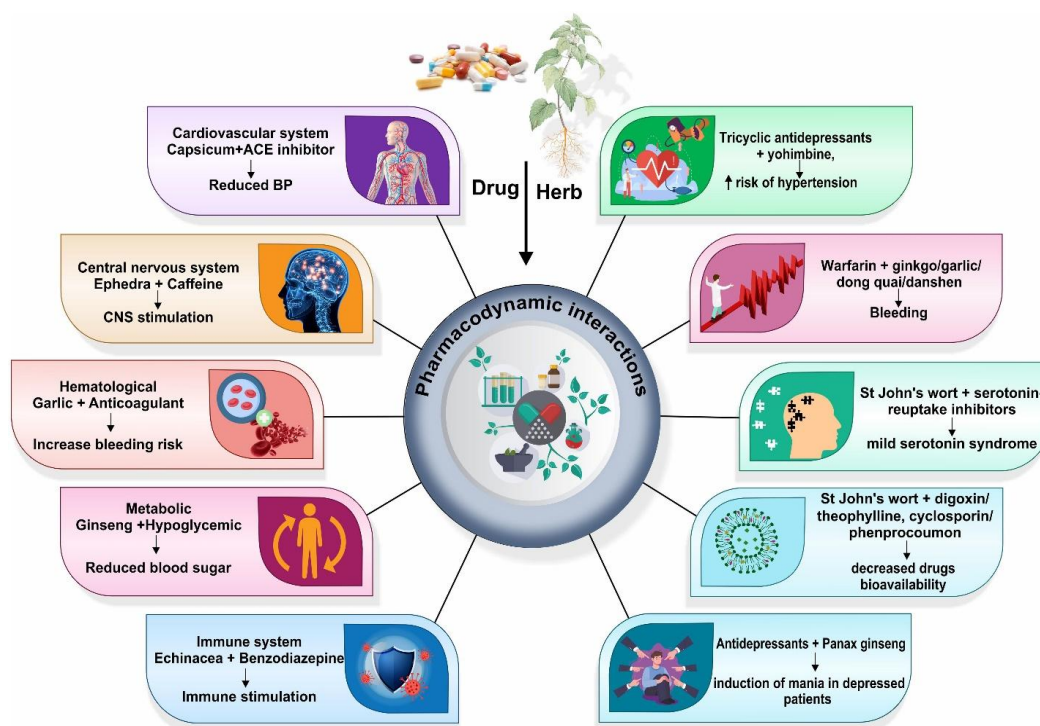


Figure:5 Future of medical plants in modern pharmacology.

7. Expanded Role in Preventive Medicine

- ❖ Use of medicinal plants in preventive healthcare through nutraceuticals and functional foods.
- ❖ Incorporation of phytochemicals into wellness and lifestyle products for holistic health management.

8. Education and Awareness

- ❖ Increased public and professional education on the benefits, safety, and applications of medicinal plants.
- ❖ Initiatives to bridge traditional knowledge with modern scientific research for wider acceptance and utilization.
- ❖ By embracing these innovations and addressing existing challenges, medicinal plants will continue to play a pivotal role in modern pharmacology, offering sustainable and effective solutions to global healthcare needs.

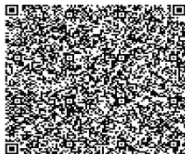
Conclusion

Medicinal plants remain indispensable in modern pharmacology. They not only offer a treasure trove of bioactive compounds but also inspire synthetic analogs for innovative treatments. As we continue to integrate traditional knowledge with cutting-edge science, medicinal plants will undoubtedly play a pivotal role in shaping the future of medicine.

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Tocilizumab, An Anti iL – 6 Receptor Monoclonal Antibody To Treat Severe Covid-19 Patients During Pandemic.

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Abstract

Tocilizumab (TCZ), a humanized monoclonal antibody targeting interleukin-6 receptors (IL-6R), has demonstrated efficacy in mitigating severe inflammatory responses in COVID-19, including cytokine storm syndrome (CSS). Administered via intravenous or subcutaneous routes, TCZ blocks IL-6 signaling, thereby reducing systemic inflammation. Clinical trials and observational studies have reported improved outcomes in critically ill COVID-19 patients treated with TCZ, particularly those with elevated C-reactive protein (CRP) levels. However, its effectiveness is diminished in patients with hyperglycemia or elevated IL-6 levels, suggesting the need for tailored glycemic management during treatment. Comparative analyses highlight TCZ's advantages over alternative cytokine inhibitors and anti-inflammatory agents such as corticosteroids, JAK inhibitors, and NSAIDs. Despite its therapeutic potential, TCZ and corticosteroids have been implicated in medication-related osteonecrosis of the jaw (MRONJ) in post-COVID-19 patients, raising concerns about long-term adverse effects. These findings underscore the critical importance of balancing the therapeutic benefits of TCZ and corticosteroids against their potential complications, particularly in severe and ICU-managed COVID-19 cases.

Key Words :Tocilizumab (TCZ), COVID-19, Interleukin-6, Cytokine storm, corticosteroids, medication – related osteonecrosis of jaw.

1. Introduction

Tocilizumab, an interleukin-6 (IL-6) receptor inhibitor, has emerged as a key therapeutic option for managing severe COVID-19, particularly in patients with cytokine release syndrome (CRS). This chapter examines its mechanism of action, clinical applications, and associated risks, including its role in reducing inflammation, improving survival outcomes, and potential adverse effects such as medication-related osteonecrosis of the jaw (MRONJ). Additionally, the use of corticosteroids, another widely implemented treatment during the pandemic, is discussed in the context of post-COVID-19 complications, highlighting the need for cautious application to mitigate long-term adverse effects.

2 .An Overview of Tocilizumab

Tocilizumab (TCZ) is a recombinant humanized monoclonal antibody belonging to the immunoglobulin G1 κ subclass, specifically targeting soluble and membrane-bound interleukin-6 receptors (IL-6R). It is commercially available under the trade name RoActemra in the European Union (EU) and Actemra in the United States. Initially introduced in Japan in 2008, TCZ was subsequently approved for use in the EU in 2009 and the US in 2010 [1].

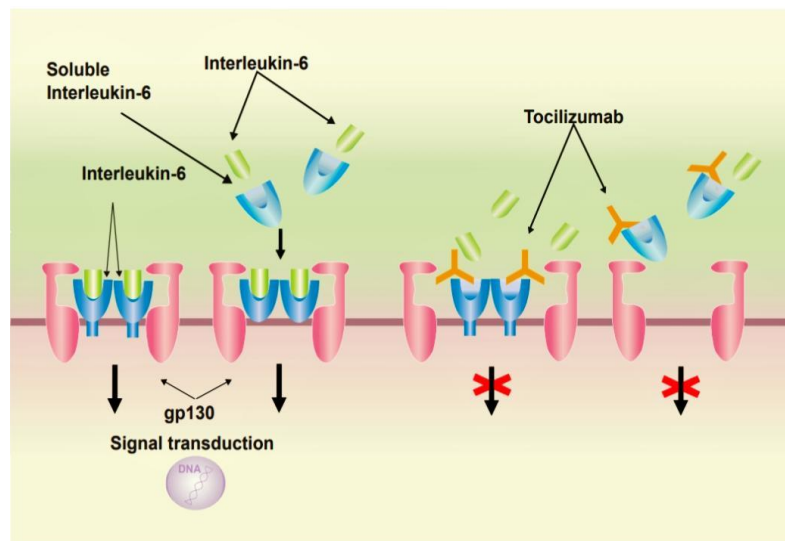


Figure 1: Interleukin-6 (IL-6) signaling begins when IL-6 binds to the IL-6 receptor (IL-6R) on the cell surface or its soluble form (sIL-6R). This interaction causes gp130 dimerization, which activates signaling. Tocilizumab blocks IL-6 signaling by preventing its binding to IL-6R or sIL-6R.

Tocilizumab functions by inhibiting the interaction of IL-6 with its receptors, thereby mitigating the cytokine's pro-inflammatory effects. This inhibition occurs through competitive binding to both the soluble and membrane-bound forms of the IL-6 receptor, effectively reducing IL-6-mediated signaling and its associated inflammatory activity [1].

Tocilizumab dosing regimens employed in these clinical trials encompassed both banded weight-based approaches (e.g., 400 mg for individuals weighing 41–65 kg, 600 mg for those weighing 66–90 kg, and 800 mg for patients exceeding 90 kg) and continuous weight-based strategies (e.g., 8 mg/kg, capped at a maximum dose of 800 mg). Nevertheless, in the absence of robust exposure–response data to inform dose optimization for individuals with COVID-19, the most appropriate tocilizumab dosing strategy for this patient population remains indeterminate [2]. Tocilizumab is administered intravenously at varying doses (median: 400 mg; interquartile range: 320–480 mg). Studies in rheumatology demonstrate comparable efficacy and safety between subcutaneous and intravenous forms. The subcutaneous formulation, approved in regions such as the United States, Canada, and the European Union, is more accessible in some countries, whereas the intravenous form faces storage limitations. However, subcutaneous tocilizumab has not been extensively studied for COVID-19, and dose conversion from intravenous to subcutaneous remains unclear [3].

Inclusion criteria for the study consisted of: individuals with a confirmed diagnosis of SARS-CoV-2 via polymerase chain reaction (PCR); patients recently hospitalized within the first three days of admission and subsequently admitted to the intensive care unit (ICU) within the preceding 24 hours, as well as those not requiring ICU admission but demonstrating rapidly escalating oxygen requirements, irrespective of gender [4].

Exclusion criteria encompassed: elevated procalcitonin levels; documented hypersensitivity to tocilizumab; ongoing bacterial or fungal infections; alanine aminotransferase (ALT) or aspartate aminotransferase (AST) levels exceeding five times the upper limit of normal; platelet counts below 50,000/mm³; neutrophil counts under 500 cells/mm³; pregnancy; breastfeeding; and individuals younger than 18 years of age [4].

3.The Role of Interleukin-6 In Covid-19

Severe SARS-CoV-2 infection outcomes, including critical illness or mortality, are associated with host genetic predispositions, advanced age, male sex, comorbidities (e.g., chronic pulmonary and cardiovascular diseases, diabetes mellitus, and obesity), and elevated cytokine levels (IL-6, TNF- α ,

IFN- γ) or other serum biomarkers. However, the marked variability in clinical manifestations remains poorly understood. Emerging research links the gut microbiome to the pathophysiology of COVID-19, suggesting that gut dysbiosis may downregulate ACE2 expression, thereby influencing viral entry and replication. Furthermore, gut barrier dysfunction in COVID-19 may exacerbate systemic inflammation and increase vulnerability to respiratory infections [5].

Cytokines, defined as soluble proteins secreted by a diverse array of cells, play a pivotal role in facilitating intercellular communication and coordination within the immune system. They exhibit both pro-inflammatory and anti-inflammatory functions, while a specific subset, known as chemokines, is instrumental in directing leukocyte migration to sites of inflammation or tissue injury. In the context of cytokine storm (CS), various cell types, signaling pathways, and cytokines contribute to pathogenesis, which is influenced by the underlying etiology. Consequently, numerous studies have sought to identify cytokines associated with adverse clinical outcomes in SARS-CoV-2 infection. Notably, elevated levels of cytokines such as interleukin-6 (IL-6), interleukin-8 (IL-8), interleukin-10 (IL-10), and tumor necrosis factor-alpha (TNF- α) are strongly correlated with increased mortality risk, as these are among the most extensively studied in this domain [9].

Immune dysregulation is marked by an excessive, uncontrolled release of cytokines, leading to secondary organ damage. This phenomenon, termed cytokine release syndrome (CRS) or cytokine storm (CS), was first identified in the early 1990s as an exaggerated immune response triggered by the presence of a pathogen or inflammatory stimuli. Comparable conditions have been documented in other viral pneumonias, including severe acute respiratory syndrome (SARS) and avian influenza. CS manifests through a spectrum of clinical complications, such as acute respiratory distress syndrome (ARDS), vasodilatory shock, and disseminated intravascular coagulation, all of which have also been observed in severe cases of COVID-19 [9].

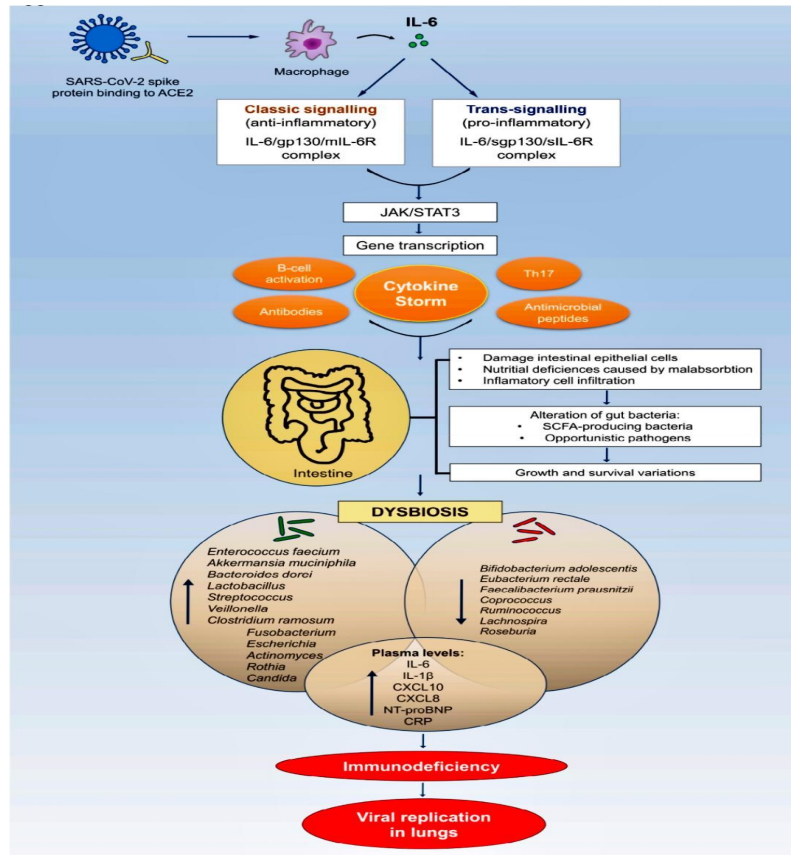


Figure 2: IL-6 signaling mediates cytokine–microbiome interactions, influencing immune response efficacy and viral replication, with dysbiosis marked by increased or decreased specific bacterial species.

4. Outcome of Covid-19 Patients Treated With Tocilizumab

a) A 42-year-old male with metastatic sarcomatoid clear cell renal cell carcinoma was hospitalized for fever, bone metastases pain, and systemic treatment planning. Initially presenting with isolated fever, he later developed mild cough and tested positive for SARS-CoV-2. Chest CT revealed bilateral ground-glass opacities consistent with COVID-19. Despite antiviral therapy with lopinavir-ritonavir, he experienced worsening dyspnea and hypoxemia requiring increased oxygen support. Intravenous tocilizumab (8 mg/kg, two doses) was administered, resulting in rapid clinical improvement, resolution of fever, reduced oxygen dependence, and decreased C-reactive protein levels.

Follow-up imaging showed partial regression of pulmonary infiltrates, and the patient fully recovered from COVID-19 symptoms [6].

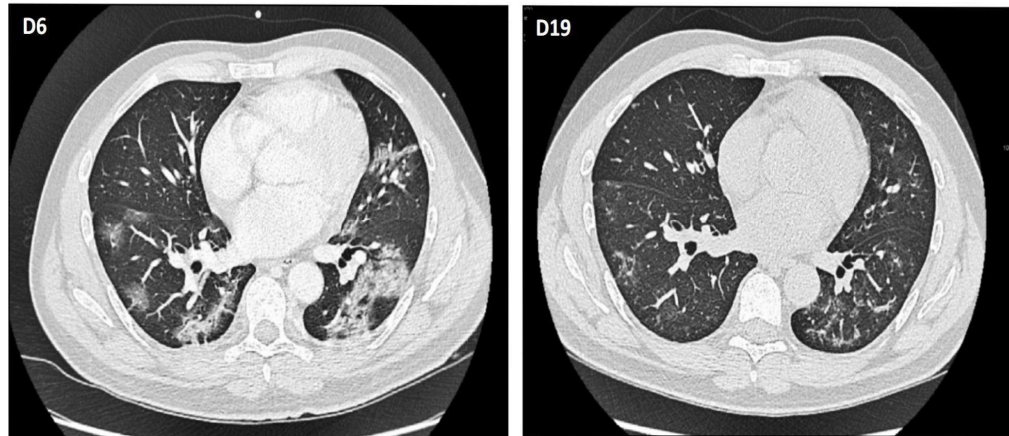


Figure 3: Pulmonary CT scan (Computerized tomography) of the patient before (day 6) and after (day 12) of tocilizumab administration shows Gradual recovery.

b) In overall survival analysis, TCZ demonstrated an advantage over SOC (HR: 0.499; 95% CI: 0.262–0.952; $p = 0.035$) based on Cox regression. While TCZ showed no benefit in severe cases, it significantly improved outcomes in critical patients [7].

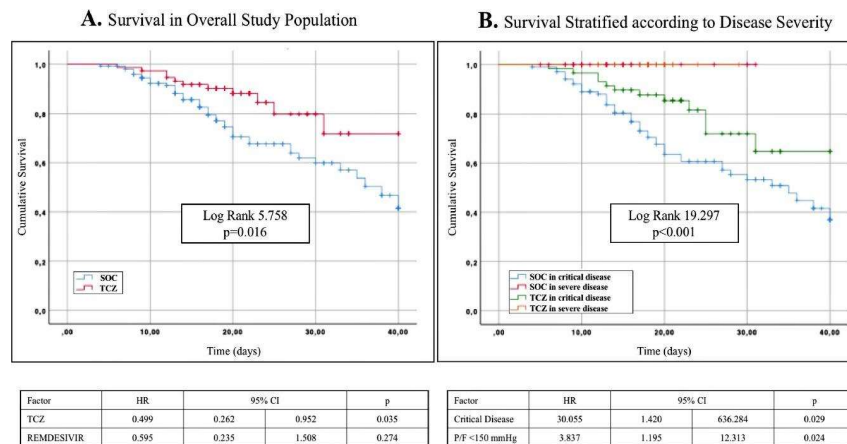


Figure 4: Kaplan-Meier Survival Curves and Cox Regression Models for Overall Survival (A) and Stratified by Disease Severity (B)

c) The fluctuations in IL-6 levels before and after tocilizumab treatment are depicted, highlighting an initial transient elevation in IL-6 concentrations shortly following tocilizumab administration. This was subsequently followed by a gradual decline over time. During the post-treatment phase, IL-6 levels remained persistently higher in non-survivors compared to survivors, with each group demonstrating distinct post-treatment trajectories [8].

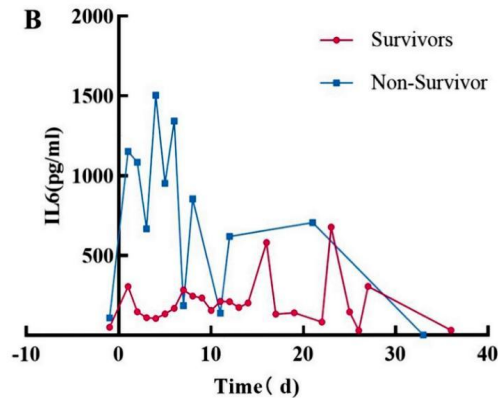


Figure 5: This figure illustrates the temporal variations in interleukin-6 levels between survivors and non-survivors.

d) Survival Analysis Stratified by Post-Treatment IL-6 Levels: Survival curves indicated significantly reduced survival outcomes in patients with elevated post-treatment IL-6 levels [8].

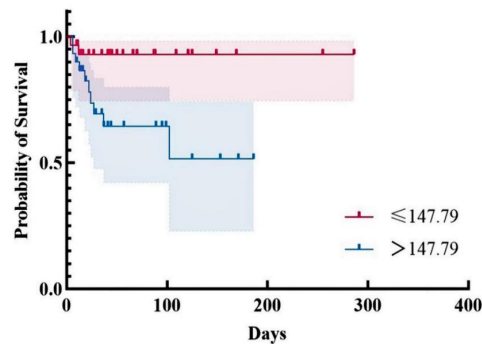


Figure 6: Analysis of Interleukin-6 Levels and Prognostic Outcomes Following Tocilizumab Treatment

e) From December 2022 to March 2023, during the COVID-19 pandemic in China, five patients with hematologic malignancies and severe COVID-19 were effectively managed using tocilizumab. This data provides a retrospective analysis of these cases to evaluate the therapeutic efficacy and safety of tocilizumab, contributing to the evidence base and sharing their center's clinical experience in treating severe COVID-19 infections [10].

Table 1: Baseline Clinical Profiles of Patients Prior to Tocilizumab Administration.

Characteristics	Case 1	Case 2	Case 3	Case 4	Case 5
Age	71	67	54	68	32
Gender	Female	Female	Female	Female	Male
Hematology malignancies	Non Hodgkin lymphoma/ Chronic Myelogenous Leukemia	Multiple myeloma	Acute leukemia	Acute leukemia	Myelodysplastic Syndrome
Treatment received by the patient	Chemotherapy+Targeted therapy	CAR-T	Chemotherapy	Chemotherapy +HLA mismatched donor stem cell transfusion	Chemotherapy
Hematology malignant state	First-line treatment	Refractory	Refractory	Complete Remission	Partial Remission
ECOG score	3	3	3	3	3
Accompanying diseases	Cerebral infarction	Chronic hepatitis B		Chronic hepatitis B	
Clinical and laboratory examinations					
Heating ($\geq 38.0^{\circ}\text{C}$)	Yes	Yes	Yes	Yes	Yes
Dyspnea with hypoxemia	Yes	Yes	Yes	No	Yes
Respiratory support	High flow oxygen mask	High flow oxygen mask	High flow oxygen mask	Low flow oxygen	High flow oxygen mask
Neutrophils ($\times 10^9/\text{L}$)	0.74	4.28	0.54	0.03	4.28
Hemoglobin (g/L)	84	105	58	46	48
Platelet ($10^9/\text{L}$)	61	176	103	41	15
Alanine transaminase (U/L)	75	15	24	17	53
Creatinine ($\mu\text{mol/L}$)	75	54	41	50	58
C-reactive protein (mg/L)	20.4	22.14	81.02	114.59	172.78
IL-6 (pg/mL)	193.69	287.74		133.48	56.38
HBV-DNA	Negative	Negative	Negative	Negative	Negative
EB-DNA	Negative	Negative	Negative	Negative	Negative
CMV-DNA	Negative	Negative			Negative
COVID-19 nucleic acid CT value	28	N/A*	27	24	37
Treatment of chronic hepatitis B		Entecavir Tablets 0.5mg qd+Tenofovir Disoproxil Fumarate Tablets 300mg qd		Entecavir Tablets 0.5mg qd	
Treatment of COVID-19					
Nirmatrelvir/ritonavir	No	Yes	Yes	Yes	Yes
Corticosteroid	Yes	Yes	Yes	No	Yes

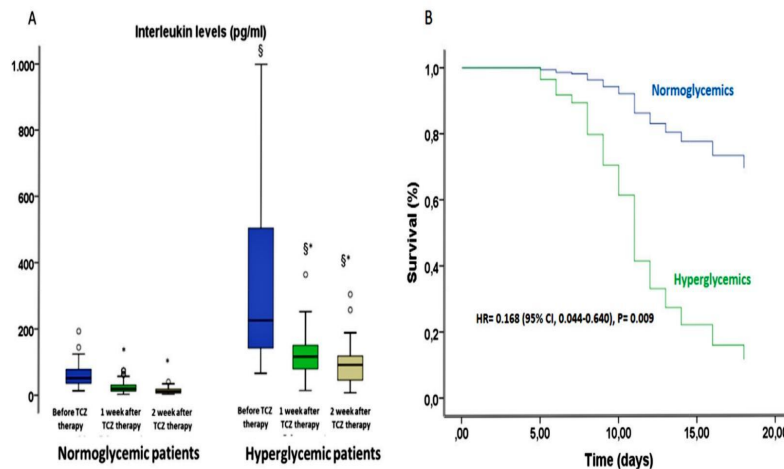


Figure 7: All patients received intravenous tocilizumab (TCZ) at 8 mg/kg, with 22.6% of hyperglycaemic and 10.6% of normoglycaemic patients needing two doses. Hyperglycaemic patients were 65.7 ± 13.4 years old on average, compared to 66.6 ± 12.2 years for normoglycaemic patients. Men accounted for 61.8% of hyperglycaemic and 72.3% of normoglycaemic groups. Illness lasted 17.6 ± 7.2 days for hyperglycaemic and 18.1 ± 6.6 days for normoglycaemic patients. Hypertension affected 61.3% of hyperglycaemic and 55.3% of normoglycaemic patients, while dyslipidaemia was more common in the hyperglycaemic group (32.3% vs. 14.9%). Smoking rates were similar. No significant differences were found in blood pressure, creatinine, or troponin levels. Both groups received TCZ and hydroxychloroquine as standard treatment. Hyperglycaemic patients had higher IL-6 levels before and after TCZ, and survival rates were lower. Data were analyzed using SPSS with $P < 0.05$ considered significant.

The study found no differences in hyperglycaemia treatment among hospitalized patients, as all received insulin therapy, and other hypoglycaemic drugs were discontinued. Subcutaneous insulin was initiated upon admission for hyperglycaemic patients, regardless of diabetes history. Higher IL-6 levels were observed in hyperglycaemic patients at admission, persisting despite tocilizumab (TCZ) treatment. Risk-adjusted analyses showed TCZ reduced severe outcomes in normoglycaemic patients but not in hyperglycaemic ones. Hyperglycaemic patients without a prior diabetes diagnosis had a higher risk of

severe disease compared to normoglycaemic and diabetic hyperglycaemic patients [12].

Previous studies suggest stress hyperglycaemia in non-diabetic patients increases in-hospital mortality, potentially due to inflammation and coagulability reduction from prior diabetes treatments. Elevated IL-6 levels in hyperglycaemic patients may weaken TCZ effectiveness. Severe COVID-19 can cause cytokine release syndrome (CRS), leading to immune dysfunction, organ damage, and inflammation. Hyperglycaemia may exacerbate CRS by promoting cytokine storms, including elevated IL-6, which correlate with severe outcomes [12].

Hyperglycaemic patients in this study had fivefold higher IL-6 levels than normoglycaemic ones. Elevated blood glucose may independently drive inflammatory responses, worsening COVID-19 outcomes. TCZ efficacy may decrease in patients with high baseline IL-6 levels, as shown in rheumatoid arthritis and COVID-19 studies. The lack of HbA1c data limits conclusions about the causes of hyperglycaemia in non-diabetic patients, which could reflect undiagnosed diabetes or stress responses [12].

These findings highlight the importance of glycaemic control in COVID-19 management, particularly for hyperglycaemic patients receiving TCZ. This has implications for clinical trials and optimizing treatment protocols for severe COVID-19 [12].

5. Comparative Analysis of Tocilizumab And Other Covid-19 Treatments In A Therapeutic Approach

Cytokine Inhibitors

- Tocilizumab: IL-6 receptor monoclonal antibody improving respiratory parameters and inflammatory markers; early administration reduces mortality.
- Sarilumab: Similar to tocilizumab, used to control pulmonary inflammation.
- Anakinra: IL-1 antagonist effective in cytokine storm syndromes, reducing respiratory distress without contraindications [11].

NSAIDs

- Nonselective COX Inhibitors: Includes aspirin, ibuprofen, diclofenac, naproxen; widely used for pain management. Adverse effects include cardiovascular (CV), gastrointestinal (GI), and renal complications.

Chronic NSAID use may offer mild protective effects against COVID-19 but could mask early symptoms.

- Selective COX-2 Inhibitors: Drugs like celecoxib and etoricoxib show anti-inflammatory benefits. Celecoxib in high doses is associated with reduced disease progression, though risks of myocardial infarction vary by dosage.
- Naproxen: Exhibits antiviral activity by inhibiting viral nucleoprotein binding; further safety evaluation is needed.
- Ibuprofen: Mixed effects in COVID-19 patients; early use may reverse lymphocytopenia, but some studies link it to worsened conditions.
- Indomethacin: Demonstrates antiviral activity; low doses may reduce cytokine storm, but renal dysfunction and coronary risks require caution [11].

Corticosteroids

- Glucocorticoids: Effective in mitigating inflammatory responses in severe COVID-19. Examples include dexamethasone and methylprednisolone, shown to reduce mortality and ICU stay in ventilated patients.
- Potential Risks: Adrenal suppression, delayed viral clearance, and increased risk of hyperglycemia, hypokalemia, and superinfections.
- Selective Applications: Administered in cases of severe inflammation, such as cytokine storms and pericardial disease [11].

JAK Inhibitors

- Baricitinib and Tofacitinib: Inhibit viral entry into lungs via JAK-STAT signaling; caution advised due to leukopenia risks.
- Fedratinib: Demonstrated efficacy in mitigating cytokine storm-associated conditions [11].

Antimalarials

- Hydroxychloroquine and Chloroquine: Exhibit antiviral properties but are associated with cardiotoxic effects, especially when combined with azithromycin. Limited evidence supports clinical efficacy [11].

Antiviral Agents

- Remdesivir: Reduces recovery time in adult patients; antiviral and anti-inflammatory effects.

- Favipiravir: Selectively inhibits RNA-dependent RNA polymerase, effective in severe COVID-19.
- Lopinavir and Ritonavir: Exert anti-inflammatory actions but with limited evidence of efficacy [11].

Colchicine

- Mechanism: Reduces cytokine release via NLRP3 inflammasome inhibition. Beneficial in systemic inflammation and pericarditis.
- Role in COVID-19: Adjunctive therapy to prevent disease progression to critical stages [11].

Herbal and Alternative Agents

- Hesperidin: Blocks viral entry via ACE2 receptor inhibition; possesses anti-inflammatory and antioxidant properties [11].

6. Impact of Tocilizumab Administration In Covid-19 Patients Requiring (Icu) Management

In this multicenter observational study of COVID-19 patients requiring ICU support, tocilizumab was associated with a reduction in hospital-related mortality, particularly in patients requiring mechanical ventilation and those under 65 years. A post-hoc analysis indicated reduced mortality in patients with baseline C-reactive protein (CRP) levels ≥ 15 mg/dL. These findings suggest that tocilizumab may be an effective treatment for severe COVID-19, pending confirmation by randomized trials [13].

Cytokine storm, primarily driven by elevated IL-6 levels, is a leading cause of death in late-stage SARS-CoV-2 infection. Tocilizumab, targeting IL-6, was associated with improved survival, especially in patients with elevated CRP levels. Reduction in CRP levels was observed in tocilizumab-treated patients, supporting its potential efficacy in inflammatory-driven COVID-19 progression [13].

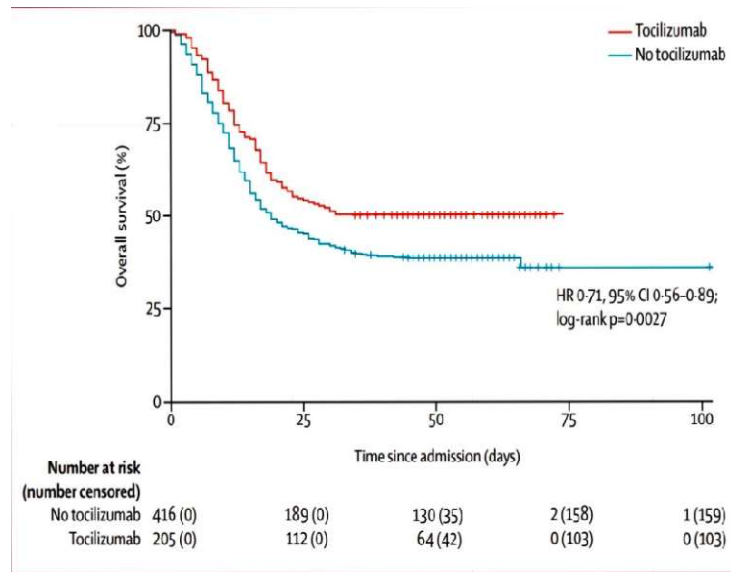


Figure 8: Survival outcomes in propensity score-matched patients

Steroid use in the recovery trial improved survival, but no similar benefit was observed in their cohort. Secondary bacterial infections were not significantly higher in the tocilizumab group, and hydroxychloroquine use had no significant impact on the findings. Limitations include the observational design, potential confounders, and regional biases, as well as the possibility of misclassification and sampling bias. Overall, tocilizumab shows promise in reducing mortality in ICU patients with severe COVID-19, warranting further investigation through randomized controlled trials [13].

7. Medication Related Osteonecrosis of The Jaw (Maxillary Osteonecrosis) In Post-Covid-19 Patients

Tocilizumab:

Medication-related osteonecrosis of the jaws (MRONJ) represents a significant pathological condition frequently associated with antiresorptive, immune-modulating, and antiangiogenic pharmacological agents. Initially identified in connection with bisphosphonates, the spectrum of implicated medications has since expanded. Emerging concerns have been articulated regarding the potential of tocilizumab, an interleukin-6 (IL-6) receptor-inhibiting monoclonal antibody employed in the management of conditions such as rheumatoid

arthritis and, more recently, COVID-19, to precipitate MRONJ, particularly among post-COVID-19 patient cohorts [14].

A clinical case has been documented involving a 36-year-old male presenting with tooth mobility and localized pain in the posterior region of the right maxilla subsequent to therapeutic intervention with tocilizumab and dexamethasone for COVID-19. Despite the administration of antibiotic therapy, the necrotic process persisted, necessitating more extensive surgical intervention, which ultimately resulted in significant improvement and sustained remission over a two-year follow-up period. This case underscores the imperative for heightened clinical vigilance and systematic investigation into the potential risk of MRONJ in patients receiving tocilizumab, emphasizing the critical need for healthcare professionals to anticipate, identify, and effectively manage this complication [14].

Corticosteroids:

A cohort of 20 patients (11 males and 9 females) who had recovered from COVID-19 and were diagnosed with osteonecrosis of the jaw (ONJ) within a 6-month timeframe was included in this study. The mean age of the cohort was determined to be 56.4 years (\pm standard deviation [SD]: 5.9 years). Of these, nine patients who successfully completed all follow-ups and fulfilled the inclusion criteria were selected for further analysis. The detailed characteristics of these cases are summarized in Table 1. The mean age of the analyzed subset was 53.7 years (\pm SD: 5.9 years; range: 44–64 years), with males comprising 56% (5/9) of the cohort. The majority of patients (78%, 7/9) underwent treatment in home isolation, while 22% (2/9) required hospitalization; notably, none were admitted to an intensive care unit during their COVID-19 treatment. All included patients reported the use of corticosteroids for the management of COVID-19. The average duration of corticosteroid administration was 9.6 days (\pm SD: 3.3 days; range: 6–20 days) [15].

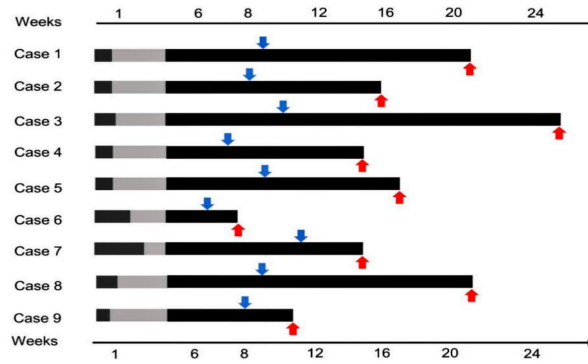


Figure 9: The timeline depicts corticosteroid use in weeks (dark gray bar), with (blue and red arrows) indicating symptom onset/tooth extraction and PCRONJ diagnosis, respectively. (The black bar represents the period between COVID-19 recovery and PCRONJ diagnosis, while the duration of COVID-19 infection (light gray bar) is undetermined.

In this study, all patients were administered corticosteroids after an average duration of 9.6 days for the treatment of COVID-19, although specific dosage data were unavailable. It was observed that 78% of the cohort received treatment at home, suggesting these individuals were predominantly in the mild or moderate stages of the disease. Notably, four patients who had been treated with corticosteroids at a mean cumulative dose of 562 mg over approximately 13 days developed avascular necrosis in the maxillae, without evidence of microbial or fungal infection. Although the precise dosage administered to these cases remains unspecified, the potential adverse effects of corticosteroid therapy on the development of post-COVID-19-related osteonecrosis of the jaw (PCRONJ) should not be underestimated [15].

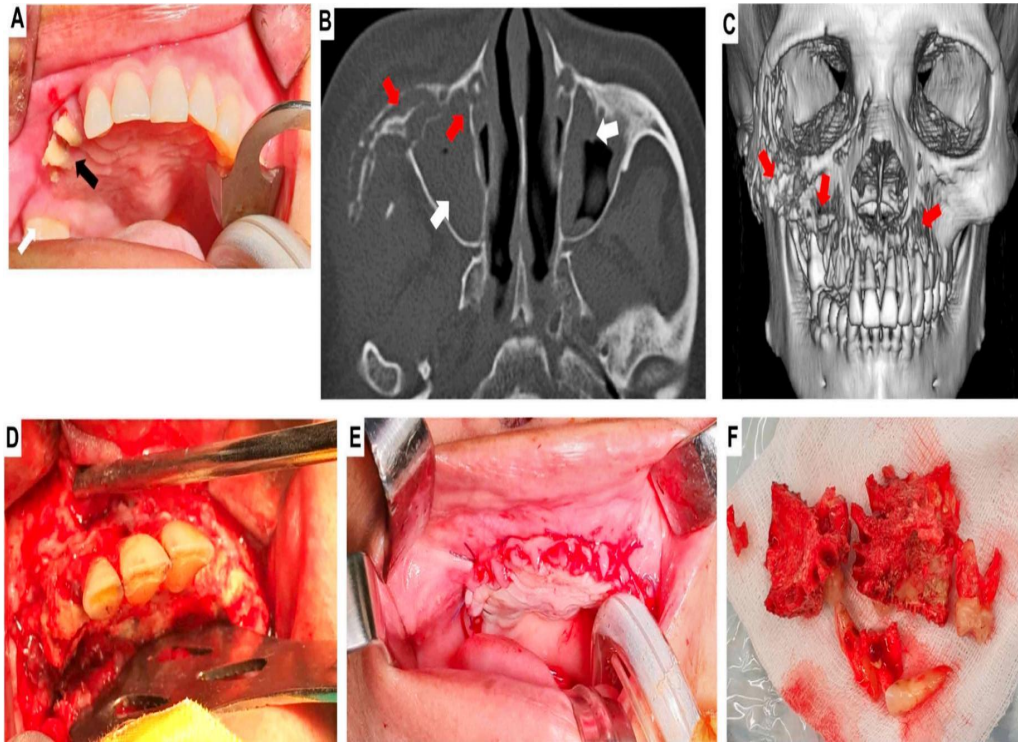


Figure 10: Case Report #8: (A) Clinical examination revealed exposed necrotic bone with purulent discharge in the right maxilla (indicated by the black arrow) and a mobile dentoalveolar segment in the anterior maxillary region.(B) Axial imaging demonstrated extensive destruction of the anterior and medial walls of the right maxillary sinus (red arrows), accompanied by significant opacification of both maxillary sinuses (white arrows).(C) Three-dimensional facial reconstruction highlighted severe bone loss in the alveolar region bilaterally, as well as destruction of the right maxilla and right zygoma (red arrows).(D) Necrotic bone was exposed through a sulcular incision to facilitate surgical intervention.(E) Primary closure of the maxillary defect was achieved following partial maxillectomy.(F) Necrotic bone fragments debrided from the maxilla were removed during the procedure.

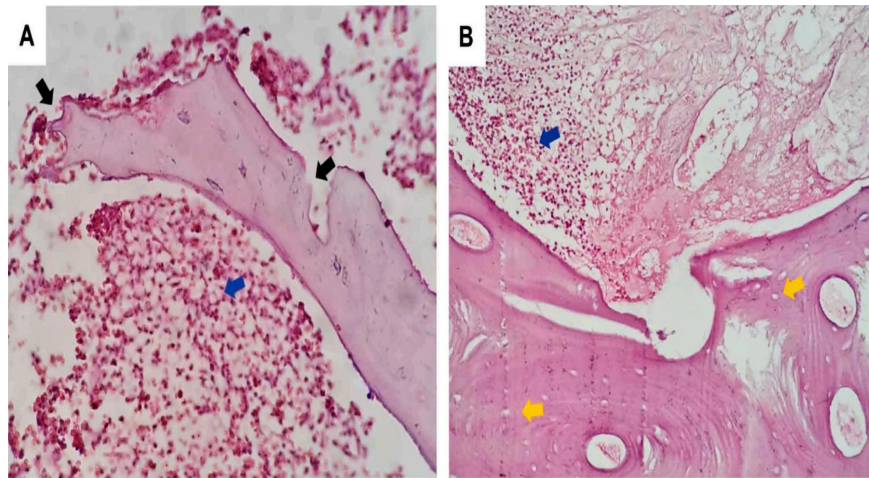


Figure 11: A histopathological analysis reveals discontinuous trabecular bone with resorptive borders (indicated by black arrows) (A) and empty osteocytic lacunae within the cortical bone (highlighted by yellow arrows) (B). Both compartments exhibit the presence of inflammatory infiltrates (marked by blue arrows). The observations were made on decalcified bone sections stained with Hematoxylin-Eosin and examined under a light microscope.

Corticosteroids, including dexamethasone and methylprednisolone, have been extensively employed globally for their anti-inflammatory properties in managing COVID-19, despite the lack of comprehensive scientific validation for this approach. Emerging research on patients recovering from COVID-19 has highlighted an association between corticosteroid administration and the onset of osteonecrosis. Steroids are known to inhibit angiogenesis, suppress bone remodeling, and induce thrombus formation through endothelial cell damage. Moreover, the jaw bones are particularly vulnerable to drug-related effects due to their heightened vascularization and metabolic activity, especially in the maxillofacial region, including the alveolar bone, which endures consistent mechanical forces. These findings provide compelling evidence to support the assertion that patients receiving corticosteroid therapy during COVID-19 treatment are at an elevated risk of developing PCRONJ. Consequently, the administration of corticosteroids in the clinical management of SARS-CoV-2 infections should be carefully restricted to severe cases and confined to periods of acute respiratory distress to mitigate potential long-term complications [15].

Conclusion

In conclusion, tocilizumab and corticosteroids have demonstrated efficacy in managing severe COVID-19 cases, particularly by mitigating cytokine storms and inflammatory complications. However, their usage is associated with potential adverse effects, including increased risk of medication-related osteonecrosis of the jaw (MRONJ). Tocilizumab has shown promise in improving survival outcomes in critical patients, especially those with elevated C-reactive protein levels and in Intensive care unit (ICU), while corticosteroids require cautious administration to minimize complications such as hyperglycemia and bone-related pathologies. These findings underscore the importance of balancing therapeutic benefits with potential risks and tailoring treatment strategies to individual patient profiles for optimal outcomes.

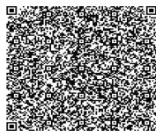
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Immunotherapy: Innovations and Clinical Applications

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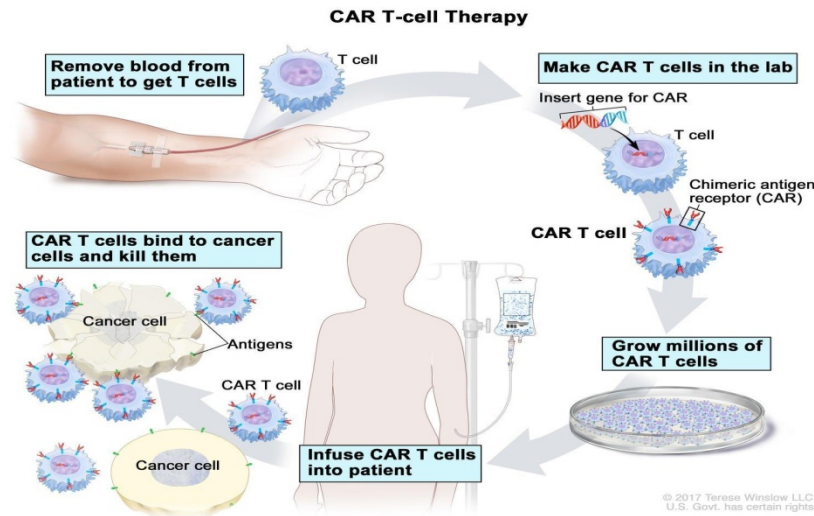
Introduction

The evolution of cancer treatment has been profoundly influenced by the development of immunotherapy, a modality that leverages the body's immune system to combat malignancies. This approach has transitioned from early experimental stages to becoming a cornerstone in oncology, complementing traditional treatments such as chemotherapy and radiotherapy. This chapter delves into the historical context of immunotherapy, underscores the immune system's pivotal role in cancer surveillance, and outlines the paradigm shift towards targeted immunotherapeutic strategies, with a focus on checkpoint inhibitors, CAR-T cell therapy, and personalized immunotherapy.

History of Immunotherapy in Cancer Treatment

- ❖ Dr. William B. Coley, who tends to be referred to as the "Father of Immunotherapy," stipulated tumor regression in patients who got erysipelas, a bacterial skin infection, in the late 19th century, and those laid the philosophical foundation for immunotherapy.
- ❖ Motivated by this evidence, Coley explicitly introduced *Streptococcus pyogenes* into cancer patients, which in certain scenes caused tumor shrinking.
- ❖ Any subsequent studies into using the immune system to combat cancer were made possible by this early type of immunotherapy.
- ❖ As the 20th century went on, emphasis turned to comprehending the elements of the immune system and how they acquainted with cancer cells.

- ❖ A noteworthy turning point was reached in 1986 when the U.S. Food and Drug Administration (FDA) stumbled upon and approved interferon-alpha 2 (IFN- α 2), making cytokine therapy a feasible cancer treatment.



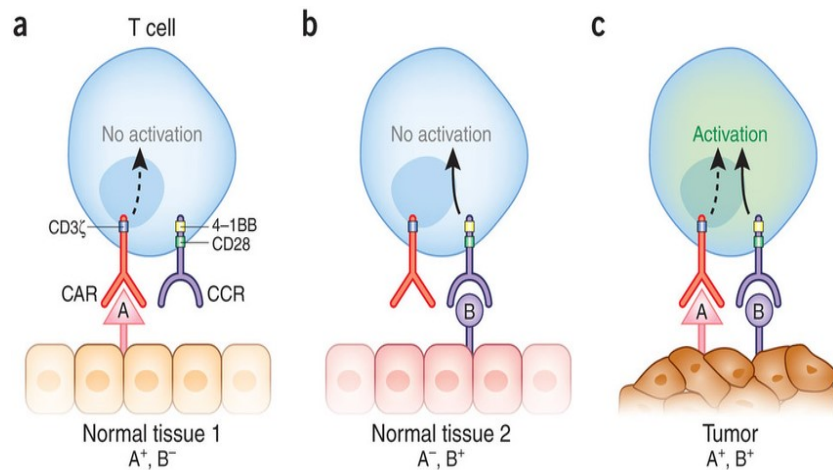
Importance of the Immune System in Recognizing and Destroying Cancer Cells

As the body's defense mechanism, the immune system conceals and gets rid of undesirable cells, particularly when it comes that are changing malignantly. T cells become crucial to this response because they are able to identify tumor-specific antigens that are displayed on the outermost regions of cancer cells. However, malignancies frequently trigger immunosuppressive microenvironments or negatively regulate antigen presentation as ways to circumvent immune identification. Designing treatments that can strengthen the immune response against cancer has required an understanding of these interconnected processes.

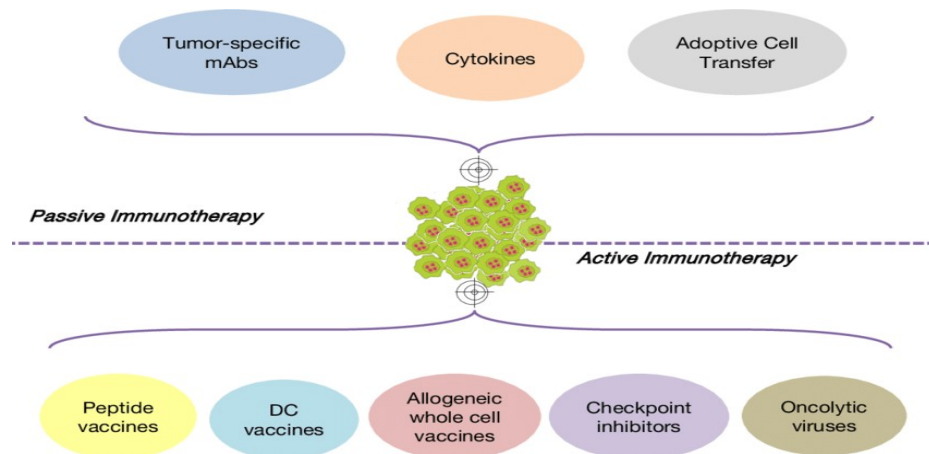
Conditions that were historically challenging to manage with conventional therapies

1. **Advancements in Checkpoint Inhibitors:** Immune checkpoints are regulatory pathways that maintain self-tolerance and modulate the duration and amplitude of physiological immune responses. Tumors often exploit these checkpoints to evade immune surveillance. Checkpoint inhibitors, such as antibodies targeting PD-1/PD-L1 and CTLA-4, have revolutionized cancer therapy by blocking these inhibitory pathways, thereby reinvigorating T cell activity against cancer cells.

2. **CAR-T Cell Therapy Updates:** Chimeric Antigen Receptor (CAR) T cell therapy involves genetically engineering a patient's T cells to express receptors specific to tumor antigens. Upon reinfusion, these modified T cells can effectively target and destroy cancer cells. This approach has shown unprecedented success in treating certain hematologic malignancies, leading to durable remissions.
3. **Personalized Immunotherapy for Cancer:** Recognizing the heterogeneity of tumors and the unique immunological landscape of each patient, personalized immunotherapy tailors treatments based on individual characteristics. This includes developing personalized cancer vaccines and identifying predictive biomarkers to select the most appropriate immunotherapeutic strategies.



Definition and Classification of Immunotherapy Approaches:



Passive Immunotherapy: This approach involves the administration of immune system components, such as monoclonal antibodies or adoptive cell transfers, that are directly active against cancer cells. These agents do not rely on the patient's immune system to initiate a response but instead provide immediate, albeit temporary, anti-tumor effects.

Active Immunotherapy: Active strategies aim to stimulate the patient's own immune system to recognize and combat cancer cells. This includes cancer vaccines and immune checkpoint inhibitors that enhance the body's natural defenses to achieve a sustained anti-tumor response.

Immunotherapy Modalities

- 1. Monoclonal Antibodies (mAbs):** These laboratory-produced molecules can bind to specific antigens expressed on cancer cells, marking them for destruction by the immune system. Some mAbs are designed to block growth signals or deliver cytotoxic agents directly to tumor cells.
- 2. Immune Checkpoint Inhibitors:** These agents target regulatory pathways in T cells to prevent cancer-induced immune suppression. By inhibiting proteins such as PD-1, PD-L1, or CTLA-4, checkpoint inhibitors reactivate T cells, enabling them to effectively attack tumor cells.
- 3. Cancer Vaccines:** Designed to elicit an immune response against specific tumor-associated antigens, cancer vaccines can be prophylactic or therapeutic. They work by presenting these antigens to the immune system, prompting the activation of T cells that target cancer cells expressing the antigens.
- 4. Adoptive Cell Therapy (ACT):** This method involves extracting immune cells from a patient, enhancing their cancer-fighting capabilities in vitro, and reinfusing them into the patient. Chimeric Antigen Receptor (CAR) T-cell therapy is a prominent example, where T cells are genetically engineered to target specific cancer antigens.

Differences Between Immunotherapy and Traditional Treatment Modalities

Aspect	Traditional Treatment (Chemotherapy/Radiotherapy)	Immunotherapy
Mechanism	Destroys rapidly dividing cells, including cancerous and some healthy cells.	Targets unique antigens on cancer cells.

Emerging Research Concept in Life Sciences

Specificity	Low specificity, leading to significant side effects.	High specificity, minimizing damage to normal tissues.
Durability	Treatment effect may not be long-lasting.	Induces long-lasting immune responses for sustained cancer surveillance.
Adaptability	Less adaptive to changes in cancer cells.	Immune system evolves to recognize and combat tumor heterogeneity.
Effectiveness	Variable; depends on the type of cancer and individual patient factors.	May not be universally effective; influenced by tumor microenvironment and individual immune system variability.
Ongoing Research	Focused on improving targeting and reducing side effects.	Aims to identify predictive biomarkers for tailored approaches.

Barrier Inhibitors: Development:

Pathway PD-1/PD-L1: T-cells express the inhibitory receptor known as programmed death-1 (PD-1), and cancer cells routinely overexpress PD-L1, its ligand. When PD-1 binds to PD-L1, T-cell activation is hindered, allowing cancer cells to evade immune destruction. As a consequence of its function in immune evasion, this immunological checkpoint is a major target in many malignant tumors.

CTLA-4 Means of transportation: Another inhibitory receptor that is expressed on T-cells, particularly amid the early phases of T-cell activation, is cytotoxic T-lymphocyte-associated antigen-4 (CTLA-4). When it comes to binding to B7 ligands on antigen-presenting cells (APCs), CTLA-4 and the co-stimulatory receptor CD28 compete. This suppresses the immune system, which authorizes malignancies to flourish unchecked.

Approved Checkpoint Inhibitors

1. Anti-PD-1 Inhibitors:

- **Pembrolizumab (Keytruda):** Approved for various cancers, including melanoma, non-small cell lung cancer (NSCLC), and head and neck squamous cell carcinoma.

- **Nivolumab (Opdivo):** Widely used for melanoma, NSCLC, renal cell carcinoma, and Hodgkin's lymphoma.

2. Anti-PD-L1 Inhibitors:

- **Atezolizumab (Tecentriq):** Effective in NSCLC, urothelial carcinoma, and triple-negative breast cancer.
- **Durvalumab (Imfinzi):** Primarily used for NSCLC and bladder cancer.

3. Anti-CTLA-4 Inhibitors:

- **Ipilimumab (Yervoy):** The first checkpoint inhibitor approved for melanoma, often combined with nivolumab for synergistic effects.

Initialization Checkpoint Objectives

- **LAG-3 (Lymphocyte Activation Gene-3):** Regulates T-cell function and is being investigated in combination with PD-1 inhibitors.
- **TIM-3 (T-cell Immunoglobulin and Mucin Domain-3):** Expressed on exhausted T-cells, TIM-3 inhibitors are in early-phase clinical trials.
- **TIGIT (T-cell Immunoreceptor with Ig and ITIM Domains):** Emerging as a promising target to enhance T-cell and NK cell activity.

Principles of CAR-T Cell Therapy

CAR-T (Chimeric Antigen Receptor T-cell) therapy is a revolutionary immunotherapy approach that involves genetically engineering a patient's T-cells to target and destroy specific cancer cells. The process integrates the following key steps:

1. **T-cell Collection:** T-cells are extracted from the patient's blood through a process called leukapheresis.
2. **Genetic Modification:** The collected T-cells are genetically engineered to express chimeric antigen receptors (CARs) on their surface. These receptors are designed to recognize specific antigens present on cancer cells.
3. **Expansion:** The modified T-cells are expanded in a laboratory to increase their number.
4. **Reinfusion:** The engineered T-cells are infused back into the patient's bloodstream, where they seek out and destroy cancer cells.

Approved CAR-T Therapies

1. **CD19-directed CAR-T Therapies:**

- *Tisagenlecleucel* (Kymriah): Approved for treating relapsed/refractory B-cell acute lymphoblastic leukemia (ALL) and certain types of large B-cell lymphoma.
- *Axicabtagene Ciloleucel* (Yescarta): Approved for diffuse large B-cell lymphoma (DLBCL), primary mediastinal large B-cell lymphoma, and transformed follicular lymphoma.

2. **Advances in CAR-T for Solid Tumors:** Despite significant success in hematological cancers, CAR-T therapy's application in solid tumors faces challenges, including tumor microenvironment barriers and antigen heterogeneity. Recent advances focus on improving CAR designs to target solid tumors effectively.

Innovations in CAR-T Technology

1. **Fourth-generation CAR-T Cells (Armored CAR-T):** These CAR-T cells are engineered to secrete cytokines or other molecules to enhance their anti-tumor activity and resistance to immunosuppression.
2. **Dual-antigen Targeting CARs:** This approach reduces the risk of tumor escape by targeting two different antigens simultaneously, thereby increasing therapeutic efficacy.
3. **Universal CAR-T Therapies:** Allogeneic (off-the-shelf) CAR-T cells are being developed to eliminate the need for patient-specific T-cell harvesting and modification, reducing costs and improving accessibility.
4. **Overcoming Challenges in Solid Tumor Microenvironments:** Strategies like incorporating cytokine release inhibitors, targeting stromal components, and enhancing CAR-T cell persistence are being explored to tackle the immunosuppressive microenvironment in solid tumors.

Clinical Trials and Outcomes

1. **Hematological Malignancies:** CAR-T therapies have shown remarkable success in treating blood cancers, with durable remission rates in diseases like ALL and DLBCL.
2. **Solid Tumors:** Emerging clinical trials are investigating the efficacy of CAR-T therapy in challenging cancers, such as glioblastoma and

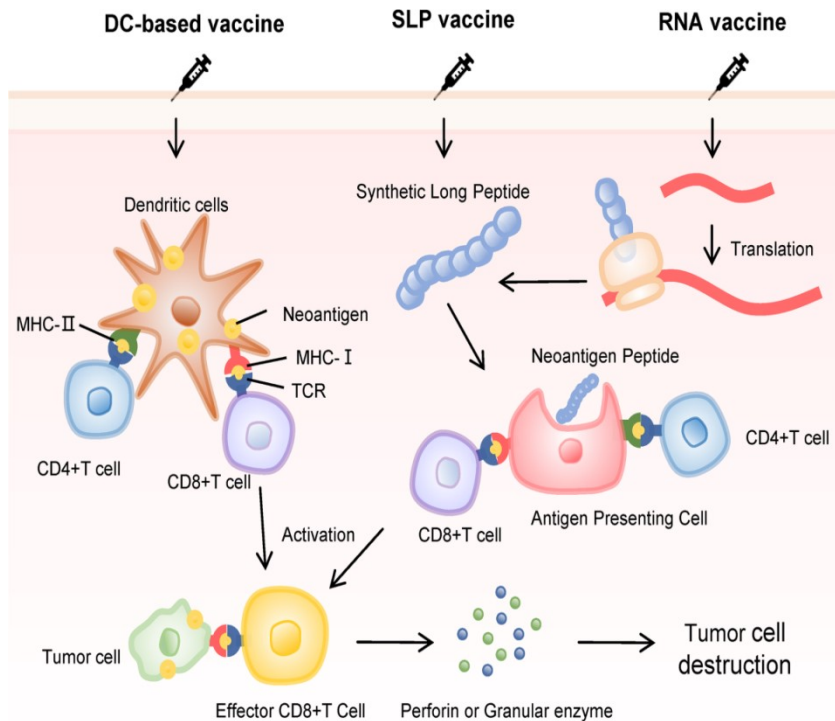
mesothelioma. These studies aim to expand the scope of CAR-T applications beyond hematologic malignancies.

Personalized Immunotherapy for Cancer

- ❖ Cancer treatment has entered an era where personalization has become critical for achieving optimal therapeutic outcomes.
- ❖ Tumor heterogeneity—where cancer cells within the same tumor exhibit significant genetic, epigenetic, and phenotypic differences—necessitates individualized approaches.
- ❖ Additionally, unique immune profiles of patients influence how their immune systems respond to therapies. This rationale underscores the need for tailored immunotherapy strategies to maximize efficacy while minimizing adverse effects.
- ❖ Personalization ensures that treatments align with a patient's specific tumor characteristics and immune microenvironment.

Neoantigen-Based Vaccines

- ✓ Neoantigens, which are tumor-specific antigens arising from somatic mutations, are gaining traction as promising targets for personalized immunotherapy.
- ✓ Advances in bioinformatics have enabled the identification of neoantigens through genomic and transcriptomic analyses. By leveraging next-generation sequencing and machine learning algorithms, researchers can predict which neoantigens are most likely to elicit an immune response.
- ✓ Early successes have been reported in melanoma and non-small-cell lung cancer (NSCLC), where neoantigen-based vaccines have demonstrated encouraging results in clinical trials.



TIL Therapy (Tumor-Infiltrating Lymphocytes)

- ✓ Tumor-Infiltrating Lymphocyte (TIL) therapy is another personalized approach that involves isolating TILs from a patient's tumor, expanding them ex vivo, and reinfusing them back into the patient.
- ✓ TILs are naturally occurring immune cells that have infiltrated the tumor and exhibit tumor-specific cytotoxicity. Recent breakthroughs in TIL therapy include the FDA's approval of therapies targeting advanced melanoma.
- ✓ Researchers are also exploring ways to improve TIL efficacy, such as combining TIL therapy with checkpoint inhibitors or using genetic modifications to enhance their functionality.

Role of Biomarkers in Personalization

- Predictive biomarkers are pivotal for tailoring immunotherapy to individual patients. For example, PD-L1 expression levels can predict the efficacy of checkpoint inhibitors like pembrolizumab and nivolumab.

- Microsatellite instability-high (MSI-H) and tumor mutational burden (TMB) are additional biomarkers that have been linked to better responses to immunotherapy.

Clinical Applications and Real-World Impact:

Success Stories in Immunotherapy: Numerous examples highlight the transformative power of immunotherapy. Patients with advanced melanoma, such as those treated with checkpoint inhibitors like nivolumab and pembrolizumab, have achieved complete remission with survival extending well beyond historical averages.

CAR-T cell therapies, including tisagenlecleucel, have demonstrated remarkable success in treating refractory B-cell malignancies, with long-term remission observed in pediatric acute lymphoblastic leukemia (ALL) and adult diffuse large B-cell lymphoma (DLBCL). Furthermore, immunotherapy's scope has expanded beyond oncology, addressing conditions like autoimmune disorders (e.g., rheumatoid arthritis) and chronic infectious diseases (e.g., HIV).

Challenges and Limitations of Current Approaches

1. **Addressing Resistance Mechanisms:** Resistance to checkpoint inhibitors and CAR-T therapy remains a formidable barrier. Tumor cells adapt by downregulating antigen presentation, upregulating alternative checkpoint pathways, or creating immunosuppressive microenvironments. Research into combination therapies, targeting multiple pathways simultaneously, offers a potential solution.
2. **Management of Side Effects:** Immune-related adverse events (irAEs), such as colitis, pneumonitis, and cytokine release syndrome (CRS), pose significant risks to patients. Early recognition, prompt management with immunosuppressive agents, and predictive biomarker development are crucial to improving tolerability and outcomes.
3. **Scalability and Cost Challenges:** The scalability of CAR-T therapies and other immunotherapies is hindered by the complexity of their production and high costs. Innovations in manufacturing processes, such as automation and the use of allogeneic cell sources, are being explored to address these issues.

Future Perspectives and Innovations:

Role of Nanotechnology: Nanotechnology is emerging as a powerful tool to enhance the delivery and targeting of immunotherapeutic agents. Nanoparticles

can be designed to deliver drugs directly to tumors, minimize off-target effects, and modulate the tumor microenvironment to enhance immune responses.

Advances in Combination Therapies: Combining immunotherapy with oncolytic viruses, chemotherapy, or radiotherapy holds promise for overcoming resistance and improving patient outcomes. Oncolytic viruses, for instance, can selectively infect and lyse tumor cells while enhancing immune system activation.

Conclusion

Immunotherapy has furnished humanity all across the world new hope by elevating the parameters of cancer treatment. It has raised awareness about long-term remissions and, in specific circumstances, possible cures for tumors that were previously intractable by utilizing the body's immune system. The continual progression of immunotherapeutic approaches and their inclusion of contemporary technology highlight its disruptive potential in spite of its drawbacks. Putting forth efforts to tackling equity and accessibility is equally important. Governments, non-profits, and the pharmaceutical sector must work together to ensure that immunotherapy reaches a variety of neighborhoods around the world.

Immunotherapy is still at the forefront of oncology; with potential applications beyond cancer into various other therapeutic areas as research and innovation continue to open up new avenues. The journey that lies ahead holds potential and calls for tenacity, ingenuity, and solidarity in order to be accomplished.


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Factors Affecting Bioremediation and Its Types

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Introduction

Bioremediation is a biological process that uses living organisms, usually microorganisms (bacteria and fungi) and plants, to degrade, remove, alter, immobilize, and detoxify waste products and pollutants from soil or water. Microorganisms secrete a militia of enzymes and cofactors that act as natural biocatalysts and facilitate the progress of biochemical pathways involved in the degradation of pesticides. Some plant species are also able to degrade pollutants and can remove contaminants from the soil or water by absorbing them via the roots and then accumulating them in the leaves. Therefore, this process is often considered as a cost-effective and sustainable method for environmental clean-up applications. Being a natural and ecofriendly process based on natural attenuation, bioremediation is highly appreciated than chemicals and physical technologies. The salient features of bioremediation are as follows:

- The majority of bioremediation treatment methods eliminate pollutants present in the soil.
- Bioremediation procedures are intended to either destroy or reduce the harmful organic compounds into less toxic and simpler molecules.
- Bioremediation major relies on indigenous microorganisms, such as bacteria and fungus. Occasionally, some plant species are used to improve biodegradation and stabilize the soils.
- In certain cases, the supply of nutrients or electron acceptors (such as ozone or hydrogen peroxide) to promote the development and reproduction of local species is necessary.

Bioremediation is a branch of biotechnology that employs the use of living organisms in the removal of contaminants, pollutants, and toxins from soil, water, and other environments. These organisms can include microbes and bacteria.

Bioremediation can be used to clean up contaminated groundwater or environmental problems, such as oil spills. Bioremediation is a biotechnical process, which abates or cleans up contamination. It is a type of waste management technique which involves the use of organisms to remove or utilize the pollutants from a polluted area.

There are several remedies where contaminated water or solid is purified by chemical treatment, incineration, and burial in a landfill. There are other types of waste management technique which include solid waste management, nuclear waste management, etc. Bioremediation is different as it uses no toxic chemicals.

Microorganisms like Bacteria and Fungi are the main role player when it comes to executing the process of Bioremediation. Bacteria are the most crucial microbes in this process as they break down the waste into nutrients and organic matter. Even though this is an efficient process of waste management but bioremediation cannot destroy 100% contaminants. Bacteria can easily digest contaminants like chlorinated pesticides or clean oil spills but microorganisms fail to destroy heavy metals like lead and cadmium.

Types of Bioremediation

Bioremediation is of three types

1. Biostimulation

In biostimulation, the bacteria are stimulated to initiate the process. The contaminated soil is first mixed with special nutrients substances including other vital components either in the form of liquid or gas. It stimulates the growth of microbes thus resulting in efficient and quick removal of contaminants by microbes and other bacteria.

2. Bioaugmentation

At times, there are certain sites where microorganisms are required to extract the contaminants, For example – municipal wastewater. In these special cases, the process of bioaugmentation is used. There's only one major drawback in this process. It almost becomes impossible to control the growth of microorganisms in the process of removing the particular contaminant.

3. Intrinsic Bioremediation

The process of intrinsic bioremediation is most effective in the soil and water because of these two biomes which always have a high probability of being full of contaminants and toxins. The process of intrinsic bioremediation is mostly used in underground places like underground petroleum tanks. In such place, it is difficult to detect a leakage and contaminants and toxins can find their way to enter through these leaks and contaminate the petrol. Thus, only microorganisms can remove the toxins and clean the tanks.

Principles of bioremediation

Environmental biotechnology is not a new field; composting and wastewater treatments are familiar examples of old environmental biotechnologies. However, recent studies in molecular biology and ecology offer opportunities for more efficient biological processes. Notable accomplishments of these studies include the clean-up of polluted water and land areas. Bioremediation is defined as the process whereby organic wastes are biologically degraded under controlled conditions to an innocuous state, or to levels below concentration limits established by regulatory authorities (Mueller *et al.*, 1996). By definition, bioremediation is the use of living organisms, primarily microorganisms, to degrade the environmental contaminants into less toxic forms. It uses naturally occurring bacteria and fungi or plants to degrade or detoxify substances hazardous to human health and/or the environment. The microorganisms may be indigenous to a contaminated area or they may be isolated from elsewhere and brought to the contaminated site (Vidali, 2001).

Contaminant compounds are transformed by living organisms through reactions that take place as a part of their metabolic processes. Biodegradation of a compound is often a result of the actions of multiple organisms. When microorganisms are imported to a contaminated site to enhance degradation we have a process known as bioaugmentation. For bioremediation to be effective, microorganisms must enzymatically attack the pollutants and convert them to harmless products.

As bioremediation can be effective only where environmental conditions permit microbial growth and activity, its application often involves the manipulation of environmental parameters to allow microbial growth and degradation to proceed at a faster rate. Like other technologies, bioremediation has its limitations. Some contaminants, such as chlorinated organic or high aromatic hydrocarbons, are resistant to microbial attack. They are degraded

either slowly or not at all, hence it is not easy to predict the rates of clean-up for a bioremediation exercise; there are no rules to predict if a contaminant can be degraded.

Bioremediation techniques are typically more economical than traditional methods such as incineration, and some pollutants can be treated onsite, thus reducing exposure risks for clean-up personnel, or potentially wider exposure as a result of transportation accidents. Since bioremediation is based on natural attenuation the public considers it more acceptable than other technologies. Most bioremediation systems are run under aerobic conditions, but running a system under anaerobic conditions may permit microbial organisms to degrade otherwise recalcitrant molecules.

Factors affecting bioremediation

The control and optimization of bioremediation processes is a complex system of many factors. These factors include: the existence of a microbial population capable of degrading the pollutants; the availability of contaminants to the microbial population; the environment factors (type of soil, temperature, pH, the presence of oxygen or other electron acceptors, and nutrients)

Aerobic: In the presence of oxygen. Examples of aerobic bacteria recognized for their degradative abilities are *Pseudomonas*, *Alcaligenes*, *Sphingomonas*, *Rhodococcus*, and *Mycobacterium*. These microbes have often been reported to degrade pesticides and hydrocarbons, both alkanes and polyaromatic compounds. Many of these bacteria use the contaminant as the sole source of carbon and energy.

Anaerobic: In the absence of oxygen. Anaerobic bacteria are not as frequently used as aerobic bacteria. There is an increasing interest in anaerobic bacteria used for bioremediation of polychlorinated biphenyls (PCBs) in river sediments, dechlorination of the solvent trichloroethylene (TCE), and chloroform.

Ligninolytic fungi: Fungi such as the white rot fungus *Phanaerochaete chrysosporium* have the ability to degrade an extremely diverse range of persistent or toxic environmental pollutants. Common substrates used include straw, saw dust, or corn cobs.

Methylophiles: Aerobic bacteria that grow utilizing methane for carbon and energy. The initial enzyme in the pathway for aerobic degradation, methane monooxygenase, has a broad substrate range and is active against a wide range

of compounds, including the chlorinated aliphatics trichloroethylene and 1,2-dichloroethane (Allard and Neilson, 1997). For degradation it is necessary that bacteria and the contaminants be in contact. This is not easily achieved, as neither the microbes nor contaminants are uniformly spread in the soil. Some bacteria are mobile and exhibit a chemotactic response, sensing the contaminant and moving toward it. Other microbes such as fungi grow in a filamentous form toward the contaminant. It is possible to enhance the mobilization of the contaminant utilizing some surfactants such as sodium dodecyl sulphate (SDS).

In situ bioremediation

These techniques are generally the most desirable options due to lower cost and less disturbance since they provide the treatment in place avoiding excavation and transport of contaminants. In situ treatment is limited by the depth of the soil that can be effectively treated. In many soils effective oxygen diffusion for desirable rates of bioremediation extend to a range of only a few centimeters to about 30 cm into the soil, although depths of 60 cm and greater have been effectively treated in some cases (Vidali, 2001).

The most important land treatments

Bioventing

Bioventing is the most common in situ treatment and involves supplying air and nutrients through wells to contaminated soil to stimulate the indigenous bacteria. Bioventing employs low air flow rates and provides only the amount of oxygen necessary for the biodegradation while minimizing volatilization and release of contaminants to the atmosphere. It works for simple hydrocarbons and can be used where the contamination is deep under the surface. In situ biodegradation involves supplying oxygen and nutrients by circulating aqueous solutions through contaminated soils to stimulate naturally occurring bacteria to degrade organic contaminants. It can be used for soil and groundwater. Generally, this technique includes conditions such as the infiltration of water-containing nutrients and oxygen or other electron acceptors for groundwater treatment (Vidali, 2001).

Biosparging

Biosparging involves the injection of air under pressure below the water table to increase groundwater oxygen concentrations and enhance the rate of biological degradation of contaminants by naturally occurring bacteria. Biosparging increases the mixing in the saturated zone and thereby increases

the contact between soil and groundwater. The ease and low cost of installing small-diameter air injection points allows considerable flexibility in the design and construction of the system. Bioremediation frequently involves the addition of microorganisms indigenous or exogenous to the contaminated sites. Two factors limit the use of added microbial cultures in a land treatment unit:

1. Nonindigenous cultures rarely compete well enough with an indigenous population to develop and sustain useful population levels and
2. Most soils with long-term exposure to biodegradable waste have indigenous microorganisms that are effective degraders if the land treatment unit is well managed.

Ex situ bioremediation

These techniques involve the excavation or removal of contaminated soil from ground. Landfarming is a simple technique in which contaminated soil is excavated and spread over a pre-prepared bed and periodically tilled until pollutants are degraded. The goal is to stimulate indigenous biodegradative microorganisms and facilitate their aerobic degradation of contaminants. In general, the practice is limited to the treatment of superficial 10-35 cm of soil. Since landfarming has the potential to reduce monitoring and maintenance costs, as well as clean-up liabilities, it has received much attention as a disposal alternative. Composting is a technique that involves combining contaminated soil with nonhazardous organic amendments such as manure or agricultural wastes. The presence of these organic materials supports the development of a rich microbial population and elevated temperature characteristic of composting.

Biopiles are a hybrid of landfarming and composting. Essentially, engineered cells are constructed as aerated composted piles. Typically used for treatment of surface contamination with petroleum hydrocarbons they are a refined version of landfarming that tend to control physical losses of the contaminants by leaching and volatilization. Biopiles provide a favorable environment for indigenous aerobic and anaerobic microorganisms (Von *et al.*, 1998).

Bioreactors

Slurry reactors or aqueous reactors are used for ex situ treatment of contaminated soil and water pumped up from a contaminated plume. Bioremediation in reactors involves the processing of contaminated solid material (soil, sediment, sludge) or water through an engineered containment system. A slurry bioreactor may be defined as a containment vessel and

apparatus used to create a three-phase (solid, liquid, and gas) mixing condition to increase the bioremediation rate of soil bound and water soluble pollutants as a water slurry of the contaminated soil and biomass (usually indigenous microorganisms) capable of degrading target contaminants. In general, the rate and extent of biodegradation are greater in a bioreactor system than in situ or in solid phase systems because the contained environment is more manageable and hence more controllable and predictable. Despite the advantages of reactor systems, there are some disadvantages. The contaminated soil requires pretreatment (e.g., excavation) or alternatively the contaminant can be stripped from the soil via soil washing or physical extraction (e.g., vacuum extraction) before being placed in a bioreactor.

Advantages of bioremediation

- ❖ Bioremediation is a natural process and is therefore perceived by the public as an acceptable waste treatment process for contaminated material such as soil. Microbes able to degrade the contaminant increase in numbers when the contaminant is present; when the contaminant is degraded, the biodegradative population declines. The residues for the treatment are usually harmless products and include carbon dioxide, water, and cell biomass.
- ❖ Theoretically, bioremediation is useful for the complete destruction of a wide variety of contaminants. Many compounds that are legally considered to be hazardous can be transformed to harmless products. This eliminates the chance of future liability associated with treatment and disposal of contaminated material.
- ❖ Instead of transferring contaminants from one environmental medium to another, for example, from land to water or air, the complete destruction of target pollutants is possible.
- ❖ Bioremediation can often be carried out on site, often without causing a major disruption of normal activities. This also eliminates the need to transport quantities of waste off site and the potential threats to human health and the environment that can arise during transportation.
- ❖ Bioremediation can prove less expensive than other technologies that are used for clean-up of hazardous waste.

Disadvantages of bioremediation

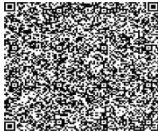
- ❖ Bioremediation is limited to those compounds that are biodegradable. Not all compounds are susceptible to rapid and complete degradation.
- ❖ There are some concerns that the products of biodegradation may be more persistent or toxic than the parent compound.

- ❖ Biological processes are often highly specific. Important site factors required for success include the presence of metabolically capable microbial populations, suitable environmental growth conditions, and appropriate levels of nutrients and contaminants.
- ❖ It is difficult to extrapolate from bench and pilot-scale studies to full-scale field operations.
- ❖ Research is needed to develop and engineer bioremediation technologies that are appropriate for sites with complex mixtures of contaminants that are not evenly dispersed in the environment. Contaminants may be present as solids, liquids, and gases.
- ❖ Bioremediation often takes longer than other treatment options, such as excavation and removal of soil or incineration.
- ❖ Regulatory uncertainty remains regarding acceptable performance criteria for bioremediation. There is no accepted definition of clean, evaluating performance of bioremediation is difficult, and there are no acceptable endpoints for bioremediation treatments.

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"A novel ray on the disease's path Recognition, and Rehabilitation of Myasthenia Gravis"

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Introduction

Myasthenia gravis (MG) is a chronic autoimmune disorder that disrupts normal communication between nerves and muscles, leading to fluctuating muscle weakness and fatigability. This condition occurs due to the production of autoantibodies that attack and block key components of the neuromuscular junction, the site where nerve cells transmit signals to muscle cells. The most commonly targeted component is the acetylcholine receptor (AChR), although in some cases, antibodies attack muscle-specific kinase (MuSK) or other related proteins. These antibodies interfere with the binding of acetylcholine, a neurotransmitter critical for muscle contraction, and trigger the destruction of these receptors, impairing neuromuscular transmission.

Clinically, MG presents with hallmark symptoms such as ptosis (drooping eyelids), diplopia (double vision), dysphagia (difficulty swallowing), and generalized muscle weakness. The muscle weakness is typically activity-dependent, meaning it worsens with exertion and improves with rest, a phenomenon known as fatigability. In severe cases, MG can affect respiratory muscles, leading to a life-threatening condition called myasthenic crisis, which requires immediate medical intervention.

The disorder affects individuals across all age groups but shows a bimodal distribution: it commonly affects younger women and older men.

Diagnosis of MG involves a combination of clinical evaluation, antibody testing, electrophysiological studies such as repetitive nerve stimulation, and imaging to detect thymic abnormalities, which are often associated with the disease.

Treatment strategies focus on improving neuromuscular transmission and suppressing the autoimmune response. Acetylcholinesterase inhibitors, such as pyridostigmine, are commonly used to enhance communication at the neuromuscular junction. Immunosuppressive therapies, including corticosteroids and drugs like azathioprine, help reduce antibody production. In some cases, thymectomy (surgical removal of the thymus gland) is performed, particularly when a thymoma is present. Although MG is a chronic condition, advances in treatment have significantly improved patient outcomes, allowing many individuals to lead relatively normal lives. Ongoing research into the pathophysiology of MG aims to develop more targeted therapies and improve the quality of life for those affected.

Historical Perspective and Discovery:

The earliest known description of myasthenia gravis dates back to 1672 by Oxford physician Thomas Willis, who documented cases of patients with fluctuating muscle weakness. However, it wasn't until 1877 that Samuel Wilks provided a modern clinical description of the disease. A significant advancement in MG treatment occurred in 1934 when Dr. Mary Broadfoot Walker discovered the therapeutic efficacy of physostigmine, an acetylcholinesterase inhibitor, marking a pivotal moment in the management of the disease.

Pathophysiology of Myasthenia Gravis:

1. Autoimmune Mechanisms

- ☐ The immune system mistakenly targets self-antigens.
- ☐ In MG, autoantibodies are produced against components of the neuromuscular junction.

2. Role of Acetylcholine Receptor (AChR) Antibodies

- ☐ Autoantibodies bind to AChRs on the postsynaptic muscle membrane.
- ☐ This binding leads to:

1. Complement-mediated damage to the muscle membrane.
2. Accelerated degradation of AChRs.
3. Functional blockade of the receptors.

☐ The result is impaired neuromuscular transmission, causing muscle weakness and fatigability.

3. Thymus and Its Role in Myasthenia Gravis:

- ☐ The thymus is involved in T-cell development and tolerance.
- 1. Hyperplasia (enlargement) with germinal centers.
- 2. Presence of myoid cells expressing AChR antigens.
- ☐ These abnormalities may contribute to the breakdown of self-tolerance, leading to the production of AChR antibodies.

Clinical Features of Myasthenia Gravis (MG):

1. Common Symptoms and Presentation

Initial Symptoms: Approximately two-thirds of individuals present with ocular symptoms, such as ptosis (drooping eyelids) and diplopia (double vision).

Progression: Symptoms may remain limited to ocular muscles (ocular MG) or progress to involve other muscle groups (generalized MG).

2. Muscle Weakness and Fatigability

Nature of Weakness: Characterized by painless weakness that worsens with activity and improves with rest.

Affected Muscle Groups: May involve ocular, bulbar, respiratory, and limb muscles.

Prevalence and Incidence of Myasthenia Gravis (MG)

Incidence: MG incidence ranges from 4.1 to 30 cases per million person-years globally, with U.S. rates recently increasing to 23.81–50.67 per million, influenced by age, sex, and race.

Prevalence: Global prevalence is estimated at 150–200 cases per million, while in the U.S., it is 14–40 cases per 100,000 individuals. Rising prevalence reflects improved diagnostics, greater awareness, and better survival outcomes.

Ocular and Bulbar Involvement

Ocular Involvement

1. Ptosis: Drooping of one or both eyelids.

2. Diplopia: Double vision due to weakness of the extraocular muscles.

Bulbar Involvement

1. Dysarthria: Slurred or nasal speech resulting from weakness in the muscles involved in speaking.

2. Dysphagia: Difficulty swallowing due to weakness in the muscles involved in chewing and swallowing.

Diagnosis of Myasthenia Gravis (MG)

1. Clinical Examination and History

Patient History

1. Assess for symptoms such as fluctuating muscle weakness and fatigability.
2. Observe distribution of muscle weakness, particularly ocular and bulbar muscles.

Physical Examination

1. Evaluate for signs like ptosis, diplopia, and muscle weakness that worsens with exertion.
2. Perform bedside tests, such as the ice pack test, to observe improvement in ptosis.

2. Laboratory Investigations

Anti-Acetylcholine Receptor (AChR) Antibody Test: Detects the presence of antibodies against AChR, present in approximately 80-85% of MG patients.

Anti-Muscle-Specific Kinase (MuSK) Antibody Test: Identifies antibodies against MuSK, found in about 5-8% of MG patients, particularly those who are AChR antibody-negative.

Anti-LRP4 Antibody Test: Assesses for antibodies against LRP4, present in a subset of seronegative MG patients.

3. Electrophysiological Studies

Repetitive Nerve Stimulation (RNS): Measures the decremental response in muscle action potentials upon repetitive stimulation, indicative of neuromuscular transmission defects.

Single-Fiber Electromyography (SFEMG): Evaluates neuromuscular jitter and blocking, highly sensitive for detecting abnormalities in neuromuscular transmission.

4. Imaging Techniques

Computed Tomography (CT) Scan: Assesses the anterior mediastinum for thymic abnormalities, such as thymoma or thymic hyperplasia.

Magnetic Resonance Imaging (MRI): Provides detailed imaging of the thymus and may help in distinguishing thymic tissue characteristics.

Management of Myasthenia Gravis (MG)

1. Pharmacological Treatments

Acetylcholinesterase Inhibitors: Medications like pyridostigmine enhance neuromuscular transmission by inhibiting the breakdown of acetylcholine, providing symptomatic relief.

Immunosuppressive Drugs: Agents such as corticosteroids, azathioprine, and mycophenolate mofetil suppress the immune response, reducing antibody production against acetylcholine receptors.

2. Surgical Interventions

Thymectomy: Surgical removal of the thymus gland can be beneficial, especially in patients with thymoma or generalized MG, potentially leading to remission or reduced medication dependency.

3. Plasmapheresis and Intravenous Immunoglobulin (IVIg) Therapy

Plasmapheresis: A procedure that filters antibodies from the blood, providing short-term improvement in muscle strength, often used in acute exacerbations.

IVIg Therapy: Administration of intravenous immunoglobulins modulates the immune system, offering temporary relief, particularly during myasthenic crises or preoperatively.

4. Management of Myasthenic Crisis

□ A myasthenic crisis, characterized by respiratory failure due to severe muscle weakness, requires prompt intensive care management, including ventilatory support and rapid immunomodulatory treatments like plasmapheresis or IVIg.

Prognosis and Long-term Care

1. Disease Course and Variability

□ MG exhibits a variable course, with some patients achieving remission while others experience persistent symptoms. Early and appropriate treatment has significantly improved outcomes.

2. Impact of Treatment on Quality of Life

□ Effective management of MG symptoms through pharmacological and surgical interventions has been shown to enhance daily functioning and overall quality of life.

3. Monitoring and Follow-up

□ Regular follow-up is essential to adjust treatments, monitor for side effects, and assess disease progression, ensuring optimal patient outcomes.

Conclusion

Myasthenia Gravis (MG) is an autoimmune neuromuscular disorder characterized by fluctuating muscle weakness and fatigability. The pathogenesis involves autoantibodies targeting components of the neuromuscular junction, notably the acetylcholine receptor (AChR), leading to impaired synaptic transmission. Clinical manifestations often include ptosis, diplopia, dysphagia, and generalized muscle weakness. Diagnosis is established through a combination of clinical evaluation, serological testing for specific antibodies, electrophysiological studies, and imaging to assess for thymic abnormalities. Management strategies encompass pharmacological treatments such as acetylcholinesterase inhibitors and immunosuppressive agents, surgical interventions like thymectomy, and therapeutic modalities including plasmapheresis and intravenous immunoglobulin therapy. The disease course is variable; however, with appropriate treatment, many patients experience significant improvement in symptoms and quality of life.

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Halophilic actinomycetes (*Streptomyces sp*) involved in antimicrobial resistance and its various applications

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Introduction

Halophilic Actinomycetes can be found in a variety of hypersaline environments. Hypersaline environments are salt pans, salt lakes, marine environments, oceans, seas, salt foods and saline soils (1). Saltern is the extreme habitat that promotes halophilic actinomycetes metabolic adaption to saline conditions. Hypersaline environments with high salt concentrations can be found throughout the salt pans. The saltpan is formed naturally by the evaporation of seawater (2). Halophiles are salt-loving microorganisms and can only survive at high salt concentrations, and they can be classified according to how much salt they need to thrive. Halophiles are divided into three categories: slight halophiles (1.7-4.8%), moderate halophiles (4.7-20) and extreme halophiles (20 - 30%) of NaCl.



Fig.1. Halophilic actinomycetes

Antimicrobial resistance

Antimicrobial resistance is one of the emerging public health problems in every day. Some pathogenic clinical bacteria are capable of causing diseases. *Staphylococcus aureus* is a frequent bacterial pathogen and it causes various infections. Pathogenic microbes such as *Enterococcus faecium*, *Enterococcus faecalis*, *Salmonella enterica*, *Staphylococcus aureus*, and *Streptococcus pneumoniae* have developed resistance to the existing antibiotics used for the microbial disease (3).

World Health Organization (WHO) published a report in 2017 on bacterial infections and human health, recommending the development of new antibiotics against the resistant developing microorganisms (4). The rapid spread of antibiotic resistance has increased research interest in the discovery of natural products, mainly from actinomycetes, which have been the primary source of antimicrobial compounds.

Novel antibiotics are being developed using natural derivative compounds. From 1981 to 2014, the Food and Drug Administration (FDA) approved a total of 1211 small molecule medications derived from natural substances (5). Totally 41 novel medicines in clinical trials around the world until April 2020, but only 13 of them have the potential to cure diseases on the WHO's critical threat list. As a result of this situation, there is currently an acute shortage of novel antibiotics and drug leads. Certain techniques, such as antibiotic modifying enzymes, antibiotic degrading enzymes, and gene transfer processes like conjugation, transformation and transduction can be used to develop antibiotics. As a result, there's a demand for naturally occurring new antibacterial compounds (6).

Actinomycetes from saline environments having lot of potential applications, including production of antibacterial agents. In this study H8 isolated from saltpan soil sample has been identified to be *Streptomyces* sp. The Strain shows broad-spectrum activity against bacteria including antibiotic-resistant bacterial pathogens *Klebsiella Pneumoniae*, *Escherchia Coli*, *Acinetobacter baumannii* and *Pseudomonas aeruginosa*. The metabolite also showed the better anticancer activity against Colon Cancer Cell line (7).

Multi drug resistant

Multidrug resistance (MDR) is a type of insensitivity developed by microbes to lethal doses of antibiotics. MDR has become a major concern for antibiotics regarding their efficacy against pathogenic disease (8). In the present scenario, resistance to antibiotics is one of the crucial issues related to

public health. Earlier, such resistance to antibiotics was limited to nosocomial infections, but it has now become a common phenomenon. Several factors, like extensive development, overexploitation of antibiotics, excessive application of broad-spectrum drugs, and a shortage of target-oriented antimicrobial drugs, could be attributed to this condition. Nowadays, there is a rise in the occurrence of these drug-resistant pathogens due to the availability.

Multidrug-resistant bacteria (MDR) that are deadly pathogenic are rising day by day and pose a very serious threat to human health. Earlier, these types of antibiotic-resistant bacterial strains were rare and limited to only nosocomial-acquired infections, but nowadays, they have become very common. This issue is more prevalent among both Gram-positive and Gram-negative bacterial species, which include *A. baumannii*, *E. coli*, *P. aeruginosa*, and *K. pneumonia* (Gram-negative), along with *S. aureus*, *S. pneumoniae*, *E. faecium*, and *E. faecalis* (Gram-positive). (B. Jubeh *et al.*, 2020)

Important features of *streptomyces sp*

Actinomycetes, particularly those of the genus *Streptomyces*, can create a large number of secondary metabolites, including antibiotics, as biologically active compounds. The majority of antibiotics that are currently in use are derivatives of actinomycetes and fungus natural product. *Streptomyces* is a Gram Positive bacterial genus that grows in a filamentous form, related to fungi and thrives in a variety of environments. *Streptomyces* morphological characterization requires the formation of a layer of hyphae capable of differentiating into a spore chain. This is a unique Gram-positive mechanism that necessitates a well-coordinated and specialized metabolism. Another distinguishing feature is a member of the genus is complicated multicellular growth (1). In which hyphae spores germinate with multinuclear aerial mycelium, which produces septa at regular intervals, forming a chain of uninucleated spores.

The potential of *Streptomyces* to synthesize Antifungals, antivirals, antitumors, and anti-hypertensives are examples of bioactive secondary metabolites is its most intriguing feature. *Streptomyces* have become an economically significant category of organisms among actinomycetes, and they are a significant source of a wide range of biologically active substances (2). However, current study has discovered that several taxa of actinomycetes are either natural or well-adapted in marine residents. Antibiotics, anticancer chemicals, immunosuppressants, antiviral, antiparasitic, and enzyme-inactivating substances are all derived from microorganisms (3). Only 150 was approximately 23,000 bioactive secondary metabolites are synthesized by

microbes have been documented for use in pharmacology, agriculture, or other sectors. Actinomycetes synthesize over 10,000 of these chemicals, accounting for 45 percent of all bioactive microbial metabolites identified, and 80 percent if just those molecules in practical application are considered. Streptomyces species produce roughly 7600 chemicals among actinomycetes.(4) Streptomyces producing antibiotics, actinomycetes play an important function inside a marine environment. Its have been associated to degradation and regeneration of organic substances. They also serve an a significant role in the mineralization of organic matter, the immobilization of mineral nutrients, nitrogen fixation, physical parameter improvement, and environmental protection (5)

Marine *Streptomyces* sp have received less attention as probiotics in aquaculture while being the numerous sources of new antibiotics. Furthermore, marine Streptomyces were recommended as prospective candidates for use in marine aquaculture according to their capacity to breakdown starch and protein macromolecules in cultivation pond water, marine Streptomyces are indicated as potential probiotic strains. Antimicrobial compounds are produced heat and desiccation tolerant spores are formed. Some a few researches on the potential Streptomyces from the sea have been used to treat a variety of illnesses. (6)

The pharmaceutical industry's economic future is dependent on the discovery or development of novel compounds with new activities or targeted towards a more specialised market. The number of secondary metabolites is continually increasing combinatorial biosynthesis generates new compounds. Since marine microorganisms, especially actinomycetes, having developed the highest genetic and metabolic diversity, the focus of efforts should be on researching actinomycetes from the marine as a source for the identification the discovery of novel secondary metabolites. (7) Recent tradition research have demonstrated that unusual actinomycetes still exist in large numbers in an ocean environment Studies on the prevalence and spread of marine actinomycetes that are not culture dependent. The information gathered from this research has also been utilised to develop selective isolation strategies that have enabled the separation of a diverse variety of marine actinomycetes (8).

Applications of halophilic actinomycetes

Dye degradation

The Synthetic dyes are extremely used in various industries like textile dyeing, paper printing, colour photographs, pharmaceuticals, food and cosmetics etc. Azodyes are integral to the textile industry and have been shown to damage

ecosystem by textile effluents during dyeing process. Strong and bright colours of the azodyes in the waste water is the serious problem of the textile waste effluent and the release of these effluents in to the water bodies cause damage to the environment. Halophilic actinomycetes now are being recognized for their degradative capacity of highly recalcitrant compounds. Actinomycetes have been shown to specifically degrade hydrocarbons. The effectiveness of microbial decolorization depends on the adaptability and the activity of selected microorganisms. (9)



Fig.2. Dye degradation

Antimicrobial compounds

- The major compounds are N-Hexadecanoic acid, N-Nonadecanol-1, 1-Methyl-1-n-butoxy-1-silacyclopentane, 2,5-Piperazinedione, 3,6-bis(2-methylpropyl) and Butanedioic acid, hydroxy-, diethyl ester present in ethyl acetate extract against for MRSA.
- This Secondary metabolites (N-Hexadecanoic acid, N-Nonadecanol-1) may be a good alternative treatment against antibiotic-resistant pathogen *Acinetobacter baumannii* (10)

Antimalarial Compounds

Malaria, as one of the leading causes of morbidity and mortality, affects millions of people every year. World Malaria Report (2012) summarizes 106 countries affected by malaria in 2011. According to the estimation of WHO (2010), 3.3 billion people were at risk of malaria in 2010 and around 91% of deaths in the African Region alone. Plasmodium, a protozoan parasite includes five different species *P. falciparum*, *P. vivax*, *P. ovale*, *P. malariae* and *P. knowlesi* causes the highly infectious disease Malaria, transmitting through the bite of infected female *Anopheles* sp mosquitoes. More prevalent and less virulent was *P. vivax* occur in throughout the world. Three species have occurred in several parts of the world. Despite developments of effective vaccines, vector control and drugs there are a steep raise to resistance for the existing antimalarial drugs, limited level successes in the vector control methods (11).

(*Streptomyces flavidofuscus*) crude extracts having highest inhibition percentage against *Plasmodium falciparum* culture determined by SYBR green assay. The major bioactive compounds n-Hexadecanoic acid and 6-Octadecanoic acid present in the TP3 ethyl acetate extract confirmed through GCMS technique. *In silico* studies confirmed the binding energy 5.0 kcal/mol values against the *Plasmodium falciparum* pflDH virulent protein.

Cytotoxic compounds


The term cancer is a very serious scared disease in the past era. According to the literature research and WHO reported in 2019, 8 million people died from different types of cancers worldwide for every year. Specifically, 1.93 million people are died from Colon and rectum cancer. It is the 3rd dangerous cancer in both sexes (12). Currently, advanced treatments for this cancer are Chemotherapy and surgery. Naturally obtained chemotherapeutic compounds have been used against cancers, in recent years emergence needed to synthesize and designing anticancer compounds (13).

This Secondary metabolites N-Hexadecanoic acid, N-Nonadecanol-1 may be a good alternative treatment Colon cancer. *Further* structural elucidation and investigations of this compound may solve the problem in treating infections caused by antibiotic-resistant bacteria and Colon cancer causing HT29 cell line. MTT assay results revealed Purified Peptide from *Streptomyces smyrnaeus* showed the IC₅₀ value of 37.87± 0.09 µg against MDA-MB-231 cell line.

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Extraction of secondary metabolites from halophilic actinomycetes and its characterization studies

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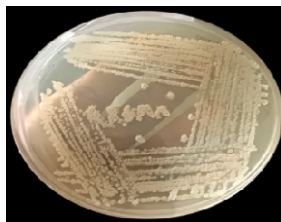
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Introduction

Marine Actinomycetes

Actinomycetes groups existing in the marine environment are important sources for the medically important drugs and other active compounds. Considering the importance of natural compounds from the marine actinomycetes. Actinomycetes particularly, *Streptomyces* species are producing wide variety of natural products with potential bioactivities. The microbial-derived metabolites hold a strong position to combat emerging and re-emerging antimicrobial drug-resistant pathogens. A diverse group of actinomycetes strains were isolated from unexplored regions of mangrove sediment. Actinomycetes are free-living microorganisms commonly found in soil, sediments and marine water. Actinomycetes are filamentous gram-positive bacteria that come under the phylum of *Actinomycetales* well defined by aerial mycelium. Actinomycetes consist 50% of guanine and cytosine content in nucleotides (1). Actinomycetes are having branching filamentous and bear chains of spores. They are produced lot of new bioactive molecules. In world, 80% of antibiotics derived from actinomycetes particularly marine actinomycetes (2).

Fig.1. Pure culture of actinomycetes



Marine actinomycetes have the greatest metabolic diversity and are distinguished by low temperature, variable salinity and low sufficient oxygen level. Marine actinomycetes are inhabiting extreme environments for adaptation and it's having unique survival strategies for growth and reproduction (3). Marine actinomycetes are rich sources of biological activities such as antimicrobial, cytotoxic compounds, anti-inflammatory agents, and antioxidant compounds. So many researchers focused on unexplored marine environments (4). Marine *Actinomycetes* represent a rich and valuable source of distinct and promising substances. The genus *Streptomyces* in particular, has been extensively studied due to its ability to produce bioactive compounds and its abundance of biosynthetic gene clusters (BGCs).

Halophilic actinomycetes

Halophilic Actinomycetes can be found in a variety of hypersaline environments. Hypersaline environments are salt pans, salt lakes, marine environments, oceans, seas, salt foods and saline soils. (4). Saltern is the extreme habitat that promotes halophilic actinomycetes metabolic adaption to saline conditions. Hypersaline environments with high salt concentrations can be found throughout the salt pans. The salt pan is formed naturally by the evaporation of seawater. Halophilic actinomycetes are producers of various bioactive compounds. Halophiles are salt-loving microorganisms and can only survive at high salt concentrations, and they can be classified according to how much salt they need to thrive. Halophiles are divided into three categories: slight halophiles (1.7-4.8%), moderate halophiles (4.7-20) and extreme halophiles (20 - 30%) of NaCl (5).

Extraction of bioactive molecules from halophilic actinomycetes

Supernatant extraction

The potential actinomycetes strain was inoculated into Starch casein broth and incubated at 28°C in a shaker at (120 rpm) for 7 days. After incubation, the broths were filtered through Whatman No. 1 filter paper. Then the filtrates were centrifuged separately at 10000 rpm for 10 min to extract the supernatant. It was transferred aseptically into screw-capped bottles and stored at 4°C for further assay.

Mass production and extraction of bioactive compounds

The pure culture of potential strains grown on 250ml of Production media (ISP4). Medium contains soluble starch-10g, Dipotassium phosphate-1g, Magnesium sulphate-1g, Sodium chloride- 1g, Ammonium sulphate- 2g, Calcium carbonate-2g, Ferrous sulphate- 1g, Magnesium chloride- 1g, Zinc

sulphate-1g and Agar-20g. The culture was incubated at 28⁰C for 7 days in the rotary shaker. After incubation, bioactive compounds were harvested by centrifugation (10,000 rpm) at 10 minutes. Collect the supernatant and add an equal volume of solvents (1:1) ratio and the mixture was shaken overnight and filtered. The filtrates were evaporated using a rotary evaporator at 50⁰C. The mass production of crude extracts stored at 4⁰C for further use (6).



Fig.2.Extraction of secondary metabolites

Characterization of bioactive molecules

Preparative High Performance Liquid chromatography

The fractions which showed the highest inhibitory action against tested pathogens were subjected to silica gel column chromatography was further separated by preparative HPLC to obtain purified compounds. The analytical study was performed using Dionex Ultimate 3000 HPLC system (Agilent) using a C18 column. C18 stationary phase in the form of USP-L1, Carbon percentage of 20%, particle pass through the size of 2 to 5 micrometers (μm), pore size of 120 Å, and manufactured with multi-stage end capping technology. The mobile phase solvent gradient was set up with 100% H₂O (HPLC grade) and 0% acetonitrile (ACN; HPLC grade) at a flow rate of 3ml/min, and the percentage of ACN was linearly increased to 100% at 25min. Final washing was performed for 15 min, using 100% ACN. Then, a second Preparative HPLC analysis run was performed for retention time [Rt] = 25 min using the same protocol as the first HPLC run, except that the solvent gradient started with 30% H₂O and 70% methanol (HPLC grade) and linearly increased (7).

GC-MS analysis

GC-MS analysis of crude extract was carried out on Shimadzu 2010 plus comprising a AOC-20i auto sampler and gas chromatograph interfaced to a mass spectrometer instrument employing the following conditions: column RTX5Ms (Column diameter is 0.32 mm, column length is 30m, column thickness 0.50 μ m), operating in electron impact mode at 70eV; Helium gas (99%) was used as carrier gas at a constant flow of 1.73 ml /min and an injection volume of 0.5 μ l was employed (split ratio of 10:1) injector temperature 270°C; ion-source temperature 200°C. The oven temperature was programmed from 40°C (isothermal for 2 min), with an increase of 8 °C/min, to 150°C, then 8°C/min to 250°C, ending with a 20min isothermal at 280°C. Mass spectra were taken at 70eV; a scan interval of 0.5 seconds and fragments from 40 to 450 Da. Total GC running time is 51.25 min.

The relative percentage amount of each component was calculated by comparing its average peak area to the total areas. Software adopted to handle mass spectra and chromatograms was a Turbo Mass Ver 5.2.0. Interpretation on GC-MS was conducted using the database of National Institute Standard and Technology (NIST) having more than 62,000 patterns. The spectrum of the unknown component was compared with the spectrum of the known components stored in the NIST library. The name, molecular weight and structure of the components of the test materials were ascertained (8).

UV-visible spectroscopy

The primary characterization of the purified fraction was carried out using UV-visible spectroscopy. The samples were periodically monitored for the colour change and measuring by UV-spectrophotometer (UV-1800 series. Shimadzu corporation Ltd, Kyoto, Japan). The spectrum was recorded throughout a range of 200- 800 nm for the wavelength corresponding to maximum absorption was determined.

FT-IR analysis

The infrared (FT-IR) spectrum of the purified fraction was determined by EXI- Spectrum One Model. The spectrum was obtained using potassium bromide (KBr) pellet techniques in the range of 4000 to 400 cm^{-1} at a resolution of 1.0 cm^{-1} . Potassium bromide (AR grade) was dried under vacuum at 100°C and 100 mg of KBr pellet was used. The spectrum was plotted between intensity and wavenumber. The FT-IR spectra were analyzed for the presence of various functional groups (9).

NMR Spectroscopic Analysis

The carbon and hydrogen (^1H and ^{13}C) NMR spectrum was studied for the purified fractions were recorded on Bruker instrument (AV300MHz) using CDCl_3 and DMSO. The solvent Tetramethylsilane as a standard with the spectral data available of the structure of the extracted pure bioactive compounds. The two dimensional structure of the compound was got using the Chem3 Draw Ultra software version 10.

***In-Silico* analysis of identified compound**

Protein Preparation

The structure of proteins three dimensional structure was downloaded from RCSB PDB database (<https://www.rcsb.org/structure>). The downloaded protein model was proceeded to energy minimization using YASARA energy minimization server (<http://www.yasara.org/minimizationserver.htm>). Furthermore, the model structure was refined using ModRefiner web server (<https://zhanggroup.org/ModRefiner/>). The prepared protein model was subjected to docking study (10).

Protein Secondary structure prediction

Protein secondary structure prediction retrieves secondary structure, families of related proteins. Solvent accessibility was performed. Prediction of subcellular localization of the protein was also analysed using <https://predictprotein.org/>

Active site prediction

The amino acids in active sites of a protein model was predicted using CASTp web server (<http://sts.bioe.uic.edu/castp/index.html?2011>). The larger volume active sites were focused and average sum of active sites volume was calculated to fix the grid box in Auto Dock Vina while doing docking step.

Molecular docking

In docking studies the identified compound selected and the structures retrieved from PubChem databases. The compound is docked with selected Proteins were retrieved from the protein data bank sources. After identification of protein active binding site, molecular docking was analysed by using Autodock vina MGL tools. In Autodock vina protein is loaded with macromolecules, and waters are removed. Then hydrogen polar bonds are added manually and add charges. Then macromolecule files are saved as PDBQT. Ligand files are input and saved as PDBQT files. Grid box are formed

in centre grid are $X = 21.26$, $Y = 0.045$ and $Z = -14.48$. And size $X = 20$, $Y = 20$ and $Z = 20$.

Then configuration file was made by using all the details correctly. Command prompts are run by using data. After the file was generated, it was split by using Vina split. Then it is analysed by using BIOVIA Discovery studio software used for protein and ligand interactions and publication-quality images. After protein and ligand docking analysis, interactions of protein-ligand sites (amino acid), and the bioactive compound against all targeted proteins shows the potent minimum binding energy and inhibitory concentration (11).

Model Building

Models are built based on the target-template alignment using ProMod3 Coordinates which are conserved between the target and the template are copied from the template to the model. Insertions and deletions are remodeled using a fragment library. Side chains are then rebuilt. Finally, the geometry of the resulting model is regularized by using a force field (12).

Molecular Visualization

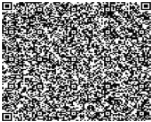
The best docked poses of protein-ligand complex was visualized using BIOVIA Discovery studio open source software. The hydrogen bond interaction, hydrophobic interaction, pi-pi interaction of protein-ligand complex was further analyzed. The amino acids shown interaction with docked protein was noted.

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Catastrophic Mass Mortality of *Hiatula diphos* in the Punnaikayal Estuary: Implications of Climate Change and Extreme Weather Events

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Abstract

The Punnaikayal Estuary, a critically important ecosystem located on the southeastern coast of Tamil Nadu, India, experienced a devastating mass mortality event of the Purple Clam (*Hiatula diphos*) in the aftermath of extreme rainfall and flooding in December 2023. This bivalve mollusk plays a vital role in the estuarine ecosystem and holds significant ecological and

economic importance. The catastrophic event resulted in the death of over 3,000 individuals, which were subsequently buried in the soil, underscoring the severity of the impact. In this study, we investigated the physiochemical alterations within the Punnaikayal Estuary that may have contributed to the mass mortality event. Drastic fluctuations in salinity levels, temperature, dissolved oxygen, pH, and turbidity were observed following the heavy rainfall and flooding. Sudden drops in salinity and temperature, coupled with reduced dissolved oxygen levels and increased turbidity, likely exceeded the tolerance limits of *Hiatula diphos*, leading to widespread physiological stress and mortality. Furthermore, the influx of freshwater and suspended sediments disrupted the benthic habitats, physically dislodging and burying the clams. The introduction of pollutants and contaminants from terrestrial sources during the flooding event may have exacerbated the lethal conditions for these bivalves. Additionally, the stress induced by the environmental changes could have made the clam population more susceptible to diseases and parasitic infections, further contributing to the mass mortality. Our findings highlight the urgent need for comprehensive monitoring programs to track physiochemical parameters and population dynamics within estuarine ecosystems.

Keywords: Estuarine ecosystem, Mass mortality, Climate change, Extreme weather events, Punnaikayal.

Introduction

Estuaries are dynamic and productive ecosystems that serve as nursery grounds for various marine species and provide valuable ecological services (Courrat *et al.*, 2009; Vasconcelos *et al.*, 2011; Sheaves *et al.*, 2015). Estuarine management can reduce flood risk, promote economic development, and conserve nature while maintaining ecosystem services and biodiversity (Boerema & Meire 2017). Invasive freshwater bivalves are prone to mass mortality events due to their low tolerance to abiotic stressors, such as extreme conditions like drought, flooding, and high and low temperatures (McDowell & Sousa, 2019).

Mass mortality episodes of bivalves are caused by climatic and anthropogenic stressors, and improving the resilience of natural systems can help mitigate these events Guillotreau *et al.*, (2017). The 2003 heat wave in Europe likely contributed to mass mortality of rocky benthic macro-invertebrates in the Northwestern Mediterranean, potentially driving a major biodiversity crisis in the region Garrabou *et al.*, (2008). A mass mortality event affecting the bivalve *Pinna nobilis* in the Spanish Mediterranean Sea has spread rapidly, causing high mortality rates up to 100% in some areas

Vázquez-Luis *et al.*, (2017). The mass mortality of the *Tagelus plebeius* population in the Bah12a Blanca Estuary is due to catastrophic burial by sandy sediment of fluvial origin Farinatiet *al.*, (1992). The mass mortality event in the Mediterranean Sea is caused by a newly described *Haplosporidian* endoparasite, *Haplosporidium pinnae*, which is closely related to temperatures above 13.5°C and salinity ranges between 36.5–39.7 psu. Cabanellas-Reboredo *et al.*, (2019).

The Punnaikayal Estuary, situated at the confluence of the Tamiraparani River and the Bay of Bengal, supports a rich biodiversity, including the Purple Clam (*Hiatula diphos*). This bivalve species plays a vital role in nutrient cycling, energy transfer, and serves as a food source for numerous organisms, including fish and migratory shorebirds. Purple clams, *Hiatuladiphos*, are popular seafood delicacies among the Taiwanese and play a role in the transmission of paralytic shellfish poisoning toxins from dinoflagellate to gastropod through a filter-feeding bivalve (Chen *et al.*, (1998); Chou *et al.*, (2005). It contains saxitoxin-like substances, which may amount to 0.013% tetrodotoxin equivalent (Chang *et al.*, 1986).

To address the critical issues raised by this catastrophic event, the present study aims to conduct a comprehensive investigation into the physiochemical alterations within the Punnaikayal Estuary that may have contributed to the mass mortality of *Hiatula diphos* following the extreme rainfall and flooding in December 2023.



Figure 1. Punnaikayal Estuary

The Catastrophic Event

In December 2023, the Tirunelveli and Tuticorin regions of Tamil Nadu experienced unprecedented heavy rainfall, with accumulations reaching up to 95 cm. The swollen Tamiraparani River discharged its waters into the Punnaikayal Estuary, leading to severe flooding. During the subsequent months of January, February, and March 2024, a significant mass mortality event of the Purple clam was observed in the estuary.

Factors Contributing to Mass Mortality

1. Freshwater influx: The heavy rainfall and flooding resulted in a substantial influx of freshwater into the estuary, drastically reducing salinity levels. *Hiatula diphos*, being a marine bivalve, is highly sensitive to prolonged exposure to low salinities, which can disrupt its osmoregulatory processes and lead to physiological stress and mortality.

2. Sediment disturbance and burial: The intense floodwaters carried large volumes of sediment and debris into the estuary, potentially burying or smothering the clam beds. *Hiatula diphos*, being a sedentary species, is vulnerable to excessive sedimentation and burial, which can impair its ability to feed, respire, and maintain its position within the substrate.

3. Habitat alteration: The heavy flooding significantly altered the physical and chemical characteristics of the estuary, such as substrate composition, water flow patterns, and nutrient dynamics. These changes rendered the habitat unsuitable for the survival and reproduction of *Hiatula diphos*.

4. Reduced water quality: Floodwaters often carry pollutants, including agricultural runoff, industrial effluents, and sewage, which can degrade water quality in the estuary. Poor water quality can have detrimental effects on the health and survival of bivalve populations like *Hiatula diphos*.



Figure 2. Bivalve Shell *Hiatula diphos*

Physiochemical Parameters

Estuarine environments are highly dynamic, with fluctuations in various physiochemical parameters, including salinity, temperature, dissolved oxygen, pH, and turbidity. Extreme rainfall events can significantly alter these parameters, causing abrupt changes that may exceed the tolerance limits of estuarine organisms like *Hiatula diphos*.

Salinity: Heavy rainfall can lead to a sudden influx of freshwater, drastically reducing the salinity levels in the estuary. Bivalves like *Hiatula diphos* are adapted to specific salinity ranges, and sudden drops or fluctuations can disrupt their osmoregulatory processes, leading to physiological stress and mortality.

Temperature: Flooding can introduce large volumes of cooler freshwater, causing sudden temperature changes in the estuary. Rapid temperature fluctuations can affect the metabolic processes and respiratory functions of bivalves, making them more susceptible to stress and mortality.

Dissolved Oxygen: Heavy rainfall and flooding can lead to increased turbidity and suspended sediments, which can reduce the availability of dissolved oxygen in the water. Bivalves rely on dissolved oxygen for respiration, and prolonged exposure to low oxygen levels can be lethal.

pH: Sudden influxes of freshwater or changes in water chemistry can alter the pH of the estuarine water, potentially affecting the acid-base balance and physiological processes of bivalves like *Hiatula diphos*.

Turbidity: Increased suspended sediments and turbidity can clog the gills of bivalves, interfering with their feeding and respiration processes, leading to stress and potential mortality.

Ecological and Economic Implications

The mass mortality of the Purple Clam has severe implications for the Punnaikayal Estuary ecosystem. This event disrupts the nutrient cycling and energy transfer processes, potentially impacting the entire food web and affecting other organisms that rely on this bivalve as a food source. Additionally, the loss of this valuable resource can have significant economic consequences for local communities that depend on clam harvesting for their livelihood and sustenance.

In addition to the physiochemical changes, other factors may have contributed to the mass mortality event, including:

Habitat destruction: Heavy rainfall and flooding can physically disrupt the benthic habitats where *H. diphos* resides, leading to dislodgement, burial, or erosion of their habitat.

Pollutants and contaminants: Flooding can introduce various pollutants and contaminants from terrestrial sources, which can be toxic to bivalves and other estuarine organisms.

Disease and parasites: Stress induced by environmental changes can make bivalves more susceptible to diseases and parasitic infections, contributing to their mortality.

Conclusion

The mass mortality of the *Hiatula diphos* in the Punnaikayal Estuary serves as a stark reminder of the vulnerability of estuarine ecosystems and their associated fauna to the impacts of climate change and extreme weather events. This catastrophic event underscores the urgent need for comprehensive monitoring, conservation efforts, and adaptive management strategies to ensure the long-term sustainability of these ecologically and economically important bivalve populations. By addressing the challenges posed by environmental changes and implementing proactive measures, we can safeguard the invaluable resources and services provided by estuarine ecosystems for present and future generations.

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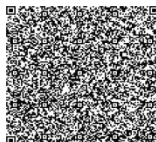
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Plants used as cosmetics

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Introduction

The history of cosmetics spans at least 7,000 years and is present in almost every society on Earth. Cosmetic body art is argued to have been the earliest form of a ritual in human culture. The evidence for this comes in the form of utilized red mineral pigments (red ochre) including crayons associated with the emergence of Homo sapiens in Africa. Cosmetics are mentioned in the Old Testament-2 king where Jezebel painted her eyelids- approximately 840 BC – and the book of Esther describes various beauty treatments as well.

Cosmetics were also used in ancient Rome, although much of Roman literature suggests that it was frowned upon. It is known that some women in ancient Rome invented make up including lead-based formulas, to whiten the skin, and Kohl to line the eyes.

By the time Queen Cleopatra came to power in the 1st century BC, Egyptian women had at their disposal a whole rainbow of cosmetics, all of which were made from rocks, minerals and plants in the region. Cleopatra used the bright green malachite paste of the ancient Egyptians on her lower eyelids. One of the most expensive beauty secrets of Cleopatra was that she slept with a golden mask on her face. Her beauty secrets are well versed as they are well documented. Apparently, Egyptian queen Cleopatra used to incorporate some of the amazing masks, treatments and baths into her beauty regime.

The giant cosmetics companies that generate billions of dollars of revenue annually were founded in the twentieth century by chemists and pharmacists in the United States. A well-known cosmetic company started by selling books door - to - door, along with colognes manufactured by a friendly pharmacist. After the First World War, the athletic look became fashionable for the first time, and the fashion statements made by Coco Channel, including

dark eyes, red lipstick, red nail polish and suntanned skin, which became popular as a contrast to the dominant fad for pale skin.

Egyptian customs were exported to Greece and Rome. Indeed, the world cosmetic is derived from the Greek Kosmetos, which means adornment or ornament.

Cosmetic products are also important in Israeli culture. Before online shopping facilities existed, people visited the Dead Sea to experience the historical context of the place and to float in the hypersaline water; they also spent large amount of money purchasing cosmetic product such as mud, clay, and water. These products have been used for thousands of years for skin care and the treatment of illnesses, and have a balanced composition of mineral salts.

Cosmetics are the substances especially prepared to enhance beauty and increase the attractiveness of the person. The prehistoric men who were cave dwellers used leaves of trees and skin of animals to protect themselves from vagaries of nature.

A cosmetic is an item intended to be rubbed, poured, sprinkled or sprayed on, introduced in to or otherwise applied to the human body or any part for cleansing, beautifying and alternating the appearance.

All cosmetic preparation has their application for long or short periods to beautify the body healthy up to some extent and has physiological impact to others. The "active life" of any cosmetic preparation begins the moment it is brought in contact with the skin, hair etc. Cosmetics that are used for hair care purpose applied orally and should not be used for therapeutic purpose. Basic feature of hair care cosmetics is as-

1. Should be easy to use
2. Should have local effect.
3. Should be harmful to hair skin and mucous membrane.
4. Should not be allergic to body.
5. Should be applied topically.

Hair care cosmeceutical formulations mainly include shampoo, gel, lotion solution and oil. In modern cosmetology, the all products of cosmetic preparations manufactured under strict quality control conditions to achieve an absence of claims on both appearance and packing.

Beneficial effects of herbs/plants as cosmeceuticals

Treat skin aging

Prevent hyper pigmentation

Prevent skin wrinkling

Treat skin ailments

Beautification

Strengthening of hair growth

Inhibit skin dryness

Result:

Table 1. Botanical name, family, habit and part used

Si no	Binomial name	Family	Habit	Part used
1	<i>Acalypha indica</i> L.	Euphorbiaceae	Herb	Leaf
2	<i>Aloe vera</i> (L.) Burm.f.	Asphodelaceae	Herb	Leaf
3	<i>Azadirachta indica</i> A.Juss.	Meliaceae	Tree	Leaf
4	<i>Beta vulgaris</i> L.	Amaranthaceae	Herb	Root
5	<i>Camellia sinensis</i> (L.) Kuntze	Theaceae	Shrub	Leaf
6	<i>Carica papaya</i> L.	Cariaceae	Tree	Fruit
7	<i>Citrus × aurantium</i> f. <i>aurantium</i>	Rutaceae	Tree	Fruit
8	<i>Citrus × limon</i> (L.) Osbeck	Rutaceae	Tree	Fruit
9	<i>Cucumis sativus</i> L.	Cucurbitaceae	Creeper	Fruit
10	<i>Curcuma longa</i> L.	Zingiberaceae	Herb	Rhizome
11	<i>Daucus carota</i> L.	Apiaceae	Herb	Root
12	<i>Hibiscus × rosa-sinensis</i> L.	Malvaceae	Shrub	Flower

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13	<i>Lavandula angustifolia</i> Mill.	Lamiaceae	Herb	Flower
14	<i>Mangifera indica</i> L.	Anacardiaceae	Tree	Fruit
15	<i>Ocimum basilicum</i> L.	Lamiaceae	Herb	Leaf
16	<i>Opuntia stricta</i> (Haw.) Haw.	Cactaceae	Shrub	Fruit
17	<i>Psidium guajava</i> L.	Myrtaceae	Shrub	Fruit
18	<i>Punica granatum</i> L.	Lythraceae	Shrub	Fruit
19	<i>Ricinus communis</i> L.	Euphorbiaceae	Shrub	Fruit
20	<i>Rosa indica</i> L.	Rosaceae	Shrub	Petal
21	<i>Santalum album</i> L.	Santalaceae	Tree	Wood
22	<i>Senna auriculata</i> (L.) Roxb.	Fabaceae	Shrub	Flower
23	<i>Solanum lycopersicum</i> L.	Solanaceae	Herb	Fruit
24	<i>Solanum tuberosum</i> L.	Solanaceae	Herb	Tuber
25	<i>Tagetes erecta</i> L.	Astraceae	Herb	Leaf
26	<i>Vitis vinifera</i> L.	Vitaceae	Climber	Fruit

***Acalypha indica* L.**

Common name : Indian acalypha

Based on literature studies, *Acalypha indica* L. has been found to possess various medicinal properties, including anthelmintic, anti-inflammatory, antibacterial, anti-cancer, anti-diabetic, anti-hyperlipidaemic, anti-obesity, anti-venom, and wound healing capabilities. Additionally, the leaves can be used for cosmetic purposes, such as removing unwanted hair.

Concoction:

A powder made from the leaves of *Acalypha indica* L. can be used to promote healthy, fresh-looking skin. To create this natural remedy, grind the leaves of *Acalypha indica* L., turmeric, and Neem together and apply the paste to your skin while bathing.

Consequence of Outcome:

The uses of *Acalypha indica* L. are diverse, ranging from removing unwanted hair to treating colds, coughs, and various skin problems.

Aloe vera (L.) Burm.f.

Common name: Aloe vera

Aloe vera Burm.f. is a widely popular plant renowned for its numerous beauty benefits and healing properties. It is a key ingredient in various cosmetic and pharmaceutical products. Leaf gel is rich in nutrients, comprising 75% nutrients, 20% minerals, 18% amino acids, and 12% vitamins.

Concoction:

To boost hair growth, massage leaf gel into your scalp and rinse after 30 minutes. For a facial treatment, mix one tablespoon of leaf gel with almond oil and massage onto your face for nearly 5 minutes. After 30 minutes, wash your face.

Consequence of Outcome:

Aloe vera (L.) Burm.f. has moisturizing properties that enrich the skin's water level, giving you a glowing, glass-like complexion with regular application.

Azadirachta indica A.Juss.

Common name: Neem

Neem leaves are rich in antioxidants, which enable them to combat acne-causing bacteria.

Concoction:

To harness the benefits of neem, grind some leaves into a paste. Mix in a pinch of turmeric powder and apply the blend to your face. Leave it on for 10 minutes before rinsing with warm water.

Consequence of Outcome:

This natural remedy helps achieve acne-free skin by preventing future breakouts and reducing the appearance of existing acne spots.

Beta vulgaris L.

Common name: Beetroot

Beta vulgaris L. is rich in folates, which help cells grow and function. It is naturally high in nitrates, which are converted into nitric oxide in the body.

Concoction:

To prepare this face pack, take 1 teaspoon of raw milk, 2-3 drops of almond oil or coconut oil, and 2 tablespoons of Beetroot juice. Mix the ingredients properly and massage them onto your face for 10 minutes. Then, wash with plain water and observe the results.

Consequence of Outcome:

For glowing skin. The red food helps you to get that nature glow to your skin and helps to keep it nourished and hydrated for skin brightening. Vitamin C is a magic nutrient well known as brightening agent for dry skin for acne. Anti-aging

***Camellia sinensis* (L.) Kuntze**

Common name: Tea plant

Camellia sinensis (L.) Kuntze is rich in antioxidants, which help fight free radicals that damage cells and cause premature aging. This plant has wonderful benefits for the skin and hair, both when applied topically and when ingested.

Concoction:

To reap the benefits of *Camellia sinensis* (L.) Kuntze, try the following: Use dry tea leaves as an exfoliator for your skin. Mix the contents of a green tea sachet with honey to create a soothing scrub. Steep leaves in a glass of boiling water, then let it cool. Refrigerate the mixture for 30 minutes before applying it to dark circles to reduce pigmentation.

Consequence of Outcome:

The antioxidants in *Camellia sinensis* (L.) Kuntze have several benefits: Reduces dark circles and helps whiten and even out skin tone.

***Carica papaya* L.**

Common name: Papaya

The enzymes papain and chymopapain in papaya have anti-inflammatory properties. Papain, a protein-dissolving enzyme, is commonly found in many exfoliating products. These products help reduce acne by removing dead skin cells that can clog pores. Additionally, papain can dissolve keratin, a protein that can build up on the skin and form small bumps.

Concoction:

To reap the benefits of *Carica papaya* L. fruit, try this face mask: Smash half a cup of overripe fruit and mix it with one tablespoon each of milk and honey. Apply the mask to your face and neck.

Consequence of Outcome:

Carica Papaya L. fruit helps achieve a glowing complexion due to its rich content of vitamin C compounds.

Citrus × aurantium f. aurantium

Common name: Sweet orange

Citrus × aurantium f. aurantium contain excellent source of fiber and a rich source of vitamin C and folate, among many other beneficial nutrients.

Concoction:

To create a face pack, mix together: 1 teaspoon of *Citrus × aurantium f. aurantium* fruit peel powder a pinch of cosmetic turmeric 1 teaspoon of natural honey and blend the ingredients into a fine paste and apply it to your face and neck. Rinse off with a gentle face cleanser or rose water after 5 to 10 minutes.

Consequence of Outcome:

The face pack offers several benefits: Absorbs excess oil from the face, removes tan and evens out skin tone, exfoliates and reveals clear, healthy skin and shrinks pores and smooths the skin.

Citrus × limon (L.) Osbeck

Common name: Lemon

Lemon oil extract serves as a source of citrus fragrance in various skincare products. It is characterized by rich amounts of antioxidants, such as glutathione, ascorbic acid, and alpha-tocopherol, which can help fight against skin-damaging lipid peroxidation.

Concoction:

Lemon Brightening Mask. To prepare it, you will need an egg white and lemon juice. Whip the egg white until it becomes frothy, then add a few drops of *Citrus × limon (L.) Osbeck*

fruit juice. Apply the mixture to the face, avoiding the eye area and lips. Hold for 10 minutes and rinse with warm water. This mask can also be applied to the hair to treat dandruff.

Consequence of Outcome:

These compounds play a physiological role in protecting the surface level of the skin

Cucumis sativus L.

Common name: Cucumber

Cucumber possesses anti-inflammatory, emollient, and moisturizing properties. It contains 95% water, vitamin C, antioxidants, silica, and fatty acids, and is widely used in lotions, creams, and cleansers for its hydrating properties.

Concoction:

Cut a cucumber in half and grind it along with the peel. Strain the juice and apply it to a clean face like a toner. This will help brighten your complexion and reduce pore size.

Consequence of Outcome:

Cucumber helps to moisturize the skin, as it is a water-rich fruit. It also helps reduce wrinkles on the face.

Curcuma longa L.

Common name: Turmeric

Turmeric is a versatile spice that can be combined with various ingredients to address different skin concerns. For hyperpigmentation and wrinkles, mix turmeric with yogurt and lemon juice for added nourishment and brightening effects. To soothe irritation, blend turmeric extract with aloe vera gel for a natural calming remedy.

Concoction:

To combat acne and antibacterial concerns, combine turmeric with warm water and honey. This mixture can help to promote glowing skin, boost healing, and potentially alleviate other skin conditions

Consequence of Outcome:

Turmeric is considered a crucial ingredient in Ayurveda for skin whitening. It is known to reduce dark patches and even out skin tone, resulting in a more radiant complexion.

Daucus carota L.

Common name: Carrot

Carrot is an excellent source of essential nutrients like potassium, antioxidants, and vitamin A. Beta-carotene, a type of carotenoid, is the nutrient responsible for carrots' vibrant colour and numerous health benefits.

Concoction:

To create a nourishing face mask, mix one cup of carrot juice with one tablespoon each of curd, gram flour, and lemon juice. Apply the mask to your face and neck for 30 minutes, then rinse with lukewarm water.

Consequence of Outcome:

The carrot face mask leaves you with glowing, spotless skin. It rejuvenates your skin by removing dead cells, making it ideal for oily skin. The mask also hydrates and improves skin complexion, while keeping bacteria that cause skin infections at bay.

Hibiscus × rosa-sinensis L.

Common name:Hibiscus

Hibiscus rosa-sinensis is rich in amino acids, antioxidants, and oligo-peptides, which help reduce fine lines and wrinkles, promoting anti-aging effects. The flowers are commonly used in various skin and hair nourishment products.

Concoction:

Hibiscus leaves serve as a natural hair conditioner due to their mucilage polysaccharides content. To utilize this benefit, boil 15-20 leaves and apply the resulting water to your hair. Consistently using this treatment for 2 weeks can lead to reduced hair fall and increased shine.

Consequence of Outcome:

Hibiscus has been shown to reduce pimples and remove dark patches, resulting in clearer skin. Additionally, it promotes shiny, thicker hair, making it an excellent natural hair care solution.

Lavandula angustifolia Mill.

Common name: Lavender

Lavender is an aromatic, pure plant that offers relaxation for the skin and mind. Its pleasant aroma makes it a popular ingredient in various scented products, such as perfumes and bath and body care items. In addition to its calming

effects, lavender possesses antiseptic and anti-inflammatory properties, which help soothe itchy skin conditions like eczema and psoriasis.

Concoction:

To harness the benefits of lavender: Treat sunburn by mixing a few drops of lavender oil with aloe gel. Prepare a face mask by boiling lavender flower petals for a few minutes. Strain the water, remove the petals, and mix with a teaspoon of oat powder to create a paste. Apply the mask to your face and rinse with water when it dries.

Consequence of Outcome:

Lavender promotes healthy, glowing skin due to its rich vitamin content. It helps to: Soothe and calm irritated skin, reduce inflammation and promote healing and improve skin tone and texture

Mangifera indica L.

Common name: Mango

Mango is rich in beta-carotene, collagen, and vitamin A, making them an excellent ingredient for reviving dull skin and achieving a radiant complexion. The beta-carotene in mangoes also helps to combat acne. The best time to apply mango is at night, as it serves as an exfoliating agent. Mangoes contain tryptophan, which aids in the production of serotonin, a hormone that promotes happiness. This is why mangoes are often referred to as the "happy fruit."

Concoction:

Apply a handful of mango pulp to your face at night and leave it on for an hour before washing it off before bed. Alternatively, boil a mango until the water is reduced to half, then use this water as a facial astringent to dry up acne naturally. Regular use of the mango astringent can help eliminate acne, pimples, and scars.

Consequence of Outcome:

Using mango on your skin can give you a golden glow on your cheeks, reduce acne spots, and leave you with a blemish-free, glowing face within just one week.

Ocimum basilicum L.

Common name: Basil

Basil has various uses beyond its medicinal properties, particularly in the cosmetic industry. It is primarily used in acne treatments, as it helps to clean oil and dirt from clogged pores.

Concoction:

To create a natural acne treatment, mix 15-20 fresh basil leaves with 2 teaspoons of curd. Apply the mixture to your face and leave it on for 15 minutes. This pack can help to make your skin spotless, acne-free, and healthy. Alternatively, combine basil leaves with 1 teaspoon of aloe gel and grind well. Store this mixture in the fridge to create a cooling icing, then apply it to your face.

Consequence of Outcome:

Basil's antibacterial and anti-inflammatory properties help prevent acne. Additionally, basil is an excellent pore cleanser, making it a valuable ingredient in skincare routines.

Opuntia stricta (Haw.) Haw.

Common name: Prickly pear

The cladodes, fruits, and seeds of Opuntia are rich in vitamins, minerals, and acids, making them a valuable source of natural ingredients for cosmetic creams. These ingredients possess powerful soothing, moisturizing, and anti-aging properties, providing nourishment and health benefits for the skin.

Concoction:

To promote hair growth and strengthen hair roots, mix Opuntia oil with 1 tablespoon of neem oil. Apply the mixture to your scalp and leave it on for 10 minutes before washing with regular shampoo. Additionally, the fruit of Opuntia can be used to heal cracked lips.

Consequence of Outcome:

The moisturizing content in Opuntia oil helps to strengthen hair, while its antioxidant properties aid in healing cracked lips.

Psidium guajava L.

Common name: Guava

Guava is a natural remedy that can help improve complexion and reduce dark spots. It brightens dull skin, hiding skin tan and leaving a radiant glow. Guava is rich in vitamins A, B, and C, which possess excellent antioxidant properties. These properties help keep the skin hydrated and work as an anti-aging agent, making the skin look instantly clear and fresh.

Concoction:

To create a nourishing face mask, blend guava leaves and peels in a blender to form a thick paste. Transfer the paste to a bowl and add one teaspoon of honey. Apply this pack to your face and leave it on for 15-20 minutes or until it dries. Rinse your face with mild, cool water and pat dry with a clean towel.

Consequence of Outcome:

Soft, crushed seeds of pink guava can also be used as a gentle exfoliating scrub, helping to remove dead skin cells and reveal smoother skin.

Punica granatum L.

Common name: Pomegranate

Pomegranate oil is a rich source of vitamin C, a vital nutrient in skincare. Pomegranate offers regenerative, antioxidant, and anti-aging properties that promote healthy, glowing skin.

Concoction:

To harness the benefits of pomegranate, create a paste by blending a few pomegranate seeds. Mix in 1 tablespoon of honey and apply the mixture to your face and neck. Leave it on for 30 minutes and repeat this process 2-3 times a week. You'll notice a visible glow in just a few days.

Consequence of Outcome:

The combination of ingredients in pomegranate extracts makes them highly effective. They help detoxify the skin by removing free radicals while also repairing and rejuvenating the skin.

Ricinus communis L.

Common name: Castor

Castor oil is a rich source of vitamin E, omega-9 fatty acids, and omega-6 fatty acids. It contains moisturizing fatty acids, particularly ricinoleic acid, which makes it an excellent skin care ingredient.

Concoction:

To create a nourishing face mask, mix an egg, water, and 1 tablespoon of castor oil in a bowl. Apply the mask to your face and neck area, gently massaging for 5-10 minutes. Let it sit for 30 minutes before rinsing. Additionally, castor oil can be applied to the eyelashes to promote thicker and healthier hair growth.

Consequence of Outcome:

Using castor oil can leave your face with a glowing complexion and reduce blemishes. It also helps to promote thicker and healthier eyelashes.

Rosa indica L.

Common name: Rose

Roses are renowned for their breathtaking beauty and fragrance. Beyond their aesthetic appeal, roses also have numerous uses in the cosmetic industry. Rose oil is rich in vitamins and possesses exceptional hydrating, antibacterial, and antifungal properties, making it an invaluable ingredient in skincare.

Concoction:

To create a nourishing face mask, combine fresh and dried crushed rose petals, 1 teaspoon of yogurt, and half a teaspoon of honey. Mix the ingredients and apply the mask to your face using gentle, finger-tip motions. Leave it on for 15-20 minutes before rinsing with cold water.

Consequence of Outcome:

The vitamin C present in rose petals helps to brighten and even out skin tone, leaving your face with a radiant glow. Regular use of this mask can also reduce blemishes, dark spots, and hyperpigmentation, resulting in a smoother, more luminous complexion.

Santalum album L.

Common name: Sandal

Sandalwood oil is rich in antioxidants that help maintain skin cell structure and buoyancy. It reduces dryness, replenishes moisture, and increases skin elasticity. The antioxidants in sandalwood oil fight free radical formation, promoting healthy skin.

Concoction:

Sandalwood oil has a soothing effect on inflammation caused by pimples and acne. To harness this benefit, mix sandalwood oil with turmeric powder and camphor. Apply the mixture to affected areas and leave it on overnight. Rinse with plain water the next morning.

Consequence of Outcome:

Sandalwood oil is commonly used to treat inflammatory and eruptive skin diseases, making it a valuable ingredient in skincare routines.

Senna auriculata (L.) Roxb.

Common name: Cassia

Cassia flowers work wonders as a face pack, preventing black spots, treating uneven skin tone, and improving skin complexion. Regular use of dried flowers keeps the skin blemish-free. Cassia flowers are rich in physicochemical properties, including moisture, proteins, and fats, which help combat microbial infections.

Concoction:

To harness the benefits of cassia, grind fresh flowers and apply the paste to your face. Rinse after 30 minutes to achieve a brighter complexion. Alternatively, combine dried cassia flowers, rose petals, and orange peels, and grind them into a fine powder. Use this mixture as a full-body cleanser during bathing to reduce body odor.

Consequence of Outcome:

Regular use of cassia flowers enhances beauty and reduces body odour. Applying this mixture to the underarms can also help remove dark patches, leaving skin looking smoother and more even-toned.

Solanum lycopersicum L.

Common name: Tomato

Tomatoes are an excellent source of folate, vitamin C, and potassium. In terms of phytonutrients, tomatoes are particularly rich in carotenoids.

Concoction:

To harness the benefits of tomatoes, combine 1 tomato with 2 tablespoons of cucumber paste and 1 tablespoon of honey. Apply the mixture to your face and leave it on for 15-20 minutes. Rinse off with lukewarm water.

Consequence of Outcome:

Tomatoes contain flavonoids and pectin, which aid in exfoliation and improve skin texture. They also help fight cellular damage, penetrate moisture into the skin, and reduce the risk of sunburn.

Solanum tuberosum L.

Common name:Potato

Potatoes are a nutrient-rich food, providing a good source of vitamins B1, B2, and B6, as well as essential minerals. Additionally, potatoes contain dietary antioxidants, which may help prevent diseases.

Concoction:

To harness the benefits of potatoes for your skin, mix potato juice, rice flour, lemon juice, and honey to form a thick paste. Apply the blended paste to your face and leave it on for 15-20 minutes or until it dries.

Consequence of Outcome:

This potato-based face mask can help eliminate unsightly spots, marks, and blemishes. It also reduces inflammation caused by bruises, rashes, and ulcers, and helps minimize puffy eyes. Furthermore, it can delay the signs of aging, leaving your skin looking smoother and more radiant.

Tagetes erecta L.

Common name: Marigold

Marigold flowers are rich in carotenoids, compounds similar to vitamin A, which can help rejuvenate dull and damaged skin. The flowers also possess antiseptic properties, making them effective in treating rashes, acne, and other skin issues. Furthermore, marigold oil is a valuable ingredient in the preparation of sunscreens, conditioners, and lip balms.

Concoction:

To harness the benefits of marigold, start by washing a few petals in tap water. Then, mix the petals with 1 teaspoon of honey and 1 teaspoon of raw milk. Apply the mixture to your face and leave it on for 15 minutes. Rinse with water once it dries to reveal soft and supple skin.

Consequence of Outcome:

One of the remarkable benefits of marigold flowers is their ability to remove warts from the face.

Vitis vinifera L.

Common name: Grapes

Grapes are an excellent source of potassium, a mineral that helps maintain fluid balance in the body.

Concoction:

To harness the benefits of grapes for your skin, mash 8-9 black grapes in a bowl. Add 1 tablespoon of fuller's earth powder and 1 tablespoon of rose water to the bowl. Mix the ingredients well and apply the mixture to your face.

Consequence of Outcome:


Rich in vitamin C and antioxidants, grapes can help revitalize your skin. They offer protection against cancer-causing ultraviolet radiation and free radicals, which can cause wrinkles and dark spots. Regular use of grapes can leave your skin looking smoother, brighter, and more radiant.

Conclusion

The knowledge of medicinal plants used by local communities appears to be deeply rooted in their culture and tradition. This study identified numerous plants used to treat dermatological disorders and as cosmetics. Interestingly, some plants were found to have dual uses, serving both curative and cosmetic purposes. Further extensive ethnobotanical and ethnopharmacological studies may lead to the discovery of new plants and compounds for skin care and treatment. In India, over 70% of the population relies on herbal cosmetics for their healthcare needs. The demand for herbal cosmetics in personal care systems has increased significantly, and they are now an integral part of daily life. A healthy appearance, characterized by healthy teeth, shiny hair, and glowing skin, is essential for overall well-being. Allopathic treatments for various health issues can be expensive and inaccessible to low-income populations. In contrast, herbal medicines offer a cost-effective and affordable alternative. Although herbal cosmetics are generally considered safe, it is essential to acknowledge that some ingredients may pose health risks. Herbal cosmetics offer several advantages, including low cost, minimal side effects, and environmental sustainability. As a result, they have a promising future ahead, particularly when compared to synthetic cosmetics. Effective management of herbal resources and good manufacturing practices will be crucial in driving significant growth in the herbal cosmetics industry.

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Commercial Applications of Aeroponics

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Abstract

Aeroponics developed into a modern agricultural technique of considerable commercial viability, giving viable, scalable alternatives for modern agriculture and water-scarce regions. Among the other economic potentials offered by aeroponics are lower water and nutrients input, faster plant growth, higher yields, and small footprint cultivation. This chapter deals about the commercial application of this crop, where one can find its potential in multifaceted applications. Through an aeroponic system, one can produce high-value crops that include vegetables, leafy green crops, and medicinal plants; it helps farmers to produce more profits while employing fewer resources. Additionally, due to its versatility, it may be grown all year-round and reduces the risk of soil-borne diseases and thus eliminates the need for chemical pesticides.

The benefits of aeroponics, particularly in terms of spatial efficiency and water savings are driving its integration into vertical farming, urban agriculture, and greenhouse practices. Farmers were able to produce uniform products throughout the year, regardless of the climatic conditions, thereby reducing their dependence on geographical areas for their produce. Aquaponics appear to have the potential in regard to overcoming many of the constraints set by traditional methods. To address the growing demand for organic and pesticide-free produce, and to boost profitability, the chapter also explores case studies of companies that have successfully implemented aeroponics.

Key Words: Commercial viability, Water saving, Increased crop yield, Sustainable production, Precision agriculture.

1. Introduction

Aeroponics has revolutionized agriculture in the present day by meeting some of the very vital challenges of agriculture. Among its major advantages is resolving the issue of resource scarcity, especially when it comes to water. A considerable amount of water is wasted in conventional farming due to inefficient irrigation techniques (Garzón *et al.*, 2023). Aeroponics addresses this issue while using a lot less water by supplying nutrients in the form of a mist straight to the plant roots. Areas with low soil fertility or water availability should use this option. Fertilizers and pesticides are necessary for aeroponic systems because they operate in a controlled environment that protects surrounding ecosystems (Shrouk, 2017). For a concept to be profitable, it must be effective and scalable.

Healthy crops are produced when systems are in place because the growing environment is fully controlled. When nutrients are given straight to the roots, plants can grow faster than with soil methods, increasing operational efficiency.

Additionally, this approach uses resources, which ultimately results in lower costs. With the help of aeroponics technique, there is no season for farming and crop produce can be certain in the farm through out every season and regardless of the soil conditions. This has an added advantage in vertical farming which is space saving in urban areas. It is a preferred option for farmers and enterprises interested in maximizing the output in a small footprint as it enables active growing of plants in a vertical fashion. The advantages of these systems overtakes the disadvantages in most cases; the installation costs of aeroponic systems are high, but there is greater production efficiency in the long run due to minimal labor costs, high crop yield and low water and fertilizer requirement. This is especially advantageous in metropolitan environments where spatial constraint is a prominent challenge. For farmers and businesses aiming to optimize output in minimum growing area by growing crops in layered fashion, it is a fairly attractive option (Vacharadze, 2021). An aeroponics system is a smart investment because it gradually uses less water and fertilizer, increasing crop yields and lowering labor costs, even with the possible setup costs.

In direct response to the growing necessity of climate-resilient farming, aeroponics gives methods where plants can grow free from the pitfalls of soil and weather, in climate-controlled environments such as a greenhouse or vertical farm, where light, humidity, and temperature are closely controlled. Aeroponics was developed due to the growing requirement for farming

practices that are resistant to climate change (Yeboah *et al.*, 2022). The soil and weather constraints are not a limiting factor in plant growth when the crop is grown in highly controlled environments such as greenhouses or vertical farms that regulate the temperature, light, and humidity levels. These factors ensure that food is produced even in unpredictable or harsh weather conditions to advance global food security (Lakhia *et al.*, 2020).

Due to this reason of global food security, it ensures that food production can be continued even in erratic or harsh weather. Aeroponics can be the perfect fit for this vision, and thus, sustainability is a rapidly expanding global priority. In efficient use of resources and space, aeroponics has come as a shining green light amidst environmentally friendly farming techniques that have been getting popular over time. Carbon footprint during transporting food from rural areas to cities can be reduced if brought through vertical farming within urban settings. Aeroponics uses less energy and resources for food production in an environment-friendly manner and makes the getting of food easier and sustainable in the urban neighborhoods (Rivera *et al.*, 2022).

2. Principles and mechanisms of aeroponics

Plants grown aeroponically rely not on soil but on a spray system that feeds nutrients straight to their roots. This is a novel approach to plant cultivation. This method involves completely free hanging plant roots in the atmosphere that are misted with nutrient-rich water vapor or fine mist on a regular basis. This system allows for more oxygen to be penetrated into the roots, thus allowing enhanced ability to drink water and nutrient from the soil in comparison to traditional agricultural and soil-based systems. The best conditions for growth consist of a combination of oxygen and moisture, resulting in healthier growth for the plants.

Misting serves as the primary controlling mechanism in aeroponics systems. It aids in regulating the nutritional solution's timing and concentration. Pressure pumps evenly apply a tiny mist to every plant to guarantee that it gets the right amount of water and nutrients. This approach conserves resources because neither over- nor under-provisioning occurs (Li *et al.*, 2018). Besides, since there is no soil, less possibility of pests and diseases spread through the soil is provided, which reduces the chances of health complications and completely rules out the application of toxic chemical treatments. Exposing plant roots to air is the main component of aeroponic systems. The level of oxygen surrounding the roots would be higher when roots are in contact with the atmosphere. This enhances growth and vigor. Root aeration prevents the roots from being anoxic or flooded -- two conditions that

can occur in soil-based or hydroponic systems (Tunio et al.,2021). Therefore, more oxygen availability enhances the faster rate of nutrient uptake and overall productivity, thus accelerating the metabolic processes of the plant.

In aeroponic systems, nutrient uptake is highly controlled so that the plants receive a blend of those trace elements and macronutrients in the most suitable proportion: potassium, phosphorus, and nitrogen. The formulation of the nutrient solution drives even greater control over the growth process depending on the specifics of the crops under cultivation. Versatility means aeroponics can be used as an efficient and easy method to grow varied crops in minimal amounts of water and nutrients.

Aeroponics differs from other forms of soil-based farming and even soilless cultivation methods like hydroponics. Since the soil provides a major supply of nutrients for traditional farming, high yields of land, water, and chemicals are required in order to produce crops at a rate that is adequate. In addition, this type of agriculture is susceptible to meteorological phenomena such as pest infestations, droughts, and degraded soil, all of which can adversely affect crop productivity. In contrast, aeroponics uses a fine mist of nutrient-rich water to hold the plant roots in the air without the need for soil (Blok *et al.*,2017). This approach dramatically reduces water usage because the mist hits the roots directly without causing excessive runoff or evaporation.

In hydroponic production, plants are grown in a nutrient-rich solution that continually flows over their root systems or is taken up by a growth medium, which can include such materials as perlite or coconut fiber. While hydroponics shares some similarities with aeroponics-directly related to the lack of soil and efficient delivery of nutrients-it essentially remains a system of submersion in water or a substrate for maintaining the root structure. With the use of aeroponics, the roots are left completely exposed to air and thus increase oxygen absorption with plants growing faster. This has thus made it possible for a system to come out faster and stronger compared to hydroponics; additionally, it uses less water and resources.

From a business perspective, aeroponics has many advantages over hydroponics and normal agriculture. The increased level of control over the growing environment allows for better nutrient delivery and reduces the requirement for chemical additives such as pesticides. Aeroponics systems also require much less physical space than traditional soil-based methods, making them very applicable to urban agriculture or land-scarce regions(Eldridge *et al.*,2020). Aeroponics offers a method of vertical farming, allowing for increased density in crop production in such small spaces to higher densities

compared with traditional methods. Even though hydroponics is inherently space efficient, aeroponics is even more compact due to the flexibility in layout and also in the absence of a growing medium.

In terms of resource efficiency, aeroponics is particularly noteworthy. Aeroponics goes one step further by spraying the roots and using only the amount of water required for plant growth, even though both hydroponics and aeroponics use less water than conventional farming. This method conserves water resources and also does not cause nutrient leaching—a prevalent problem that also exists within hydroponic and soil-based methods (Shrouk 2017). Second, the implementation of a closed-loop mechanism within an aeroponic system allows for the re-circulation of water and nutrients to aid in the reduction of waste and to further contribute to the long-term sustainability of the system. Lastly, the major differences between these agricultural methods truly emerge from a perspective involving the management of resources, control of resource use, and general scale-up possibilities. Especially in commercial applications, where time, space, and water are limiting factors, aeroponics is surprisingly efficient and very resource-frugal, especially as compared to both conventional soil-based cultivation and hydroponics.

3. Commercial applications of aeroponics

Aeroponics is rapidly finding various commercial uses in food production and agribusiness sectors, especially in the scarcity of then endemic high valued crops like leafy vegetables, herbs, and strawberry among others. This is because such crops are very much environmental condition dependant and thus require growth control and cultivation precision which aeroponics aims to achieve. For example, leafy greens draw a constant level of nutrients and growth moisture which can be supported by the misting systems of aeroponics (He et al., 2022). The end product is a healthier produce, with a higher content of nutrients and quicker growth – thus, this system is well-liked by commercial farmers who focus on quality and healthy products. In the same vein, culinary herbs, such as basil, mint, and cilantro, which are also produced subject to demand can be grown all-year that is without interference with season through aeroponics.

3.1 Urban Farming and Vertical Farming Initiatives

Aeroponics has also found its commercial application in one of them – urban agriculture especially vertical farming. In most urban centers, the scarcity of land presents a unique challenge in agricultural activities and aeroponic systems come in handy to efficiently utilize space. Vertical farms,

which are containment structures far away from the traditional farms or in some instances constructed within the skyscrapers, consist of overhead layers of crops grown on aeroponic systems, and this enables mass food production where there is limited space (Kodmany *et al.*, 2018, Linhares *et al.*, 2021). This method combats the very problem of the land issue for agriculture in cities as well as bringing in the fresh food supply to the customers hence cutting the costs of transportation and carbon emissions as well. Cities such as New York, Singapore and, Tokyo ventured into growing vertical farms featuring aeroponics and successfully established local food systems that are both ecologically and economically sustainable.

Businesses and communities all across the world are increasing food production via the use of a new technology called aeroponics (Lakkireddy *et al.*, 2012). Leading the way in commercial aeroponic farming in the US are businesses like AeroFarms and Plenty, who grow plants in vertical indoor farming structures and distribute the goods to retail chains and restaurants. More specifically, AeroFarms has established one of the biggest vertical farms in New Jersey wherein a variety of salads are cultivated by the use of aeroponically grown practices, knowing that they will grow fast and yield great amounts of crops with limited resources. Focusing on Europe, the Netherlands is one of those countries which is also practicing aeroponics as a means of staying ahead of the game in agriculture. Dutch growers have a reputation of being open-minded and resourceful and thus have embraced aeroponics in their quest to supply an ever-increasing volume of organic and green where possible foodstuffs.

In countries with less arable land but overpopulation, particularly in Asia, aeroponics is becoming increasingly popular as a solution to food shortages. Japan and Singapore are among the first countries to adopt this relatively new technique of growing food in urban environments using aeroponic systems. To further boost local food production, Singapore has commenced urban farming projects under the “30 by 30” policy that strives to grow 30% of the country’s edible products by the year 2030. The company ComCrop is building roof-top urban farms with a special emphasis on aeroponic systems growing herb and leafy greens to aid the food production efforts in the city-state (Wood *et al.*, 2020).

3.2 Pharmaceutical and Medicinal Plant Cultivation

Aeroponics has been developed as a good means of farming especially for crops that are considered to have high commercial values such as pharmaceutical and medicinal plants because it provides an environment within

the growth chambers that is conducive to growing plants that possess therapeutic purposes and are of high degrees of potency. This is quite different in traditional farming since the cultivation of medicinal plants has to deal with changes in soil and climatic conditions which sometimes lead to poor consistency in potency or low production. Growing plants in aeroponics system allows growers to play around with all parameters and variables in the growing process, which include light, temperature, humidity and nutrients amongst others, so that active compounds in the plants are allowed to reach their optimal levels every time without fail. This is even more important where plants like ginseng, echinacea and cannabis, which are expensive and grow for their active ingredients, alkaloids and cannabinoids, that depend on how accurate the farming conditions are done (Movahedi *et al.*,2020). This is very ideal for the pharmaceutical industry because such a degree of control leads to the production of plant material that is more refined and has less variation which is critical in the production of effective therapeutic agents.

Not only medicinal herbs but also a wide range of plants with commercial value are cultivated using aeroponics. For example, the supply of certain bioactive substances, namely flavonoids and glycosides, has provoked the interest of medicine companies into new ways of cultivating specific vegetation, such as the use of aeroponic system. These systems allow pharmaceutical companies to grow and harvest such plants more rapidly and with better yields of the target active ingredients while maintaining contaminant-free conditions for the plants. The sterile environment in the aeroponic system also minimizes the infestation of pests and diseases, which is very vital for the production of such medicine – any inception of disease or growth of undesirable organisms can render the product useless.

3.3 Biotechnology and Research Applications

The technology of aeroponics has also captured a sizable segment of the biotechnology and research industries, especially in the growing of plants for genetic engineering or genetic studies. Because it will allow the researcher to work in a sterile and controlled environment, it is the best platform for the researchers doing plant genetics, gene editing, or developing any new bioengineered plants. With aeroponics, scientists have this ability to regulate factors such as supply of nutrients, which eases the study of the plants by allowing an investigation into modified genes and the characteristics that the gene modulates. This system encourages the quick generation of transgenic tissues, which helps the researchers in the rapid turnover of their experiments and collection of data.

In terms of tissue culture and plant propagation techniques, aeroponics provides an excellent, if not the best, approach that allows plants to be grown in a more rapid and effective manner than the existing means of growing plants. The system involves the placing the plant roots in the air and spraying the nutrients in a misting form. This system enhances the root growth which in turn promotes the growth of the plant and helps in the process of propagation. Aeroponics is used in many biotechnology laboratories for developing plants when there is a need to carry out research that calls for a large number of similar plants. In tissue culture, which involves taking small parts of a plant and making them whole plants again, aeroponics helps to provide the best conditions aimed at ensuring cell growth and cell differentiation, thus being more effective than any old methods. This application is thus considered to be advantageous in the maintenance and propagation of even the most sensitive of plants and those whose growth is impossible to media such as soil.

3.4 IoT-Based Applications in Aeroponics

The integration of Internet of Things (IoT) technology into aeroponic systems has expanded their commercial applicability by supporting real-time monitoring, automation and data analysis. Within such structures, IoT sensors continuously measure various parameters, including temperature, humidity, nutrients and light. This information is then uploaded to cloud servers to allow the users to track and control the grow tent's conditions using a smartphone or a computer. IoT systems also include automated misting that enhances crop growth through precise amounts of nutrients to the plants and efficient use of resources. Predictive analytics also come in handy because they prevent potential losses in production by warning the growers of possible risks before they happen (Simanungkalit *et al.*, 2023).

In extensive undertakings like urban vertical farms, the use of IoT allows for the better management of crop growing in numerous layers with little physical involvement of the personnel. IoT along with climate and forecast models can be used in aeroponic farms to control the temperature and humidity levels indoors and promote growth of crops in all seasons (Guzmán *et al.*, 2022). The capacity to gather and process enormous amounts of growth data allows growers to improve their processes over time, hence boosting productivity and profitability.

3.5 Commercial Flower and Horticulture Industry

The concept of Aeroponics has become quite popular over the years with emphasis on the flower as well as the horticulture industry. Particularly, growing perfection for the conditions is key. This technique has enabled the

growing of quality ornamental flowers through a mist based delivery of nutrients. Growers are able to achieve which means they are able to tell the moisture, nutrient levels, and light – all leading to vivid, healthy flowers all year round (Sharma *et al.*, 2018). This system aids in constant nursery cycle allowing a steady supply of hard to find blooms like roses, tulips and lilies for horticulturalists and decorators. Crafted for enhancing floriculture production, aeroponics minimised the odds of fluctuations in flower dimensions, color variations and stem rigidity, making them win over their rivals(Wandl *et al.*, 2017).

3.6 Nurseries and Landscaping

In the context of horticulture and landscaping, aeroponic systems allow the cultivation of young plants or trees in an accelerated manner. The possibility of simulating perfect growth conditions irrespective of the quality of external soil or the existence of weather means stronger rooting in a shorter time flap. For example, landscaping can include vegetation of all kinds, be it decorative or robust trees, and they will be growing vegetatively when their planting becomes possible. Due to high efficiency of aeroponics, area requirements for soil growing are also lowered, which makes this method very popular in city nurseries or extensive landscaping works (Zobel *et al.*, 1976).

3.7 Sustainable Agricultural Demonstration

Sustainable agricultural exhibitions nowadays tend to focus more on the use of aeroponic systems, where water consumption is less than the normal and productivity is higher. These exhibitions are also designed for the people and the policymakers on why resources, waste, and crops health are better managed in aeroponics. As all elements of growth can be controlled by the farmer, it is, therefore, appropriate to present the system as a panacea for such problems as global warming, food insufficiency, and need for sustainable farming practices (Wimmerova *et al.*, 2022).

3.8 Space Exploration and Closed-Loop Systems

Employing aeroponics in the exploration of space is a recent development that revolves around its applicability. In order to alleviate the growing need for food in space missions, NASA and other space agencies are venturing into the use of aeroponic systems to cultivate foods in microgravity environments. These systems however, operate within very closed systems, that are of utmost importance for outer space travel due to the recycling of water and nutrients (Jurga *et al.*, 2018). Such systems are developed for the provision of clean, wholesome and healthy food to the astronauts while doing

away with wastage of life-sustaining resources. On the other hand, closed loop aeroponic systems are fast becoming popular within the dry regions of the Globe, as they present efficient means of agricultural practice with minimal resources, that is sustainable in nature (Salís *et al.*, 2020).

4. Economic implications of aeroponics

The viability of aeroponics as a sustainable agricultural technique lies in its resource use and management efficiency. While the costs of investing in basic infrastructure are usually high, it doesn't impede the adoption of this technique. Vertical farming or growing more crops in the same amount of land, as is in the case with aeroponics, makes it possible to optimize land-use reducing the areas under cultivation and minimizing the cost of land purchase. Also, because the system is designed in such a way that it reduces the cycle duration of crops, then within the same period, more crops can be grown hence more returns on the investment. The absence of a growing season also encourages multiple plantings within a year, leading to faster paybacks and better cash flow. Moreover, this practice makes it unnecessary to apply any pesticides, thus making chemical inputs and overall costs cheaper. This system and others also restrict loss of yield through damage to healthy and uniform crops, thus increasing usable produce and decreasing losses. In summary, all these reasons support the fact that aeroponics is affordable and feasible in the long term especially in regions with very little land, water and other resources or where they are expensive (Tokunaga *et al.*, 2015, Acheampong *et al.*, 2020, Babatunde *et al.*, 2021).

One of the most remarkable benefits of aeroponics in large-scale farming is effectiveness, more so in water and nutrient usage. It is known that aeroponic systems utilize significantly less water compared to the traditional soil-based agriculture penetrating up to 90% in some instances. This is particularly advantageous in areas with limited water supply enabling the farmers to grow crops while saving precious resources. Nutrient efficiency is again important— as the nutrients in the form of mist are supplied directly to the roots, there is very little wastage, which ultimately saves a lot of money in the long run. Further, the fact that plants can be grown in a vertical format or even in packed arrangements makes aeroponics incredibly space-efficient. More produce can be grown by farmers in a smaller area hence reducing the cost of land and increasing the yield per square foot.

From an energy consumption standpoint, aeroponics systems do require an active electricity source for components such as a misting system, lights, and monitoring sensors, but due to recent developments in technology, such

systems have become more energy efficient. For example, they can make use of smart sensors and IoT devices which can continuously monitor the health of the plants and the environment such that nutrients are supplied only when necessary and at the rate required thus eliminating or minimizing wastage of energy (Simanungkalit *et al.*, 2023). Moreover, energy costs are often compensated by the elevated and accelerated growth associated with aeroponics allowing growers to realize several output cycles in a short span unlike in conventional agriculture where such is not the case.

Looking at the return on investment and cost of operations, it should be noted that for aeroponics technology, the cost involved at the start is higher. For instance, one has to put in place the various elements, such as misting systems, environmental control, and nutrient supply systems as well as fitting several sensors in order to facilitate growth management. Despite this, the operational sustainability period is on the increase since the input sources including water, nutrients and labor are less than before. In particular, labor expenses tend to decrease because growing plants using aeroponics does not involve much physical work as in soil farming – the plants are automatically checked for their health, and nutrients are delivered to the plants only when required and for as long as they are needed without any manual input (Mehra *et al.*, 2018). The typical care and maintenance is mostly done by inspecting the different parts of the system and making sure the spray nozzles are clear of any clogs which is easier than managing an actual farm (Narasegowda *et al.*, 2020).

Aeroponics has yielded favorable results for many commercial growers in the areas of return on investment (ROI) and profitability. Perceptibly, the outset investment might be high but enhanced efficacies, which allows for quicker and more heightened production of high-value crops translates to faster returns. For example, given the existing market, aeroponic systems that take one year to set up can make huge profits in less than one year through the production of herbs and leafy greens whose growth periods are not long (Chittibomma *et al.*, 2023). Moreover, aeroponics offers the chance of production throughout the year, allowing the growers to have no worries over the seasons that limit their counterparts practicing subsistence farming, increasing profits even more.

5. Challenges and limitations in commercial aeroponics

5.1 Technical and Operational Challenges

In the case of commercial aeroponics, the actual maintenance and utilization of the systems poses real-life technical challenges. Messing systems

responsible for the optimum supply of nutrients and water at the root of the plants usually break down. Problems such as blocking, pump breakdown, or pressure inconsistency demoralize the entire process causing stress on the crop or immense losses. In this regard, any electrical pump fails for even a few seconds can cause great damage to the plants, as the system is designed to supply nutrients at all times. Besides, such systems have the risk of power blackout because they require constant electric supply in order to function, which means production may come to a halt or very expensive diesel generators have to be procured.

Continuous monitoring is another challenge that requires precise management of the environment—humidity, temperature, and nutrient concentration all need to be carefully regulated (Min *et al.*, 2022). A fluctuation in any of these can hinder plant growth, so advanced sensors and monitoring tools must be installed to maintain optimal conditions. But this infrastructure not only increases the initial setup cost, it also requires a highly skilled team to manage, troubleshoot, and maintain it. Training personnel to handle such complexity in large-scale operations can be a significant time and financial investment, especially for growers unfamiliar with these technologies.

5.2 Market Penetration Barriers

The steep upfront investment is one of the most substantial barriers to the broad adoption of aeroponics. Unlike traditional farming methods, which are more straightforward and lower in cost, aeroponic systems require significant capital to set up. The technology behind automated misting, sensors, and environmental controls doesn't come cheaply, making the financial entry point daunting for small to mid-sized farmers (Lakhia *et al.*, 2018). Even with the promise of long-term savings and higher yields, many growers hesitate, concerned about the substantial initial costs and the risks involved in shifting from traditional methods to high-tech solutions.

The other impediment is the technical know-how required for the use of aeroponic systems. These advanced systems include a lot of expertise transferrable knowledge on misting systems, environmental control systems, and nutrient controls, which many people in the agriculture sector are unaccustomed to. Training existing personnel or bringing in professionals can be expensive, thus deterring many from making the leap into a new technology. There is also the apprehension of breakdowns which may be too serious to manage and could result in crop losses if not managed in time. In regions where the access to such technical care or spare appliances is limited; this threat is more unbearable and obstructs the market growth of aeroponics.

5.3 Regulatory and Standardization Issues

Aeroponic as a form of farming is still in its infancy as it is young and cannot compete with farming, and the lack of ready-made rules and regulations is a major hindrance to its growth. Many nations do not have a well-informed prohibition related to the practice of aeroponics, this resulting to so much ambiguity on matters like food safety regulations, environmental safeguards, and sometimes even operational licenses. This sometimes creates an operational environment which is hostile to business expansion in most cases if not most because, such vagueness in regulation always promote delays or outright bar entry into the defined market. It is understandable that for most would-be users, this uncertainty encourages a new type of risk in a very risky proposition.

The absence of sector-wide specifications for aeroponic units serves to complicate things also. In the absence of defined, social designs of the system or operational performance criteria, different levels of efficiency tend to be the order of the day. This difference in ability tends to create difficulties for firms when it comes to maintaining efficiency and reliability, which is the key aspect of production when it comes to going large scale production. The biome in which they operate could be greatly improved with the implementation of such protocols, making it accessible for the novices and explaining why confidence in embracing the technology is possible. In addition, it would also be reassuring to the customers who will be assured that whoever grows their vegetables hydroponically/tower garden, the vegetables will always be of the same quality and safety level (Thomas *et al.*, 2023).

5.4 Limited Crop Variety

The restrictions on the type of crops that can be grown with the commercial aeroponic system are some of the serious issues faced in the production of commercial aeroponics. Even though one can market aeroponics claiming success with fast-growing high-turnover crops, like salads, herbs, some fruits among others, the same practice fails with many staple crops and bigger crops. This is because not every plant can cope with the method of spraying water and nutrients as a fine mist. This is because plants that require large root systems, or have high degrees of development in internal nutrition, are not well suited to the light, aerated systems of aeroponics. Tuber bearing plants such as potatoes or even tubers, and tall or heavy vegetation present in a farm maybe need deeper rooting which aeroponics cannot effectively offer without increasing costs in plant support (Yep *et al.*, 2019). Also, it still poses a

challenge for commercial growers to be able to optimize nutrient solutions for an extended list of plants. This in turn means that commercial growers will not only lack the necessary tools to grow certain crops, but will also be unable to grow these crops for other markets or grow them and continue in the same market.

5.5 Technological Complexity

Aeroponics has encountered a challenge in its commercial adoption owing to the system's business and operational complexity. Atmospheric pressure and moisture control, nutrient and water vapour delivery systems, and plant status indicators are all inbuilt in the design of an aeroponics system. Such technologically advanced systems are composed of an arrangement of various components, including sensors, atomist sprayers, air conditioning systems, and information integrators, which should all be brought together to achieve effective performance of the system (Garzón *et al.*, 2023). This is not easy for a lot of farmers especially, the inexperienced high tech due to the complexity of operation. This increases the divide between those who wish to effectively operate the system or cope with the system and those who will remain passive managers of the system. Ultimately, operational risk is elevated in addition to the curve in knowledge.

6. Current case studies and successful implementations

6.1 AeroFarms (United States)

AeroFarms, located in Newark, New Jersey, is a forerunner in the commercial application of vertical farming and aeroponics. Their 70,000sf facility is one of the world's largest indoor aeroponics farms designed to cultivate leafy greens like arugula, kale and watercress. It is operational throughout the year and can grow as much as 2 million pounds of greens every year. The aeroponic system they utilize delivers water and nutrients directly to the root zone in the form of a fine mist which saves water and also helps ensure that the plants get exactly what they need to grow their best. Unlike traditional farming, there's no soil, water usage is 95% lower, and pest control chemicals are not used in AeroFarms' way of farming as well. This has been made possible by the extensive use of real-time health monitoring of the plants through sensors and artificial intelligence to improve the conditions for growth. The facility's growing system is designed as a closed loop which allows the operator to produce higher yields in a much shorter growing period than conventional techniques, achieving up to 390 times output per square foot. By applying innovative aeroponic technology in combination with curated

processes, AeroFarms occupies an admirable position with respect to sustainable urban farming especially where land and resources are scarce (Howitt *et al.*, 2022).

6.2 Lettus Grow (United Kingdom)

Lettus Grow, based in Bristol, is an emerging company that has revolutionized the concept of aeroponics with its modular farm systems. Their aeroponic innovation comes in the form of “rolling beds”, which mist the roots of the plants with nutrients while spraying a carpet of water and nutrients to all the plants in the bed. Their systems offer high level of customization therefore they can fit objects of many sizes from lettuces grown in hydroponics to strawberries grown in soil (Estuita *et al.*, 2022). LettUs Grow’s technology enhances aeration of the root zone which plays a vital role in plant growth promoting faster growth as compared to traditional soilless systems. The company claims that its systems can achieve up to 98% less water usage and man less energy costs due to the accuracy of how the nutrients are delivered. Such efficiency, in addition to the mobility of systems makes it easy to adopt in areas also known as cities where space is limited. The startup has also collaborated with urban farmers, restaurants, and grocery who wish to sell locally grown food, healthy produce which does not require long transport. The growing conditions on site can also be remotely supervised and controlled using their integrated software system, thus minimizing the need for personnel to be on site and cutting down operational costs.

6.3 Infarm (Germany)

With its headquarters in Berlin, Infarm innovatively proposes the idea of integrating aeroponic farming systems into public spaces like restaurants and supermarkets. They are horizontal compact systems within an enclosed farm that employs mist irrigation control for delivering nutrients at the root. Infarm's aeroponic farming technology allows the plants to grow up to three times their growth potential, using only 5% of the water resource, and occupying less than 10% of the land. Infarm grows and sells herbs, micro veggies, and kales from the nearest geographical direction in order to let their clients enjoy the freshest products possible. The system from Infarm is very much modular and adaptable because the vertical farming systems can be stacked in various arrangements and also positioned in structures as big as hypermarkets to as small as fast food joints. A cloud based application has also been developed where growth parameters and environmental aspects of the plants in question are managed from and for a distance. Providing these methods of control in all aspects to plant growth management is real time and focuses on light,

humidity, nutrients, and even plant growth alternates. Infarm, having made a rather successful and aggressive entry in the European market, signing such contracts with holdings like Marks & Spencer, Carrefour and EDEKA (Cointet *et al.*, 2019).

6.4 Pinduoduo Smart Greenhouses (China)

Pinduoduo, one of the top e-commerce platforms in China, has contacted smart agriculture technology by building and operating commercial-sized aeroponic greenhouses. The developed system includes sophisticated Internet of Things sensors and artificial intelligence combined with data analytics for real time control of the growing conditions. The system specifically targets controlled-environment agriculture for high-valued crops such as leafy vegetables and herbs which are in high demand in urban areas of China. Pinduoduo's smart greenhouses combine misting technology with AI algorithms that adjust temperature, humidity, and nutrient levels to ensure optimal plant growth. Their aeroponic method allows crops to grow faster while using 90% less water than traditional farming, and the controlled environment means there is no need for chemical pesticides. Pinduoduo's initiative is part of a broader push toward modernizing China's agricultural sector and improving food security in densely populated urban areas. By integrating aeroponics with e-commerce, Pinduoduo aims to streamline the supply chain, bringing fresh produce from farm to consumer more efficiently. This initiative also focuses on reducing the environmental impact of agriculture, which is crucial given China's limited arable land and water resources. These companies represent the cutting edge of aeroponics and urban agriculture, each adapting the technology to different scales, crops, and markets while demonstrating the versatility and sustainability of aeroponic systems in modern farming (Chang *et al.*, 2019).

6.5 Sky Greens (Singapore)

Sky Greens from Singapore is yet another example that successfully combines vertical farming and aeroponics. Agriculture in Singapore takes up only a limited portion of the land and so measures such as aeroponic farming have been embraced in order to solve food integrity issues. The company operates the world's first low-carbon vertical farm which is hydraulic driven utilizing the aeroponic method of growing vegetables in rotating stackable towers. Such towers enhance space and allow for rapid urban crop production.

The system in place at Sky Greens uses 95 percent less water than conventional farming, conserves energy by relying on sunlight and uses renewable energy to power vertical tower rotation. The use of aeroponics

means that the plants receive nutrients in a gaseous form, which results in quicker growth rates and greater crops within a smaller area(Zhao et al., 2024). Looking around East Asia, much has been provided in the way of administrative support for the creation of salad bars and for sections of sited supermarkets dedicated to fresh vegetable produce including III phase systems. Such out-migration of a sizeable domestic market for traditional fresh produce has been beneficial to many surfaces who focus on fresh vegetables as a core business. Furthermore, most especially in that space around the entire civilization developing technology models entails some serious deviation from the usual practices in agriculture.

6.6 Sundrop Farms (Australia)

Sundrop Farms, situated in Australia's southern region, is a pioneer in the commercial usage of aeroponics in a renewable energy setting. By combining solar power, desalination and aeroponics in a semi-arid desert climate, Sundrop Farms has developed a sustainable closed-loop farming system. This allows the growing of crops that otherwise require extensive vegetation cover, such as tomato, with very little water and without root in any soil; hence less dependence on conventional agricultural processes and agricultural chemicals such as freshwater and pesticide. Instead of relying on rainfall, Sundrop Farms grows fresh crops all year round using geothermal energy with the unique capacity of vertical farming. Cultivating more than 15,000 tons of tomatoes a year while consuming less than conventional farming 90 above less freshwater, the farm has proved that large scale commercial production using aeroponics is feasible especially in water scarce regions. Sundrop Farms has proven how modern technology can provide solutions to food insecurity particularly in harsh climatic conditions by becoming the leader in sustainable developments in agriculture (Paul,2015).

7. Conclusion

Aeroponics has the potential to revolutionize the practice of farming for it provides solutions to problems brought about by urbanization, scarcity of resources and environmental conservation. It is a method of growing plants in air with mist delivered to the roots of the plants without using any soil, thus saving water and doing away with the use of pesticides. Aeroponics systems can be adapted in every possible condition from large scale commercially productive farms to small urban spaces and grow food in places that are not suitable for conventional soil-based farming. Pioneering companies like AeroFarms and LettUs Grow deploy such systems, incorporating cutting-edge

technologies, including AI and modular systems, to improve the environmental conditions and productivity.

With the increasing preference for sourcing produce locally, what the pilot projects also show is how Potager System, for instance, employs aeroponics, which can be optimistic for food systems. Projects like Pinduoduo's smart greenhouses and Sundrop Farms' solar-powered operations showcase that this kind of agriculture can be successful even in the hostile climates. Combining sustainable solutions with technologies where aeroponics alone ensures that there is food security but also that there is diversification of agricultural practices. The outlook on food production is bright, and with the increasing awareness and acceptance of the systems, the role of aeroponics systems in the food supply chain will be more pronounced, more so in the sustainable and effective chain.

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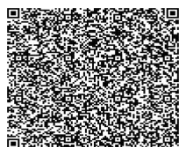
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Emerging Research Concept in Life Sciences

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