

BIOSYNTHESIZE NANOPARTICLES AS ADVANCED NANOCARRIERS FOR TARGETED THERAPEUTIC INTERVENTIONS IN DISEASE MANAGEMENT

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Abstract: *Biosynthesized nanoparticles have emerged as cutting-edge nanocarriers for targeted therapeutic interventions in illness management, transforming medication delivery and treatment techniques. These nanoparticles provide a biocompatible and sustainable substitute for conventional synthetic processes since they are made utilizing biological systems including fungi, bacteria, and plants. They are perfect candidates for precision medicine because of their innate capacity to lower toxicity and increase bioavailability. Because of their distinct surface characteristics, biosynthesized nanoparticles are easily functionalized with therapeutic drugs, allowing for targeted distribution to sick cells or certain organs while reducing off-target effects. Additionally, the large surface area-to-volume ratio and nanoscale size promote better cellular absorption and regulated drug release, enhancing therapeutic efficacy. These nanoparticles' natural origin lowers their immunogenicity and adverse response risk, which enhances patient outcomes even more. Biosynthesized nanoparticles have demonstrated great promise in cancer therapy, including the ability to target tumor cells specifically, carry chemotherapy medications, and even combine with imaging agents for theragnostic uses. They also show promise in improving therapy specificity and lowering adverse effects in other diseases like infectious diseases, cardiovascular ailments, and neurological disorders. This study covers the developments in biosynthesized nanoparticles as nanocarriers, highlighting their importance in altering illness management through tailored therapeutics while evaluating the limitations and future prospects for their clinical translation.*

Keywords: Biosynthesized Nanoparticles, Advanced Nanocarriers, Controlled Release Systems, Biocompatibility, Therapeutic Interventions, Disease Management, Biogenic Nanoparticles, Tumor Targeting, Drug Encapsulation

Introduction

Biological entities like plants, bacteria, fungus, or algae are used to manufacture biosynthesized nanoparticles, also known as biogenic or green nanoparticles. In contrast to traditional chemical or physical techniques, which frequently call for expensive equipment, strong chemicals, or high temperatures, biosynthesis makes use of natural biological processes to produce nanoparticles (Pandit et al., 2022). This green chemistry-based ecologically friendly method decreases the production of harmful byproducts and lessens the demand for toxic solvents. Biosynthesized nanoparticles are extremely attractive for applications in medicine, agriculture, and environmental remediation because they have various benefits, such as better biocompatibility, decreased toxicity, and cost-effectiveness (Ahmed et al., 2022). The manufacture of nanoparticles using green chemistry methods makes use of natural reductants, stabilizers, and capping agents found in biological systems. For example, phytochemicals such as terpenoids and flavonoids, which are abundant in plant extracts, can both stabilize and decrease metal ions to produce nanoparticles (Ovais et al., 2018). This helps to

achieve sustainable development goals and lessens the environmental impact of producing nanoparticles. Biosynthesized nanoparticles are a possible substitute for traditionally synthesized nanomaterials because they frequently show improved surface functionalization, which may be tuned for particular purposes including medication delivery, antibacterial activity, and pollution removal (Moritz et al., 2013).

A technique known as "green synthesis," which produces nanoparticles in an environmentally acceptable and sustainable manner, depends heavily on biological processes, including bacteria, fungus, plants, and algae (Khan et al., 2022). In contrast to traditional chemical and physical techniques, which frequently call for dangerous ingredients and harsh environments, biological synthesis uses an organism's inherent metabolic processes to turn metal ions into nanoparticles. For instance, a large range of phytochemicals produced by plants, such as phenolic compounds, terpenoids, and flavonoids, function as stabilizing and reducing agents during the creation of nanoparticles (Yousaf et al., 2018). Because of their great versatility and low growth needs, bacteria may also operate

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as biofactories. They can do this by using their proteins and enzymes to catalyze the reduction of metal ions, such as gold or silver, into nanoparticles (Shankar et al., 2016). Fungi are very good at mass-producing stable nanoparticles because of their strong cell wall structures and capacity to produce a lot of extracellular enzymes. Because they grow quickly and contain bioactive substances that aid in the creation of nanoparticles, algae both macro and microalgae have demonstrated great promise in this field as well (Vargas-Estrada et al., 2020). These biological systems yield nanoparticles with distinct sizes, shapes, and properties that can be customized for a range of applications in agriculture, medicine, and environmental remediation. They also offer an environmentally friendly substitute for conventional nanoparticle synthesis methods (Mao et al., 2007). In the biosynthesis of nanoparticles, sustainability and environmental friendliness have attracted a lot of interest as alternatives to conventional chemical and physical processes, which frequently entail hazardous byproducts, excessive energy consumption, and toxic reagents (Kumar et al., 2021). The use of natural organisms such as plants, bacteria, fungus, and algae to manufacture biosynthesized nanoparticles (NPs) is a more environmentally friendly method that is in line with the ideas of sustainable development and green chemistry (Khan et al., 2022). This environmentally beneficial technique, which frequently takes place in moderate settings without the use of dangerous chemicals, makes use of the biological machinery of organisms to convert metal ions into nanoparticles. For instance, the use of plant extracts introduces bioactive substances that function as natural reducing and stabilizing agents, such as proteins, enzymes, and phytochemicals, obviating the need for harmful substances from the outside (Shafey et al., 2020). By adding biocompatibility and improved functionality, this biogenic method not only lessens the environmental impact of the nanoparticles but also makes them more suited for medication administration, environmental cleanup, and biomedical applications (Das et al., 2021). Furthermore, compared to conventional processes, the production of biosynthesized nanoparticles frequently occurs at room temperature and pressure, which dramatically reduces energy usage. A circular economy may be achieved through these environmentally friendly methods since they are scalable and have an abundance of biological resources that

can be used again. Biosynthesized nanoparticles stand out as a crucial invention in improving sustainable nanotechnology at a time when environmental safety is a top priority for both companies and researchers (Bharti et al., 2022). The aim is to create effective and biocompatible drug delivery systems using nanoparticles that can more precisely target specific diseased areas, therefore enhancing treatment results and reducing adverse effects. Furthermore, the project aims to explore how biosynthesized nanoparticles might help overcome traditional therapy barriers such as low bioavailability and drug resistance.

Mechanisms of Biosynthesis; Insights from Nature

The biosynthesis of nanoparticles is an intriguing process that offers important lessons from nature. It reveals intricate biological processes that help different species create nanoparticles. Metal ions from the soil are usually taken up by plants during the formation of nanoparticles (Haverkamp et al., 2009). Plant metabolites, such as proteins, enzymes, and secondary metabolites, subsequently decrease and stabilize the metal ions. Benefits of this plant-mediated method include reduced toxicity, biocompatibility, and the capacity to create nanoparticles with particular morphological and chemical characteristics suited to the intended use. On the other hand, the enzymatic mechanisms that define microbial pathways especially those involving bacteria and fungi are essential to the reduction of metal ions into nanoparticles (Ameen et al., 2021). Microorganisms generally contain distinct metabolic pathways that enable them to decrease metal ions extracellularly or intracellularly, with diverse enzymes like reductases and oxidases being crucial in these activities. These biosynthetic pathways differ significantly in terms of yield, efficiency, and the kinds of nanoparticles that are synthesized (Li et al., 2011). Microbial biosynthesis can produce nanoparticles more quickly and with a wider variety of sizes and morphologies than plant-mediated biosynthesis, which is frequently slower and may produce lower quantities of nanoparticles. Knowing these biological mechanisms helps to clarify the natural processes that lead to the formation of nanoparticles and provides opportunities for environmentally friendly and sustainable methods of synthesising nanoparticles that can be used in materials science, environmental remediation, and medical fields (Albrecht et al., 2006).

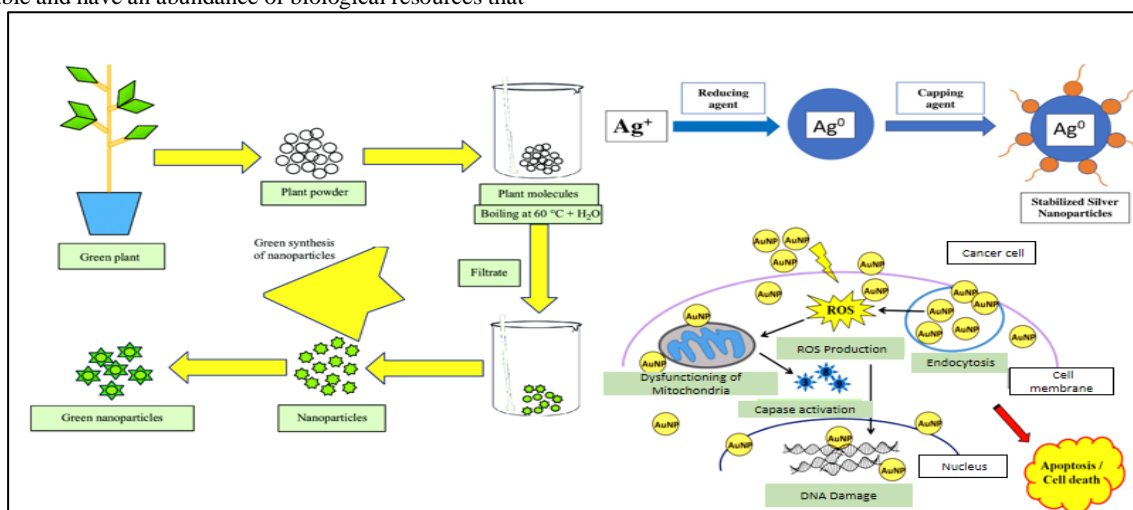


Fig 1: Mechanisms of Biosynthesis; Insights from Nature

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Therapeutic Applications in Cancer Treatment Nanoparticle Characterization Techniques

Understanding the physical and chemical characteristics of biosynthesized nanoparticles is essential as these attributes directly affect their effectiveness in biomedical applications. To investigate important features including size, shape, morphology, and surface chemistry, a variety of methodologies are used (Albanese et al., 2012). One of the most sophisticated techniques for characterizing materials is X-ray diffraction (XRD), which helps identify the crystalline structure and phase composition of nanoparticles, revealing information about their purity and crystallinity. Researchers can see the form and size distribution of nanoparticles by using high-resolution imaging of surface morphology made possible by scanning electron microscopy, or SEM. This is enhanced by Transmission Electron Microscopy (TEM), which provides information on interior structures and atomic-level details on the size, shape, and aggregation of nanoparticles (Kim et

al., 2018). To analyze the surface chemistry of nanoparticles and determine the functional groups and bonding interactions that are crucial for their reactivity and biocompatibility, Fourier Transform Infrared Spectroscopy, or FTIR, is a necessary tool. Particle size distributions in solution are commonly evaluated using Dynamic Light Scattering (DLS), which yields hydrodynamic size data that is crucial for comprehending the behavior of nanoparticles in biological settings (Rahtar et al., 2019). Scaling up biosynthesized nanoparticles for therapeutic purposes remains challenging despite the progress made in these characterization approaches. Large-scale manufacturing is hampered by problems including resolving possible toxicity issues, guaranteeing repeatability, and preserving size and shape consistency throughout synthesis. To successfully overcome these obstacles and enable the seamless integration of biosynthesized nanoparticles into clinical practice, ongoing advancements in synthesis techniques and characterization processes are needed (Lim et al., 2015).

Table 1: Overview of Nanoparticle Characterization Techniques

Characterization Technique	Description	Key Advantages	Limitations	Applications
Thermogravimetric Analysis (TGA)	Measures weight changes as a function of temperature to analyze thermal stability and composition.	Provides thermal stability information Can analyze degradation behavior	Requires solid samples Limited to thermal characteristics	Stability assessments Composition analysis
UV-Vis Spectroscopy	Monitors the absorption of UV-visible light to identify the electronic transitions in nanoparticles.	Non-destructive A simple and quick analysis	Limited to optically active materials Cannot provide size or shape information	Optical characterization Concentration determination
Atomic Force Microscopy (AFM)	Uses a cantilever with a sharp tip to measure surface forces and generate high-resolution topographical maps.	Provides 3D surface profiles Can operate in various environments	Slow scanning speed Requires conductive or insulating substrates	Surface roughness analysis Nanostructure imaging
Dynamic Light Scattering (DLS)	Measures the Brownian motion of particles in suspension to determine hydrodynamic size distributions.	Quick and easy measurements Analyzes particle size in solution	Sensitive to concentration and refractive index Limited to submicron sizes	Size distribution analysis Stability assessments
Fourier Transform Infrared Spectroscopy (FTIR)	Analyzes the infrared absorption spectra to identify functional groups and chemical bonds in nanoparticles.	Provides information on surface chemistry Non-destructive analysis	Limited to surface analysis Requires interpretation of spectra	Functional group identification Interaction studies
Transmission Electron Microscopy (TEM)	Uses electron beams to transmit through thin samples, providing detailed images of internal structures and sizes.	Atomic-level resolution Reveals internal structures	Requires extensive sample preparation High cost	Size and shape analysis Aggregation studies
X-ray Diffraction (XRD)	A technique used to determine the crystalline structure of nanoparticles by analyzing the diffraction patterns of X-rays.	Identifies crystalline phases Assesses purity and crystallinity	Requires crystalline samples Limited to bulk characterization	Material identification Phase analysis

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Therapeutic Applications in Cancer Treatment

Biosynthesized nanoparticles have become a viable therapeutic approach in the treatment of cancer, especially when it comes to improving the delivery accuracy of radiation and chemotherapy. These biocompatible and low-toxicity nanoparticles, which are made from biological systems like fungus, bacteria, or plants, are perfect for use in medical applications (Rónavári et al., 2021). Biosynthesized nanoparticles reduce harm to healthy tissues and minimize adverse effects by acting as drug carriers in chemotherapy, improving the precision of tumor targeting. For example, these nanoparticles improve medication solubility and enable regulated, sustained release of chemotherapeutic drugs directly at the tumor site by encasing them (Mattheolabakis et al., 2012). In a similar vein, biosynthesized nanoparticles can serve as radiosensitizers in radiotherapy, increasing the efficacy of radiation doses to cancer cells while shielding neighboring

healthy cells from unintentional harm. The potential of biosynthesized nanoparticles to improve treatment results for cancer has been shown in clinical studies. Studies on patients with breast and lung cancer have demonstrated considerable tumor suppression with less toxicity when gold and silver nanoparticles synthesized from plant extracts were used. Furthermore, to further optimize treatment regimens, biosynthesized nanoparticles allow the integration of multimodal treatments, such as combining radiation and chemotherapy. As research advances, these nanoparticles might revolutionize oncology by providing promise for more individualized, efficient, and safe cancer treatments (Kemp et al., 2021).

Biosynthesis of Nanoparticles (Using biological systems to produce biocompatible nanoparticles) Drug Encapsulation in Nanoparticles (Chemotherapy) Nanoparticles as Radiosensitizers (Radiotherapy) Improved Treatment Efficacy Case Studies from Clinical Trials.

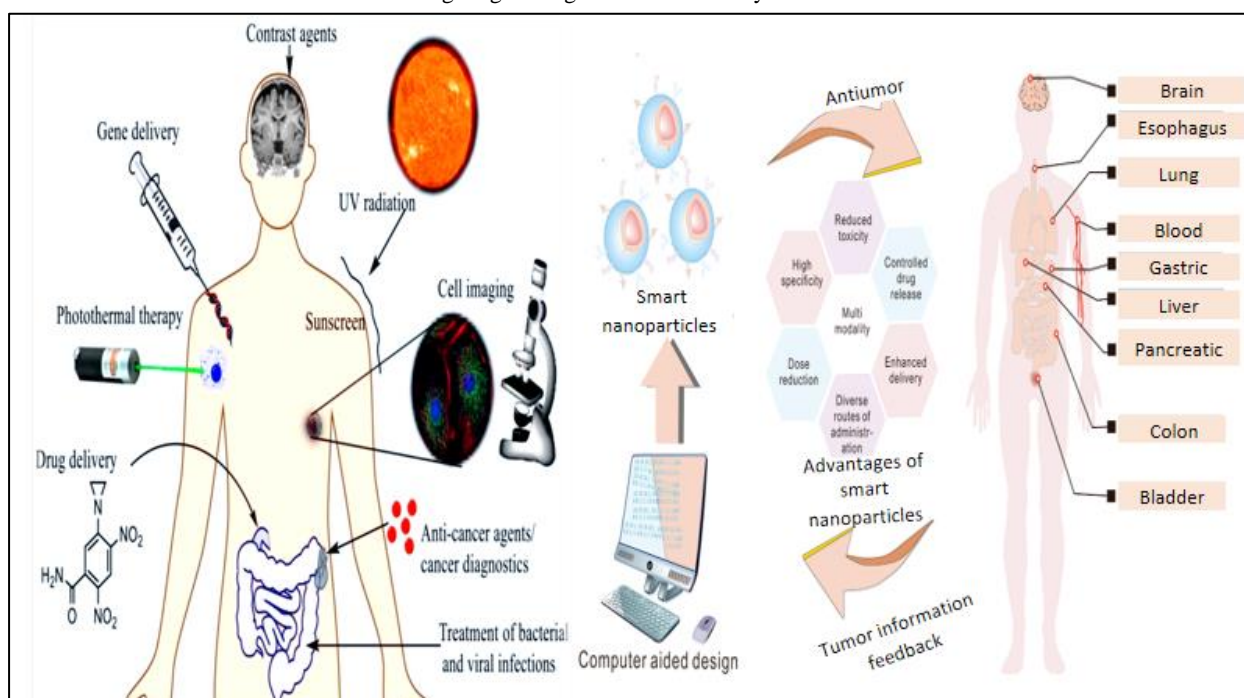


Fig 2: Therapeutic Applications in Cancer Treatment

Applications in Antimicrobial and Antiviral Therapy

Nanoparticles that are biosynthesized have shown great promise in antiviral and antibacterial treatment because of their diverse characteristics and multifunctional capacities. These biological processes involving fungus, bacteria, and plants provide nanoparticles that can be useful transporters of antiviral and antibiotic drugs (Stan et al., 2021). These medications can be stabilized, made more bioavailable, and delivered more precisely by encasing them in nanoparticles, which enables controlled release at infection sites. Additionally, the inherent antibacterial and antiviral qualities of biosynthesized NPs can work in concert to increase the potency of the medications they carry. They are very useful in the fight against difficult viral infections and bacteria that are resistant to drugs due to their dual action. The disruption of microbial cell membranes, the production

of reactive oxygen species (ROS), and the binding of NPs to microbial DNA or proteins, which inhibit vital biological activities, are the mechanisms by which NPs improve antimicrobial action (Nisar et al., 2019). Biosynthesized NPs can obstruct viral entrance, replication, and fusion with host cells in viral infections. The utilization of biosynthesized nanoparticles is a noteworthy approach to surmount conventional drug resistance since it circumvents conventional resistance mechanisms including enzymatic degradation and efflux pumps. Since viruses and bacteria that are resistant to drugs continue to be major dangers to global health, the use of biosynthesized nanoparticles offers a viable path toward the development of stronger antiviral and antibacterial treatments, which might completely change the way infections are treated (Gurunathan et al., 2020).

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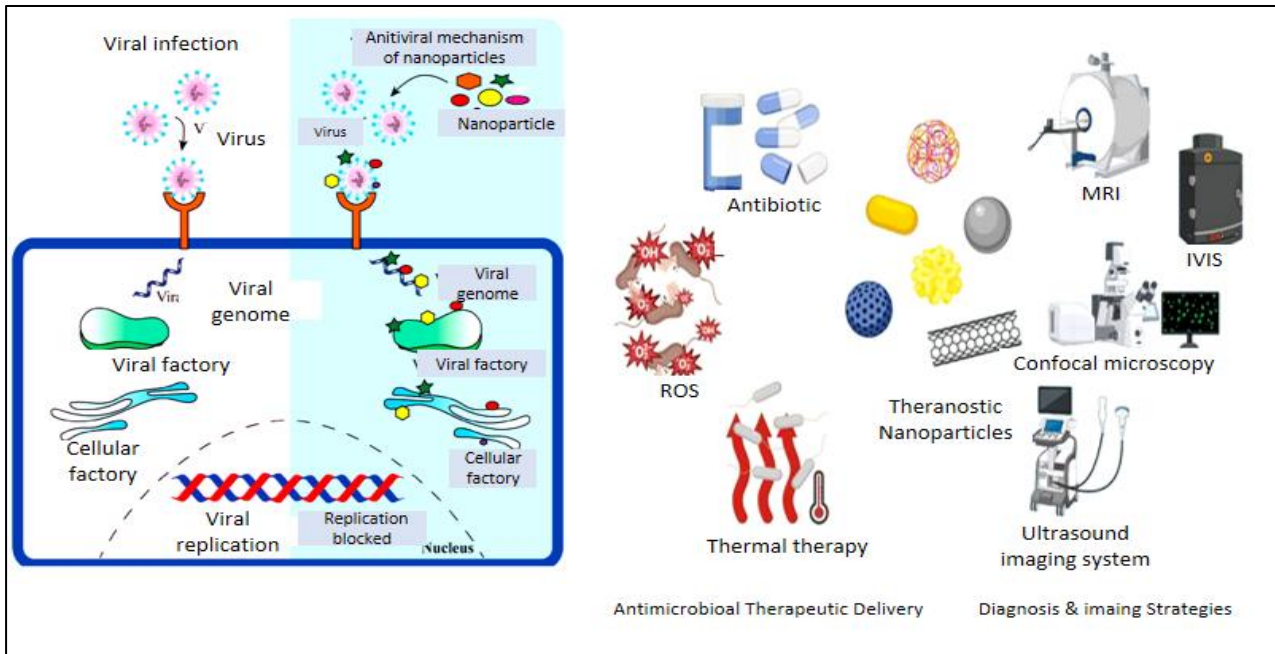


Fig 3: Applications in Antimicrobial and Antiviral Therapy

Cardiovascular Therapeutics: Novel Applications

The development of biosynthesized nanoparticles, which provide tailored therapy options for a variety of cardiovascular illnesses, has significantly advanced cardiovascular therapies. These biologically sourced nanoparticles, which include decreased toxicity, biocompatibility, and the capacity to cross biological barriers, are perfect for the delivery of cardiovascular medications because of their special qualities (Pala et al., 2020). Nanocarriers, in particular, have enhanced therapeutic effectiveness, minimized systemic adverse effects, and guaranteed accurate delivery to afflicted regions, revolutionizing the administration of anti-hypertensive and anti-thrombosis drugs. More regulated therapy is possible using nanocarriers because they may be made to release medication in response to certain

cardiovascular system signals, such as pH or enzyme activity. Vascular tissue engineering is a promising field in which biosynthesized nanoparticles are being investigated for their potential to promote the repair of injured blood arteries (Hosseini et al., 2020). These nanoparticles can promote the repair and remodeling of vascular tissues by acting as scaffolds or by delivering growth factors and other therapeutic agents. New developments in this area indicate great promise for bettering patient outcomes in diseases where conventional treatments frequently fall short, such as heart failure, hypertension, and atherosclerosis. Because biosynthesized nanoparticles may combine tissue engineering and therapeutic delivery, they open up new possibilities for cardiovascular treatment (Fathi-Achachelouei et al., 2019).

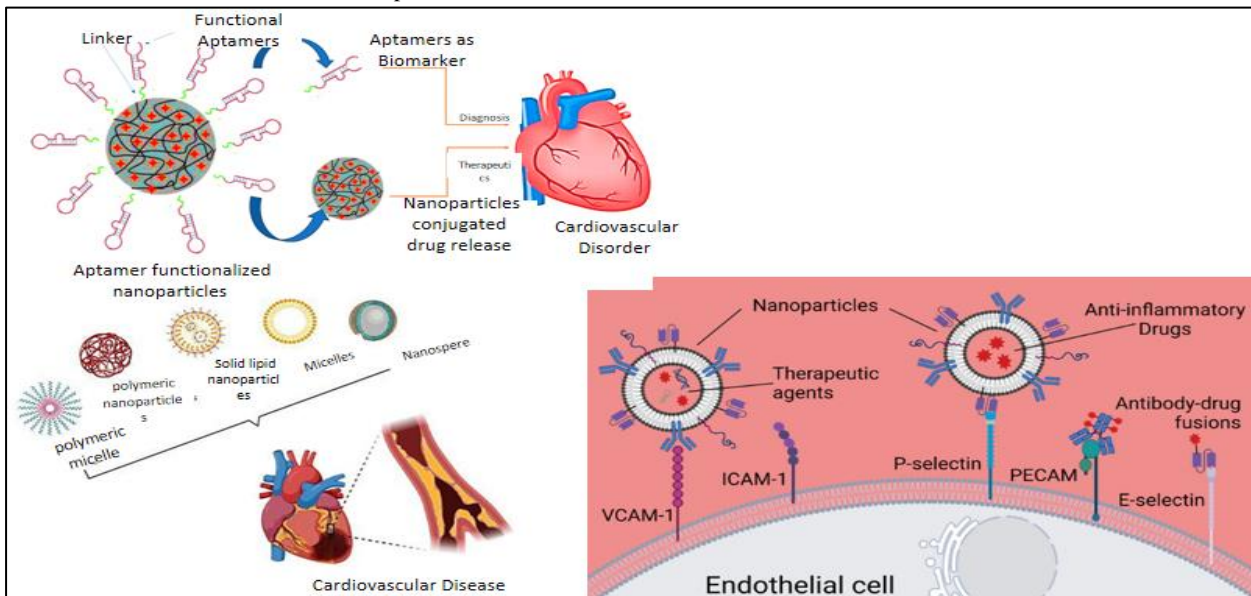


Fig 4: Cardiovascular Therapeutics: Novel Applications

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Biosynthesized Nanoparticles in Gene Therapy

When it comes to gene therapy, biosynthesized nanoparticles are showing promise, especially when it comes to delivering gene-editing tools like CRISPR/Cas9. Because of their size, biocompatibility, capacity to modify the surface, and resistance to immune responses, biosynthesized nanoparticles are perfect vehicles for delivering CRISPR/Cas9 components to specific cells and tissues (Xu et al., 2021). These nanoparticles are less hazardous for use in therapeutic settings since they were created via biological processes, such as the synthesis of bacteria, fungus, or plant extracts. Gene therapy involves the precise editing of faulty genes that cause genetic illnesses and malignancies by introducing DNA, RNA, or CRISPR-associated proteins into cells using biosynthesized nanoparticles. Their capacity to cross cellular membranes and transport gene-editing instruments straight to the nucleus improves the effectiveness of gene editing and

raises optimism for the treatment of diseases including muscular dystrophy, cystic fibrosis, and inherited malignancies (Grunwald et al., 2020). The widespread use of biosynthesized nanoparticles in gene therapy is still fraught with difficulties, notwithstanding recent advancements. Controlling the stability, biodistribution, and release of tools for gene editing are still important issues. Another challenge that has to be overcome is scaling up the manufacturing of biosynthesized nanoparticles for consistent quality and efficacy in clinical settings. Future studies will probably concentrate on lowering off-target effects, strengthening the stability of CRISPR/Cas9 systems in the circulation, and improving nanoparticle characteristics to increase targeting specificity. Through the use of biosynthesized nanoparticle-mediated gene therapy, these efforts may lead to extremely effective, minimally intrusive therapies for genetic disorders and malignancies (Gavas et al., 2021).

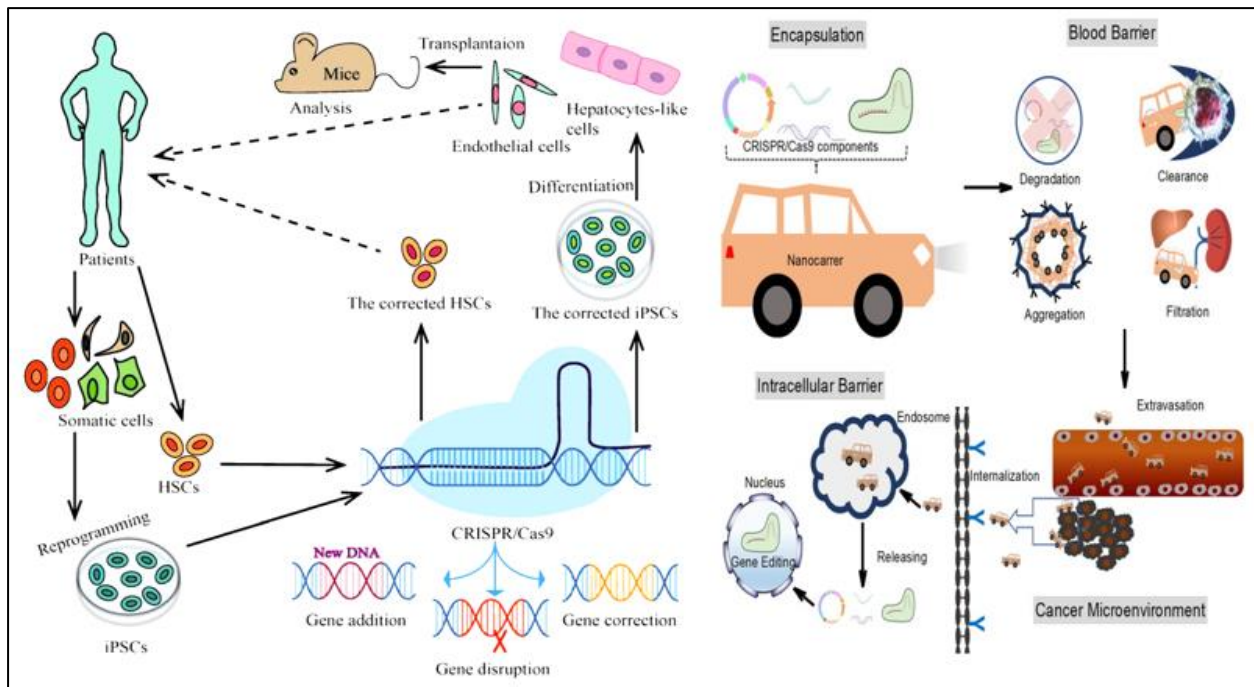


Fig 5: Biosynthesized Nanoparticles in Gene Therapy

Nanoparticle-Based Vaccines: The Future of Immunization

As a revolutionary method of vaccination, vaccines based on nanoparticles are emerging as the direction of vaccine technology. The utilization of biosynthesized nanoparticles, which are created by biological systems like bacteria, fungi, or plants, is one of the most promising advances in this sector (Koul et al., 2021). The great biocompatibility of these biosynthesized nanoparticles lowers the possibility of unfavorable responses, and their nanoscale size enables targeted antigen delivery to the immune system. They are being praised for their capacity to improve antigen presentation and delivery efficiency, which can greatly boost immune responses. In contrast to conventional vaccinations, systems based on nanoparticles may be designed to more closely resemble infections, eliciting a strong and focused immune response at lower dosages

(Gregory et al., 2013). Furthermore, nanoparticles can be engineered to contain many adjuvants or antigens, increasing their immunogenicity and providing broader disease protection. Because of this, they are extremely useful in the creation of vaccinations for future generations, especially when it comes to pandemic preparation. Because nanoparticle-based systems may be readily adapted to new pathogens such as viruses and bacteria, they provide the benefit of expedited vaccine development. Given their capacity for large-scale manufacturing and their ability to stabilize and preserve antigens, they may be essential in helping societies respond quickly and effectively to pandemics in the future. Biosynthesized nanoparticles have the potential to completely change the vaccination landscape by making it possible to create vaccines that are more effective, long-lasting, and readily adjustable (Bezbaruah et al., 2022).

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Table 2: Nanoparticle-Based Vaccines

Category	Description	Examples	Key Benefits	Applications
Biosynthesized Nanoparticles	Nanoparticles are engineered using biological sources, such as bacteria, fungi, and plants, which offer biocompatibility and reduced toxicity.	Plant-based nanoparticles, bacterial-derived nanoparticles	Biocompatibility, reduced risk of adverse effects, environmentally friendly production.	Used in the creation of vaccines with reduced allergenic or toxic side effects, suitable for populations with compromised immunity.
Antigen Presentation	Nanoparticles can be engineered to mimic pathogens, allowing for precise and effective antigen presentation to immune cells.	Virus-like particles (VLPs), metallic nanoparticles	Increased antigen uptake by dendritic cells, improved activation of T-cells and B-cells.	Essential for diseases where immune evasion is a concern, such as tuberculosis and certain cancers.
Stability and Shelf Life	Nanoparticles provide stability to vaccines, protecting them from degradation and extending shelf life, which is crucial for global distribution.	Nanocapsules, dendrimer-based nanoparticles	Increased shelf life, resistance to environmental degradation, and fewer cold-chain requirements.	Important for vaccines distributed in regions with limited refrigeration and infrastructure, aiding global immunization efforts.
Multiple Antigen Delivery	Nanoparticles can carry multiple antigens and adjuvants simultaneously, providing broader protection against multiple strains or types of pathogens.	Gold nanoparticles with multiple antigen-loading capabilities, hybrid nanoparticles	Broader immunity coverage, improved vaccine efficacy in complex diseases.	Used in vaccines for multi-strain viruses like influenza and for pathogens with high mutation rates such as the coronavirus.
Adjuvant Capabilities	Nanoparticles can serve as adjuvants themselves or be used to deliver traditional adjuvants, boosting the immune system's response to the antigen.	Aluminum-based adjuvant nanoparticles, silica nanoparticles	Enhanced immune activation, reduced need for synthetic adjuvants, and lowered risk of adverse reactions.	Applied in vaccines for difficult-to-immunize populations such as the elderly or immunocompromised individuals.
Reduced Dosage Requirements	Due to the efficient delivery and enhanced immune response of nanoparticle-based vaccines, lower dosages are required compared to traditional vaccines.	Polymeric nanoparticles, metal oxide nanoparticles	Lower doses reduce the risk of side effects, cut down costs, and improve accessibility for low-resource settings.	Particularly important in vaccines where repeated doses are required (e.g., Hepatitis B) or where there are supply constraints.
Personalized Vaccine Development	Nanoparticles can be tailored for personalized vaccine development, adapting vaccines to individual patient profiles for enhanced efficacy.	Peptide-conjugated nanoparticles, mRNA-loaded nanoparticles	Increased vaccine efficacy, reduced risk of adverse reactions, and optimized immune responses tailored to the patient's immune system.	Critical for cancer vaccines and other conditions requiring highly individualized treatment, providing a path to precision medicine in vaccination.
Environmental Considerations	Biosynthesized nanoparticles reduce the environmental impact of vaccine production by using sustainable and eco-friendly materials.	Plant-based nanoparticles, biodegradable polymer nanoparticles	Environmentally sustainable production methods reduce toxic waste and less reliance on synthetic chemicals.	Increasingly relevant for global health organizations aiming to lower the environmental footprint of vaccine development and production.

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Biosynthesized Nanoparticles in Regenerative Medicine

Since they are biocompatible, have low toxicity, and are simple to functionalize, biosynthesized nanoparticles have become a viable tool in regenerative medicine, especially in tissue engineering and wound healing. These biologically sourced nanoparticles, which come from bacteria, fungus, and plant extracts, have demonstrated enormous promise in fostering tissue regeneration and repair (Nandhini et al., 2024). Biosynthesized nanoparticles speed up wound healing by promoting angiogenesis, collagen production, and cell proliferation. Additionally, they play a key role in the creation of scaffolds for cartilage restoration and bone regeneration. Researchers have shown better osteogenic differentiation by including biosynthesized nanoparticles in these scaffolds, which helps with cartilage tissue repair and the regeneration of broken bones (Christy et al., 2020). By encouraging cell migration and vascularization, these nanoparticles improve the integration of skin transplants with host tissues. Moreover, their newfound significance in stem cell treatment is very fascinating. The results of regenerative treatments can be greatly enhanced by using biosynthesized nanoparticles as carriers for growth factors and genes that promote stem cell differentiation and proliferation. This includes using them to improve transplanted stem cells' integration and survival in injured tissues, opening up new therapeutic options for a variety of degenerative illnesses. Biosynthesized nanoparticles are expected to become a mainstay of regenerative medicine as research progresses, providing novel, efficient, and long-lasting approaches to tissue mending and regeneration (Bose et al., 2020).

Nanocarriers for Hormone Therapy in Chronic Disease Management

In the field of hormone therapy for the treatment of chronic diseases, notably diabetes and thyroid problems, nanocarriers have shown great promise. The precise targeting and regulated release of hormones made possible by these nanoparticle-mediated delivery methods improve the effectiveness of hormone therapy by preserving stable hormone levels for extended periods (Hashem et al., 2020). For example, nanocarriers can be utilized to administer insulin more precisely in diabetes treatment, imitating the body's natural release patterns and enhancing glucose control while lowering dosage frequency. Comparably, formulations based on nanoparticles provide a way to provide levothyroxine in thyroid diseases more efficiently, guaranteeing that patients receive constant hormone levels something that might be problematic with conventional oral administration. Hormone treatments typically cause systemic adverse effects; however, by encasing hormones into nanoparticles, the therapy becomes more biocompatible and less likely to degrade before reaching the target location (Mirza et al., 2021). Additionally, endocrine problems can be specifically modified to be treated using precision-engineered nanocarriers, offering a customized approach. Particle size, surface characteristics, and release kinetics may all be altered thanks to this tailoring, which promotes focused and prolonged hormone release and improves therapeutic results. Nanocarrier systems have the potential to completely transform hormone treatment as this field of study develops since they can increase therapeutic accuracy in the management of chronic illnesses, minimize side

effects, and improve patient adherence (Ingersoll et al., 2008).

Challenges in Clinical Translation of Biosynthesized Nanoparticles

There are several obstacles in the way of biosynthesized nanoparticles' clinical translation, mostly related to regulatory approval, scalability, and mainstream healthcare integration. Regulatory obstacles are a big worry as biosynthesized nanoparticles sometimes involve intricate biological systems that raise special questions about their safety and effectiveness (Dos Santos et al., 2014). Although strict preclinical and clinical studies are mandated by regulatory agencies like the FDA and EMA, standard operating procedures are still lacking for determining the toxicity, long-term biocompatibility, and environmental effects of these nanoparticles. Variations in raw materials might result in inconsistencies in the end product, which further complicates the process of biosynthesis. Examples of diverse natural sources utilized in biosynthesis are plant extracts and microbial agents. Scalability presents another important difficulty. While biosynthesized nanoparticles may be produced on a laboratory scale, achieving large-scale manufacturing involves overcoming challenges such as stability, uniformity in particle size, and purity all of which are essential for therapeutic effectiveness (Patel et al., 2021). Furthermore, there is still uncertainty about how cost-effective large-scale production may be. The future of biosynthesized nanoparticles is bright, even despite these obstacles. Scalability problems may be resolved by developments in nanotechnology and biotechnology, and the rising need for environmentally friendly, sustainable substitutes for chemical synthesis is propelling innovation in these fields. Biosynthesized nanocarriers have the potential to revolutionize healthcare by providing safer and more environmentally friendly solutions in drug delivery systems, cancer treatments, and other personalized medicine fields if regulatory frameworks can adjust to these developing technologies (Sainz et al. 2015).

Ethical and Environmental Considerations in Biosynthesized Nanoparticles

With the rising breadth of nanotechnology, ethical and environmental problems in the manufacture and deployment of biosynthesized nanoparticles have gained increasing attention. The effects of large-scale biosynthetic processes on the environment, especially the possible release of nanoparticles into ecosystems, are a major source of worry (Pérez-Hernández et al., 2021). Despite being biosynthesized, these particles can interact with their surroundings in ways that might damage nearby ecosystems, cause them to bioaccumulate in living things, or affect biodiversity. Adopting sustainable manufacturing processes and conducting comprehensive environmental evaluations are necessary for mitigating these hazards. It's also important to consider the moral ramifications of employing biological systems like fungi, plants, or bacteria to produce nanoparticles. These biological agents are frequently subjected to genetic modification or manipulation, raising concerns about the proper handling of living systems and the potential ecological fallout from unleashing altered species (Bonneuil et al., 2014). In addition, the regulatory frameworks of biosynthesized nanoparticles are in a state of constant evolution; yet, they are vital for guaranteeing environmental security as well as

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ethical accountability. Governments and international organizations are starting to focus on the sustainability of producing nanoparticles and are promoting the synthesis of these particles using green chemistry and renewable resources. To minimize unforeseen repercussions and guarantee that biosynthesized nanotechnology complies with ethical and environmental norms, strong and enforced laws are still required. This will promote both ecological balance and technological progress (Yadav et al., 2019).

Summary

Because of their special qualities, biosynthesized nanoparticles (NPs) have become cutting-edge nanocarriers for targeted therapeutic interventions, transforming the management of illness. These biocompatible, environmentally beneficial nanoparticles provide an alternative to traditional chemical synthesis techniques. They are sourced from biological sources such as microbes, plants, and biopolymers. The exact distribution of therapeutic substances is made possible by their enormous surface area, tiny size, and functionalization, which maximizes efficacy and minimizes adverse effects. Biosynthesized NPs have the potential to be very useful in cancer therapy, cardiovascular illnesses, and neurological disorders because they target certain cells or tissues, hence improving drug bioavailability and reducing off-target toxicity. Furthermore, their capacity to go through biological barriers including the blood-brain barrier opens up new therapeutic options for diseases like Alzheimer's and brain cancers. Because of their natural nature, the use of biosynthesized nanocarriers further lowers the danger of immunological rejection and presents a possibility for scalable and sustainable manufacturing processes. With further investigation, the use of biosynthesized nanoparticles in precision medicine may improve patient outcomes and result in safer, more individualized, and more efficient treatment plans for managing illnesses.

Declarations

Data Availability statement

All data generated or analyzed during the study are included in the manuscript.

Ethics approval and consent to participate

Approved by the department Concerned.

Consent for publication

Approved

Funding

Not applicable

Conflict of interest

The authors declared absence of conflict of interest.

References

Ahmed, S. F., Mofijur, M., Rafa, N., Chowdhury, A. T., Chowdhury, S., Nahrin, M., ... & Ong, H. C. (2022). Green approaches in synthesising nanomaterials for environmental nanobioremediation: Technological advancements, applications, benefits and challenges. *Environmental Research*, 204, 111967.

- Albanese, A., Tang, P. S., & Chan, W. C. (2012). The effect of nanoparticle size, shape, and surface chemistry on biological systems. *Annual review of biomedical engineering*, 14(1), 1-16.
- Albrecht, M. A., Evans, C. W., & Raston, C. L. (2006). Green chemistry and the health implications of nanoparticles. *Green chemistry*, 8(5), 417-432.
- Ameen, F., Alsamhary, K., Alabdullatif, J. A., & ALNadhari, S. (2021). A review on metal-based nanoparticles and their toxicity to beneficial soil bacteria and fungi. *Ecotoxicology and Environmental Safety*, 213, 112027.
- Bezbaruah, R., Chavda, V. P., Nongrang, L., Alom, S., Deka, K., Kalita, T., ... & Vora, L. (2022). Nanoparticle-based delivery systems for vaccines. *Vaccines*, 10(11), 1946.
- Bharti, B., Kumar, R., Kumar, H., Li, H., Zha, X., & Ouyang, F. (2022). Advanced applications and current status of green nanotechnology in the environmental industry. In *Green functionalized nanomaterials for environmental applications* (pp. 303-340). Elsevier.
- Bonneuil, C., Foyer, J., & Wynne, B. (2014). Genetic fallout in bio-cultural landscapes: Molecular imperialism and the cultural politics of (not) seeing transgenes in Mexico. *Social Studies of Science*, 44(6), 901-929.
- Bose, S., Sarkar, N., & Banerjee, D. (2020). Controlled Delivery of Natural Medicinal Compounds from Tissue Engineering Scaffolds for Critical-Sized Bone Defect Repair. Available at SSRN 3708576.
- Christy, P. N., Basha, S. K., Kumari, V. S., Bashir, A. K. H., Maaza, M., Kaviyarasu, K., ... & Ignacimuthu, S. (2020). Biopolymeric nanocomposite scaffolds for bone tissue engineering applications—A review. *Journal of drug delivery science and technology*, 55, 101452.
- Das, S., Bhardwaj, A., & Pandey, L. M. (2021). Functionalized biogenic nanoparticles for use in emerging biomedical applications: a review. *Current Nanomaterials*, 6(2), 119-139.
- Dos Santos, C. A., Seckler, M. M., Ingle, A. P., Gupta, I., Galdiero, S., Galdiero, M., ... & Rai, M. (2014). Silver nanoparticles: therapeutic uses, toxicity, and safety issues. *Journal of Pharmaceutical Sciences*, 103(7), 1931-1944.
- Fathi-Achachelouei, M., Knopf-Marques, H., Ribeiro da Silva, C. E., Barthès, J., Bat, E., Tezcaner, A., & Vrana, N. E. (2019). Use of nanoparticles in tissue engineering and regenerative medicine. *Frontiers in bioengineering and biotechnology*, 7, 113.
- Gavas, S., Quazi, S., & Karpiński, T. M. (2021). Nanoparticles for cancer therapy: current progress and challenges. *Nanoscale research letters*, 16(1), 173.
- Gregory, A. E., Titball, R., & Williamson, D. (2013). Vaccine delivery using nanoparticles. *Frontiers in cellular and infection microbiology*, 3, 13.
- Grunwald, P. (2020). Genome Editing and Gene Therapies: Complex and Expensive Drugs. In *Pharmaceutical Biocatalysis* (pp. 281-374). Jenny Stanford Publishing.
- Gurunathan, S., Qasim, M., Choi, Y., Do, J. T., Park, C., Hong, K., ... & Song, H. (2020). Antiviral potential of nanoparticles—can nanoparticles fight against coronaviruses?. *Nanomaterials*, 10(9), 1645.
- Hashem, N. M., & Gonzalez-Bulnes, A. (2020). State-of-the-art and prospective of nanotechnologies for smart reproductive management of farm animals. *Animals*, 10(5), 840.
- Haverkamp, R. G., & Marshall, A. T. (2009). The mechanism of metal nanoparticle formation in plants: limits on accumulation. *Journal of Nanoparticle Research*, 11, 1453-1463.
- Hosseini, M., & Mozafari, M. (2020). Cerium oxide nanoparticles: recent advances in tissue engineering. *Materials*, 13(14), 3072.
- Ingersoll, K. S., & Cohen, J. (2008). The impact of medication regimen factors on adherence to chronic treatment: a review

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- of the literature. *Journal of Behavioral Medicine*, 31, 213-224.
- Kemp, J. A., & Kwon, Y. J. (2021). Cancer nanotechnology: current status and perspectives. *Nano convergence*, 8(1), 34.
- Khan, F., Shahid, A., Zhu, H., Wang, N., Javed, M. R., Ahmad, N., ... & Mehmood, M. A. (2022). Prospects of algae-based green synthesis of nanoparticles for environmental applications. *Chemosphere*, 293, 133571.
- Khan, F., Shahid, A., Zhu, H., Wang, N., Javed, M. R., Ahmad, N., ... & Mehmood, M. A. (2022). Prospects of algae-based green synthesis of nanoparticles for environmental applications. *Chemosphere*, 293, 133571.
- Kim, B. H., Yang, J., Lee, D., Choi, B. K., Hyeon, T., & Park, J. (2018). Liquid-phase transmission electron microscopy for studying colloidal inorganic nanoparticles. *Advanced Materials*, 30(4), 1703316.
- Koul, B., Poonia, A. K., Yadav, D., & Jin, J. O. (2021). Microbe-mediated biosynthesis of nanoparticles: Applications and future prospects. *Biomolecules*, 11(6), 886.
- Kumar, J. A., Krithiga, T., Manigandan, S., Sathish, S., Renita, A. A., Prakash, P., ... & Crispin, S. (2021). A focus to green synthesis of metal/metal based oxide nanoparticles: Various mechanisms and applications towards ecological approach. *Journal of Cleaner Production*, 324, 129198.
- Li, X., Xu, H., Chen, Z. S., & Chen, G. (2011). Biosynthesis of nanoparticles by microorganisms and their applications. *Journal of nanomaterials*, 2011(1), 270974.
- Lim, E. K., Kim, T., Paik, S., Haam, S., Huh, Y. M., & Lee, K. (2015). Nanomaterials for theranostics: recent advances and future challenges. *Chemical Reviews*, 115(1), 327-394.
- Mao, Y., Park, T. J., Zhang, F., Zhou, H., & Wong, S. S. (2007). Environmentally friendly methodologies of nanostructure synthesis. *Small*, 3(7), 1122-1139.
- Matheolabakis, G., Rigas, B., & Constantinides, P. P. (2012). Nanodelivery strategies in cancer chemotherapy: biological rationale and pharmaceutical perspectives. *Nanomedicine*, 7(10), 1577-1590.
- Mirza, Z., & Karim, S. (2021, February). Nanoparticles-based drug delivery and gene therapy for breast cancer: Recent advancements and future challenges. In *Seminars in cancer biology* (Vol. 69, pp. 226-237). Academic Press.
- Moritz, M., & Geszke-Moritz, M. (2013). The newest achievements in synthesis, immobilization and practical applications of antibacterial nanoparticles. *Chemical Engineering Journal*, 228, 596-613.
- Nandhini, J., Karthikeyan, E., Rani, E. E., Karthikha, V. S., Sanjana, D. S., Jeevitha, H., ... & Priyadharshan, A. (2024). Advancing engineered approaches for sustainable wound regeneration and repair: Harnessing the potential of green synthesized silver nanoparticles. *Engineered Regeneration*, 5(3), 306-325.
- Nisar, P., Ali, N., Rahman, L., Ali, M., & Shinwari, Z. K. (2019). Antimicrobial activities of biologically synthesized metal nanoparticles: an insight into the mechanism of action. *JBIC Journal of Biological Inorganic Chemistry*, 24, 929-941.
- Ovais, M., Khalil, A. T., Islam, N. U., Ahmad, I., Ayaz, M., Saravanan, M., ... & Mukherjee, S. (2018). Role of plant phytochemicals and microbial enzymes in the biosynthesis of metallic nanoparticles. *Applied microbiology and biotechnology*, 102, 6799-6814.
- Pala, R., Anju, V. T., Dyavaiah, M., Busi, S., & Nauli, S. M. (2020). Nanoparticle-mediated drug delivery for the treatment of cardiovascular diseases. *International journal of nanomedicine*, 3741-3769.
- Pandit, C., Roy, A., Ghotekar, S., Khusro, A., Islam, M. N., Emran, T. B., ... & Bradley, D. A. (2022). Biological agents for the synthesis of nanoparticles and their applications. *Journal of King Saud University-Science*, 34(3), 101869.
- Patel, D. M., Patel, N. N., & Patel, J. K. (2021). Nanomedicine scale-up technologies: feasibilities and challenges. In *Emerging Technologies for Nanoparticle Manufacturing* (pp. 511-539). Cham: Springer International Publishing.
- Pérez-Hernández, H., Pérez-Moreno, A., Sarabia-Castillo, C. R., García-Mayagoitia, S., Medina-Pérez, G., López-Valdez, F., ... & Fernández-Luqueño, F. (2021). Ecological drawbacks of nanomaterials produced on an industrial scale: collateral effect on human and environmental health. *Water, Air, & Soil Pollution*, 232, 1-33.
- Rahdar, A., Amini, N., Askari, F., & Susan, M. A. B. H. (2019). Dynamic light scattering: A useful technique to characterize nanoparticles. *Journal of Nanoanalysis*, 6(2), 80-89.
- Rónavári, A., Igaz, N., Adamecz, D. I., Szerencsés, B., Molnar, C., Kónya, Z., ... & Kiricsi, M. (2021). Green silver and gold nanoparticles: Biological synthesis approaches and potentials for biomedical applications. *Molecules*, 26(4), 844.
- Sainz, V., Coniot, J., Matos, A. I., Peres, C., Zupančič, E., Moura, L., ... & Gaspar, R. S. (2015). Regulatory aspects on nanomedicines. *Biochemical and biophysical research communications*, 468(3), 504-510.
- Shafey, A. M. E. (2020). Green synthesis of metal and metal oxide nanoparticles from plant leaf extracts and their applications: A review. *Green Processing and Synthesis*, 9(1), 304-339.
- Shankar, P. D., Shobana, S., Karuppusamy, I., Pugazhendhi, A., Ramkumar, V. S., Arvindnarayan, S., & Kumar, G. (2016). A review on the biosynthesis of metallic nanoparticles (gold and silver) using bio-components of microalgae: Formation mechanism and applications. *Enzyme and Microbial Technology*, 95, 28-44.
- Stan, D., Enciu, A. M., Mateescu, A. L., Ion, A. C., Brezeanu, A. C., Stan, D., & Tanase, C. (2021). Natural compounds with antimicrobial and antiviral effects and nanocarriers are used for their transportation. *Frontiers in pharmacology*, 12, 723233.
- Vargas-Estrada, L., Torres-Arellano, S., Longoria, A., Arias, D. M., Okoye, P. U., & Sebastian, P. J. (2020). Role of nanoparticles on microalgal cultivation: A review. *Fuel*, 280, 118598.
- Xu, X., Liu, C., Wang, Y., Koivisto, O., Zhou, J., Shu, Y., & Zhang, H. (2021). Nanotechnology-based delivery of CRISPR/Cas9 for cancer treatment. *Advanced drug delivery reviews*, 176, 113891.
- Yadav, S. K., Lal, S., Yadav, S., Laxman, J., Verma, B., Sushma, M., ... & Sharma, V. (2019). Use of nanotechnology in agri-food sectors and apprehensions: an overview. *Seed Res*, 47(2), 99-149.
- Yousaf, Z., & Saleh, N. (2018). Advanced concept of green synthesis of metallic nanoparticles by reducing phytochemicals. *Nanobotany*, 17-36.



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