

The contribution of cyanobacteria in the development of nanobiotechnology: a mini-review

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Abstract This review shows how adaptable cyanobacteria are for biotechnological uses; they are potent sources of bioactive substances, biofertilizers, bioplastics, energy, food, and pharmaceuticals. In the last decades, they have been applied to the synthesis of nanoparticles and in the bioremediation process. Diazotrophic cyanobacteria are known for their ability to fix nitrogen which making them important in the global nitrogen cycle. Cyanobacteria play a key role in the ecosystem by producing oxygen for other organisms. Moreover, they play a crucial role in the environment through their ability to perform photosynthesis, and nitrogen fixation in some species, and release important organic compounds. Their metabolic activities also contribute to the formation of a sizable portion of the world's limestone through the development of travertine and stromatolites. One of the notable features of some cyanobacteria is their filamentous growth habit; others are unicellular, which can range from unicellular to quasi-multicellular. The distribution of cyanobacteria is influenced by several factors, including environmental conditions, nutrient availability, and biotic interactions. Cyanobacteria have been found to interact with several types of metal nanoparticles (NPs), leading to the discovery of new potential uses and applications for these NPs. The interaction between cyanobacteria and metal NPs has been studied in different fields such as medicine, environmental protection, and industrial production, showing promising results in terms of anti-pathogenic, cytotoxicity, and antioxidant properties. To conclude, different strains of cyanobacteria and various types of metal NPs they interact with have a wide range of potential applications.

Keywords Cyanobacteria . Green nanotechnology . Biotechnology . Applications . Nanoparticles

Cyanobacteria and its applications: an overview

Cyanobacteria have a broad range of purposes in medicine, pharmacy, biotechnology, and renewable energy as they can be utilized in a variety of items, including cosmetics, detergents, soaps, toothpaste, shampoos, and bioenergy (Zahra et al. 2020). Utilizing cyanobacteria in nanoparticle biosynthesis was chosen for various reasons, including ease of culturing, quick generation time, and ability to withstand mild pH, temperature, and pressure conditions (Elsayed 2017; Hussein et al. 2022). It was discovered that cyano-

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bacteria can produce natural bioactive substances such as phenolic compounds, which have antioxidant, anti-inflammatory, and anti-microbial properties (Komárek 2013). The following sections discuss in more detail the production, characteristics, and main applications of cyanobacteria.

Isolation and purification of cyanobacteria

Diazotrophic cyanobacteria are known for their ability to fix nitrogen that is making them important in the global nitrogen cycle. One common method for isolating cyanobacteria is the utilization of enrichment culture techniques (Hotos et al. 2023). This involves growing the organisms in a medium specifically designed to promote the target organism's growth while inhibiting other microorganisms' growth (Reeve et al. 2023). For cyanobacteria, this can involve the use of media that is high in nitrogen, such as ammonium or nitrate, and low in phosphorus (Markou and Georgakakis 2011). Another method for isolating cyanobacteria is the use of dilution plating (Rippka 1988). This involves spreading a known amount of a mixed culture onto a solid medium and then diluting the culture by spreading a known amount of the diluted culture onto new plates. This process is repeated until a single colony is obtained. This method is particularly useful for isolating rare or difficult-to-grow strains of cyanobacteria. A third method for isolating cyanobacteria is with filtration. This involves passing a sample through a filter with a known pore size to capture the target organism while removing other microorganisms (Nnadozie et al. 2015).

Overview of cyanobacteria

Cyanobacteria are a diverse group of microorganisms that come in various shapes, such as unicellular, filamentous, and colonial forms. They can be found in a wide range of environments, including hot and cold temperatures, alkaline and acidic conditions, and freshwater, marine, saline, and terrestrial habitats (Sanchez-Baracaldo et al. 2005). Cyanobacteria play a key role in the ecosystem by producing oxygen for other organisms. They possess a primitive nucleus, gas vesicles, and thylakoids that are loosely distributed, along with hexagonal carboxysomes, phycobilisomes, ribosomes, and numerous storage granules (Gallardo 2014). They are characterized by the presence of chlorophyll, accessory pigments, and the PSII reaction center. The diazotrophic cyanobacteria can utilize inorganic nitrogen via permeases and fix nitrogen using nitrogenase enzymes as in diazotrophic, utilizing photosystem I to create adenosine triphosphate (ATP). They hold potential as natural biofertilizers (Glazer 1977; Kalyanasundaram et al. 2020), but their effectiveness can be limited by physiological changes and variations in inoculum depending on the ecological zone.

Importance of cyanobacteria

Cyanobacteria play a crucial role in the environment through their ability to perform photosynthesis, fix nitrogen in some species, and release important organic compounds. Their metabolic activities also contribute to the formation of a significant portion of the world's limestone through the development of travertine and stromatolites (Shridhar 2012). As primary producers in both aquatic and terrestrial ecosystems, cyanobacteria make up a significant portion of the biomass in many aquatic environments (Gladyshev et al. 2013). Additionally, they are important for health and industry as they can produce a variety of valuable natural compounds (Saranraj and Sivasakthi 2014). Furthermore, their role in ecology is notable, particularly in rice-growing countries, where they act as natural biofertilizers through their nitrogen-fixing abilities (Mishra and Pabbi 2004).

Main characters of cyanobacteria

Cyanobacteria are a diverse group of microorganisms that can be classified into five major sub-groups based on their morphological and physiological characteristics. This classification system is based on the dichotomous key presented in the second edition of Bergey's Manual of Systematic Bacteriology, which takes into account factors such as the type of cell division, the presence of differentiated cells, and the morphology of the organism (Pinevich 2008). One important feature of some cyanobacteria is their filamentous



growth habit; others are unicellular, which can range from unicellular to quasi-multicellular. The color of cyanobacterial cultures can vary from blue-green to brown. The shape and size of vegetative cells can vary depending on the species, with some having barrel-shaped cells and others having ovoid cells. The size of vegetative cells can range from 2.5-5.5 micrometers in diameter and 4.0-7.0 micrometers in length. Another essential feature of some filamentous cyanobacteria is the presence of specialized cells called heterocysts, which participate in nitrogen fixation.

Heterocyst is typically ovoid, with diameters ranging from 3.0-6.0 micrometers in width and 3.5-6.5 micrometers in length. They can be found in both internal and terminal positions within the filament. Some cyanobacteria also possess another specialized cell called akinetes, which are used for survival during adverse conditions. Akinetes are usually ovoid and vary from 4.0-6.0 micrometers in width and 5.0-10.0 micrometers in length (Brito 2015). Cyanobacteria are traditionally classified by phycologists based on their morphological and physiological characteristics. However, recent advances in molecular biology have allowed for the use of molecular markers in the classification of cyanobacteria. These markers can be used to differentiate between different strains and species of cyanobacteria. Overall, the classification of cyanobacteria is a complex process that requires the use of multiple criteria and techniques (Prasanna et al. 2006).

Factors that affect the distribution of cyanobacteria

Cyanobacteria are found in a wide range of environments, including freshwater, marine, and terrestrial habitats (Kultschar and Llewellyn 2018). The distribution of cyanobacteria is influenced by several factors, including environmental conditions, nutrient availability, and biotic interactions (Schulte et al. 2022). One of the primary factors that affect the distribution of cyanobacteria is the availability of light. Cyanobacteria are photosynthetic organisms that require light for growth and survival (Mischke 2003). Therefore, they are typically found in environments where light penetration is high, such as in clear water bodies or terrestrial habitats with low canopy cover. Additionally, certain species of cyanobacteria have adaptations, such as pigments that allow them to tolerate low light conditions resulting in surviving in environments with limited light penetration (Havens et al. 1998). Another key factor that affects the distribution of cyanobacteria is nutrient availability.

Cyanobacteria are known to be able to tolerate a wide range of nutrient conditions, but they typically thrive in environments where nutrient levels are high (Arias et al. 2020). For example, in freshwater systems, cyanobacteria often proliferate in eutrophic (nutrient-rich) environments, where nitrogen and phosphorus levels are elevated. In addition to light and nutrient availability, biotic interactions also play a role in shaping the distribution of cyanobacteria (Paerl and Otten 2013). Competition for resources among different organisms can limit the growth and spread of cyanobacteria. For example, in freshwater systems, cyanobacteria may be outcompeted by other photosynthetic organisms, such as macrophytes or phytoplankton. Climate change is also an important factor affecting the distribution of cyanobacteria (Salmaso 2000). Cyanobacteria can tolerate a wide range of temperatures, but certain species are more tolerant to warmer temperatures than others (Bonilla et al. 2023). As global temperatures increase, it is expected that certain species of cyanobacteria will become more dominant in certain regions, while others may decline. Overall, the distribution of cyanobacteria is influenced by a complex interplay of environmental factors, including light availability, nutrient availability, biotic interactions, and climate change. Understanding these factors is critical for predicting and managing the growth and spread of cyanobacteria in different environments.

Applications of cyanobacteria

Cyanobacteria are surprisingly the most significant class of microbes in the cosmos because they have critical environmental responsibilities, serving as a major source of oxygen, nitrogen, and carbon for the entire planet (Joye and Lee 2004). They are significant producers of biofuels, food additives, and coloring agents (Kumar et al. 2019). Cyanobacteria are utilized in the treatment of water (Svrcek and Smith 2004), the manufacture of bioplastics (Markl et al., 2018), cosmetics (Morone et al. 2019), forestry, animal feed (Grewe and Pulz 2012), biohydrogen, bioethanol, and biogas (Parmar et al. 2011). The following sections explore their applications in different sectors.



Cyanobacteria for nanoparticles biosynthesis

Cyanobacteria are a group of photosynthetic microorganisms that have been found to have the ability to biosynthesize nanoparticles (Doman et al. 2023). This process (Figure 1), known as biogenic synthesis, involves the use of enzymes and other biomolecules present in cyanobacteria to produce nanoparticles with unique properties and potential applications in various fields (Srivastava et al. 2022). One of the key features of cyanobacteria that allows for the biosynthesis of nanoparticles is their ability to produce complex mixtures of polysaccharides, proteins, and lipids; which produce extracellular polymeric substances (EPS) (Huang et al. 2023).

These EPS serve as a template for the formation of nanoparticles and can also provide a protective environment for the particles during their formation. The most widely studied type of nanoparticles biosynthesized by cyanobacteria is the metal nanoparticles, particularly those of silver, gold, and copper. The mechanisms by which these nanoparticles are formed are not fully understood, but it is believed that they involve the reduction of metal ions present in the EPS by enzymes or other biomolecules (Fulaz et al. 2019). In addition to metal nanoparticles, cyanobacteria have also been found to biosynthesize other types of nanoparticles, such as semiconductors and magnetic nanoparticles (Asmathunisha and Kathiresan 2013). The potential applications of these nanoparticles are diverse and include their implementation in medical diagnostics, environmental remediation, and renewable energy (Khan et al. 2022).

Despite the potential of cyanobacteria in the biosynthesis of nanoparticles, there are still some challenges that need to be overcome. For example, the yields of nanoparticles produced by cyanobacteria are often low, and the process can be affected by environmental factors such as pH and temperature (Rahman et al. 2020; Thakkar et al. 2010). In conclusion, cyanobacteria have shown enormous potential in the biosynthesis of nanoparticles. Using these microorganisms, unique nanoparticles with potential applications in various fields can be produced in a sustainable and environmentally friendly route. However, further research is needed to improve yields and optimize the biosynthesis process.

The synthesis of metal nanoparticles, such as silver nanoparticles (AgNPs), can be achieved through both physical and chemical methods (Abou El-Nour et al. 2010). However, these methods often have drawbacks, such as the use of expensive and toxic chemicals, difficulty in achieving mono-dispersion and thermodynamic stability, and high energy consumption with the potential for dangerous by-products (Irvani

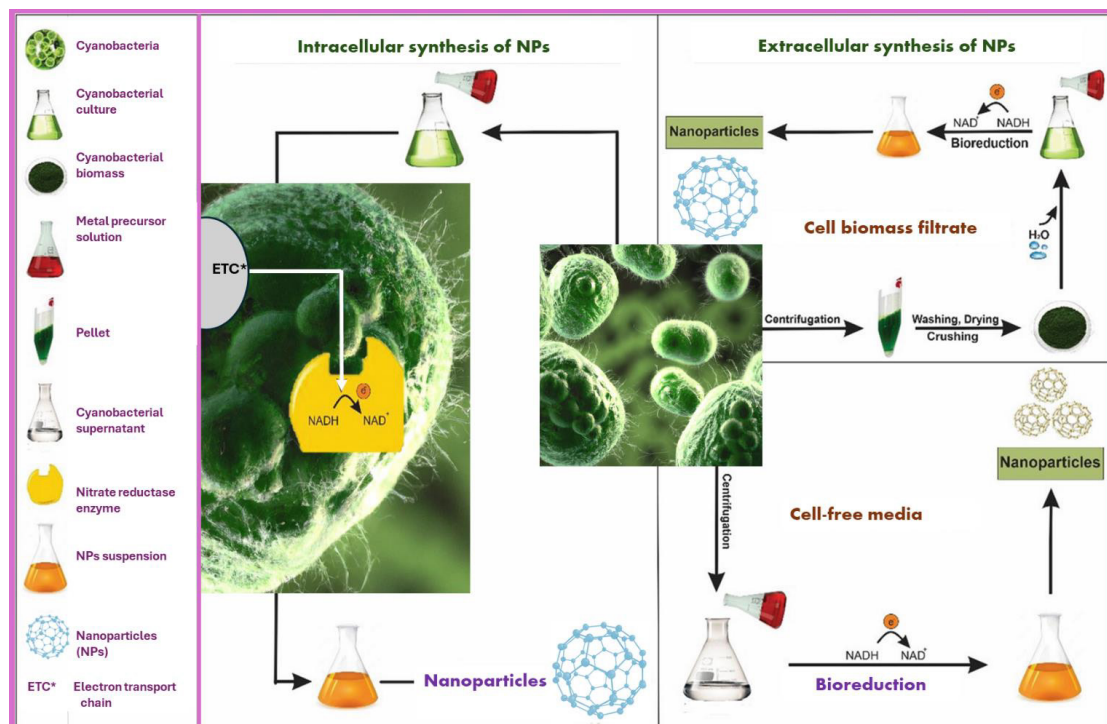


Fig. 1 A schematic flow diagram for the biosynthesis of nanoparticles from cyanobacteria (Javed et al. 2024)



et al. 2014). Green chemistry offers alternative methods for the synthesis of nanoparticles with applications in fields such as drug delivery, sterilization, and catalysis (Albrecht et al. 2006). Silver nanoparticles have strong antimicrobial properties and are commonly used in various consumer products, such as clothing and cosmetics (Mazari et al. 2021). The green method is considered safe, eco-friendly, and capable of large-scale production (Fathy et al. 2020). Additionally, it does not require high energy consumption and can be performed at room temperature under normal conditions (Abdel-Raouf et al. 2019).

The proteins, enzymes, sugars, and lipids present in cyanobacteria act as reducing and capping agents in the conversion of silver ions into nanoparticles (Roy et al. 2019). The hydroxyl group in tyrosine residues and the carboxyl group in aspartic acid and/or glutamic acid residues have been identified as the most active functional groups in this process (Makarov et al. 2014).

Biosynthesis of nanoparticles

Nano-biotechnology is a rapidly growing field within the broader field of nanoscience and technology, and nanoparticles have been shown to possess a wide range of physical, optical, electrical, and chemical characteristics (Jha et al. 2014; Sorbiun et al. 2018). This versatility makes nanoparticles highly attractive for use in biomedical science, including as antimicrobial agents, gene carriers, diagnostic agents, dental resin composites, tissue repair, optical receptors, and electrical batteries (Fathy et al. 2020).

Chemical synthesis of nanoparticles can be challenging, as it requires particular circumstances and has several drawbacks, including chemical toxicity and nanoparticle instability. Physical synthesis methods, while possible, are often energy-intensive and costly (Patel et al. 2015). Although, nanoparticles could be made in one of two approaches: top-down or bottom-up, in most cases, the bottom-up one is exploited in the biological development of nanoparticles (Ibraheem et al. 2016). However, the biological method is superior to the chemical and physical processes since it overcomes the disadvantages of other methods. It has several advantages, including low cost, stability, eco-friendly, and the fact that it does not consume energy (Patel et al. 2015).

Nanoparticles have a size range of 1-100 nm, making the interaction with microbial surfaces much easier, quicker, and more effective than microbes of typical size (Ibraheem et al. 2016). Cyanobacteria with noble metals like silver or metallic elements like copper are used in the biological technique (Hamida et al. 2020b). Bioactive chemicals produced by microorganisms perform a vital role in Ag^+ reduction and nanoparticle stabilization without toxicity or severe circumstances (Cruz et al. 2010; Jeevanandam et al. 2016; Pirtarighat et al. 2019). In general, parameters like the kind of solvent and the reducing agent for silver ions play a big influence on silver nanoparticle production (Ebrahimezhad et al. 2016). One of the most effective uses of nanoparticles is antimicrobial agents, which can be used in a variety of fields with amazing efficacy, such as pharmaceutical medication production against various microorganisms (Madkour 2017; Zhang et al. 2010).

Characterization of biosynthesized nanoparticles

Characterization is crucial to understanding their properties and potential applications. Several instruments and methods can be used to characterize these nanoparticles, including Transmission Electron Microscopy (TEM); TEM is a widely used technique for imaging and analyzing nanoparticles (Figure 2a). It uses a beam of electrons to produce high-resolution images of the particles, allowing for the determination of their size, shape, and crystalline structure (Wilson and Prud'homme 2021). Scanning electron microscopy (SEM) is another imaging technique that uses a beam of electrons to produce images of the surface of nanoparticles (Figure 2b). It is particularly useful for studying the surface morphology and topography of the particles (Titus et al. 2019). X-ray diffraction (XRD) is a technique that uses X-rays to determine the crystal structure of nanoparticles (Figure 2c). It can be used to identify the crystal phases present in the particles, as well as their size and shape (Bishnoi et al. 2017). UV-visible spectroscopy is another technique that uses light in the ultraviolet and visible range to study the optical properties of nanoparticles. It can be used to determine the absorption and scattering properties of the particles, as well as their size and



shape (Shehap and Akil 2016). Dynamic light scattering (DLS) is another technique that uses laser light to study the size and size distribution of nanoparticles in solution. It is a non-destructive method, making it useful for studying the stability of the particles over time (Hoo et al. 2008). Fourier transforms infrared spectroscopy (FTIR) is a technique that uses infrared radiation to study the vibrations of chemical bonds in nanoparticles (Figure 2d, 2e, and 2f). It can be used to identify the chemical composition of the particles and to study their chemical interactions with other molecules (Faghizadeh et al. 2016). In conclusion, the characterization of biosynthesized nanoparticles is crucial to understanding their properties and potential applications. Although, several instruments and methods can be used to study these particles, including TEM, SEM, XRD, UV-visible spectroscopy, DLS, and FTIR, however, each of these techniques has its limitations, and the choice of technique will depend on the specific questions being asked and the properties of the nanoparticles being studied.

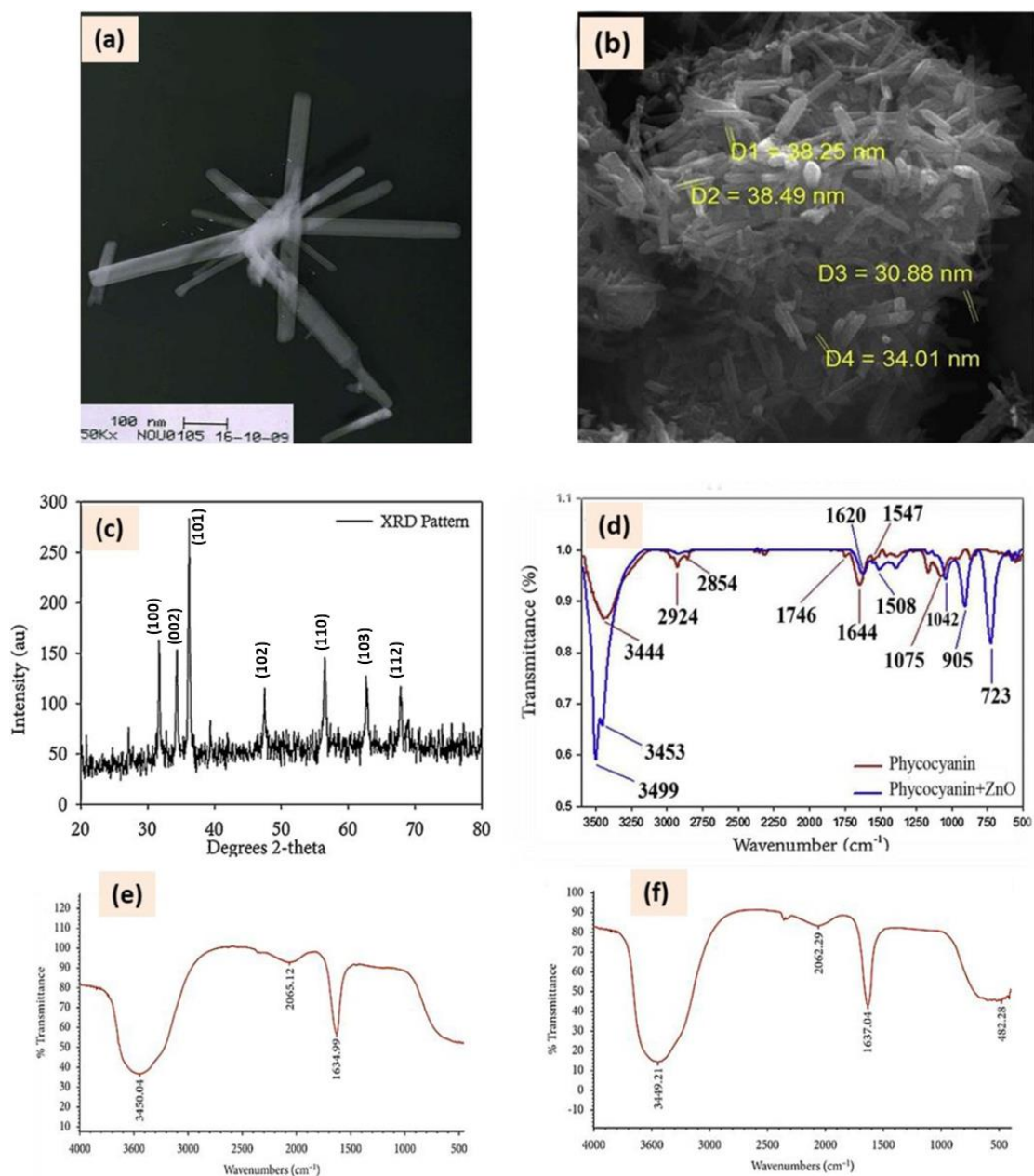


Fig. 2 TEM (a), SEM (b), XRD (c), and FTIR (d) of phycocyanin- ZnO nanorods (Davaeifar et al., 2019) and FTIR spectra of Ag-NPs synthesized extracellularly from *Desertifilum tharense* (e) and *Phormidium ambiguum* (f) (Hanna et al. 2022).



Selectivity of cyanobacteria in the biosynthesis of nanoparticles

Cyanobacteria have been found to interact with various types of metal nanoparticles (NPs), leading to the discovery of new potential uses and applications for these NPs. The interaction between cyanobacteria and metal NPs has been studied in fields such as medicine, environmental protection, and industrial production, showing promising results in terms of anti-pathogenic properties, cytotoxicity, and antioxidant properties. It can be inferred that the different strains of cyanobacteria and the various types of metal NPs they interact with have a wide range of potential applications (Table 1). The data also suggests that certain strains of cyanobacteria have a greater potential for certain applications, such as *Spirulina platensis* for lead uptake and *Anabaena variabilis* for malachite green dye removal.

It is also worth noting that some of the entries in the data have missing or unspecified NP sizes, which could affect the applicability of the findings. In the future perspective, the potential uses and applications of cyanobacteria and metal NPs in various fields such as medicine, environmental protection, and industrial production could be further explored and developed. Further research on the interactions between cyanobacteria and metal NPs, including optimization of NPs size and composition, could lead to the development of new and improved products and technologies.

From the Table interpretation, silver and gold nanoparticles are the most used metals in the biosynthesis of nanoparticles by cyanobacteria. Silver nanoparticles have strong antioxidant and antibacterial properties to kill various types of respiratory tract pathogenic bacteria (Song et al. 2019). They also have the potential to the removal of malachite green dye and suppress pathogenic bacteria growth and cytotoxic effects on cancer cells (Sathiyavimal et al. 2022). Gold nanoparticles, on the other hand, have been found to have various potential applications. They have been found to enhance the transmission of electrons through the cell membrane by absorbing light and increasing the number of photo-excited electrons (Liu and Choi 2021). They also have activities that figureht off free radicals and myocardial infarction. They also have the potential as biocatalysts, bio-selective characters, and biocompatible (Wang et al. 2021). In summary, silver and gold nanoparticles have various potential applications and are commonly used in the biosynthesis of nanoparticles by cyanobacteria. These nanoparticles have promising characteristics in biomedical, environmental, and industrial fields.

Cyanobacterial bioactive compounds

As an abundant source of bioactive chemicals, cyanobacteria have been discovered. The majority of the bioactive substances that have been identified from cyanobacteria are lipopeptides, which are made up of an amino acid fragment and a fatty acid portion. Secondary metabolites extracted from cyanobacteria have a wide spectrum of biological activity, including antibacterial, antifungal, antialgal, anti-protozoan, and antiviral properties

Medical and pharmaceutical biotechnology

Cyanobacteria comprises several secondary metabolites that are useful in the field of medical biotechnology. These microbes have received tremendous attention from researchers because of the generation of bioactive compounds that are incredibly useful in medical settings (Żymańczyk-Duda et al. 2022). Although they generate effective toxins, they also generate various metabolites that are vital in terms of their anticancer (Dash et al. 2023), antibiotic (Saeed et al. 2022), anti-inflammatory (Gómez et al. 2023), immunosuppressant (Polyak and Sukharevich, 2023) and antimicrobial (Frazzini et al. 2022) effects. Cyanobacteria come in a variety of forms, making them a potential source of chemicals with therapeutic potential for malignant diseases including cancer. Different metabolic processes can be combined to produce a wide variety of bioactive substances with enormous biological potential.

Bioplastics and biofuel production

Cyanobacteria are incredibly well-suited for producing biodegradable plastics and biofuels because they



Table 1 A collection of some cyanobacteria used in different nanoparticle biosynthesis.

| Cyanobacteria strains | Metal | NP Size (nm) | Applications | Reference |
|--|-------------------|--------------|--|-------------------------------|
| <i>Phormidium tenue</i> | Ca | 5 | Bio tagging | (MubarakAli et al. 2012) |
| <i>Cylindrocapsa stagnale</i> | Cu | 12.21 | Effectively inhibit the growth of the pathogens <i>Canida albicans</i> , <i>Klebsiella pneumoniae</i> , <i>Enterobacter cloacae</i> , <i>Pseudomonas aeruginosa</i> , and <i>Escherichia coli</i> , with minimum inhibitory concentrations of 1.5, 2.4, 1.7, 2.5, and 0.6 mM, respectively. Cytotoxic to the HepG2 cell line and effective against the larvae of <i>Aedes aegypti</i> , <i>Anopheles subpictus</i> <i>Grassi</i> , and <i>Culex quinquefasciatus</i> . | (Sombol et al. 2021) |
| <i>Synechocystis</i> sp. PCC 6803 | - | - | Enhances the transmission of electrons through the cell membrane by absorbing light and increasing the number of photo-excited electrons. | (Liu and Choi 2021) |
| <i>Lyngbya majuscula</i> | 41 | | Fight off free radicals and myocardial infarction. | (Bakir et al. 2018) |
| <i>Anabaena cylindrica</i> | < 10 | | Cyanobacteria have a bioselective character. | (Rochert et al. 2017) |
| <i>Anabaena laxa</i> | - | | Biocatalyst | (Lemartowicz et al. 2017) |
| <i>Lepolyngbya tenuis</i> | Au | 40 | Bio-compatibility | (Parial et al. 2016) |
| <i>Coleofasciculus thiomoplastes</i> | | | | |
| <i>Nostoc ellipsosporum</i> | | | | |
| <i>Phormidium willeyi</i> | 25 | | Antioxidant qualities | (MubarakAli et al. 2013) |
| <i>Synechocystis</i> sp. PCC 6803 | 13 ± 2 | | biolabeling (gold and DNA interaction) | (Foesan et al. 2011) |
| <i>Spirulina subsalsa</i> | < 20 | | Biomolecule sensing | (Chakraborty et al. 2009) |
| <i>Spirulina plantensis</i> | 10-20 | | Bio-recovery of gold | (Sayadi et al. 2018) |
| <i>Arthrospira (Spirulina) platensis</i> | Se | 100-550 | Lead uptake up to 90% | (Zincovscaia et al. 2017) |
| <i>Nostoc muscorum</i> NCCU 442 | 30 | | Antioxidant Action | (Husain et al. 2021) |
| <i>Spirulina platensis</i> | 13 | | Strong antioxidant and antibacterial properties. | (Ameen et al. 2020) |
| <i>Anabaena variabilis</i> | 17.9-26.4 | | The highest level of antibacterial activity was observed against the pathogen <i>Staphylococcus aureus</i> MTCC 902. | (Ismail et al. 2021) |
| <i>Spirulina platensis</i> | 4.5-26 | | The ability to kill seven different types of bacteria commonly found in the respiratory tract. | (Hamida et al. 2020a) |
| <i>Desertifilum</i> IPPAS B-1220 | - | | Malachite green dye was removed by 93% using <i>S. platensis</i> and 82% using <i>A. variabilis</i> AgNPs | (Sahoo et al. 2020) |
| <i>Chroococcus minutus</i> | 5-25 | | Five pathogenic bacteria growth was suppressed | (Ebrahimzadeh et al. 2020) |
| <i>Anabaena flos-aquae</i> | < 50 | | Cytotoxic effects on MCF-7, HepG2, and Caco-2 cancer cells with IC50 values of 58, 32, and 90 µg/mL, respectively. | (Tomer et al. 2019) |
| <i>Haloleptolyngbya alacalis</i> KR2005/106 | Ag | | The lowest effective dose of 100 mg was found to have the ability to kill pathogenic strains of <i>E. coli</i> and <i>S. pyogenes</i> , which are responsible for upper respiratory tract infections. | (Al Rashed et al. 2018) |
| <i>Phormidium</i> sp. | - | | Treatment of T47D cells with AgNPs for 24 h resulted in a 6.21% increase in apoptosis and a 28% increase in necrosis. | (El-Naggar et al. 2017) |
| <i>Phycocyanin extracted from Nostoc linckia</i> | 9-26 | | Monitoring water quality using ammonia. | (Soner et al. 2017) |
| <i>Nostoc</i> sp. strain HKAR-2 | 51-100 | | Antimicrobial for <i>P. aeruginosa</i> , <i>E. coli</i> , and <i>S. aureus</i> | (Roychoudhury et al. 2016) |
| <i>Lyngbya majuscula</i> | 20-50 | | Breast cancer cytotoxicity test and antibacterial against <i>S. aureus</i> , <i>Pseudomonas aeruginosa</i> , <i>E. coli</i> , and <i>Klebsiella pneumoniae</i> . | (San Keskin et al. 2016) |
| <i>Synechococcus</i> sp. | 430-450 | | Anti-cancer cytotoxic action in humans MCF-7 cells were discovered to have an IC50 of 27.5 µg/mL | (Kallamurthi et al. 2016) |
| <i>Spirulina platensis</i> | - | | Antibacterial and antifungal activities. | (Morsy et al. 2014) |
| <i>Nostoc commune</i> | 15-54 | | Both antibiotic and leukemic | (Sudha et al. 2013) |
| <i>Microcoleis</i> sp. | 44-64 | | Degradation of methylene blue | (Hifney and Abdel-Wahab 2019) |
| <i>Spirulina platensis</i> | Ti | 17.3 | Use of therapeutic thermostable silver nanoparticles | (Ebad et al. 2022) |
| <i>Desertifilum</i> sp. EAZ03 | 88 | | Surface sterilization, antibacterial and antibacterial | |
| <i>Limnodinium</i> sp. KO05 | Zn | 33 | Antimicrobial against <i>Strepococcus</i> sp., <i>B. subtilis</i> , <i>S. aureus</i> , <i>E. coli</i> , <i>P. vulgaris</i> , <i>S. typhi</i> , and <i>V. cholera</i> | |
| <i>Anabaena</i> sp. L31 | 80 | | Powerful antioxidant ability | |
| <i>Anabaena variabilis</i> | AgCl ₂ | 12-20 | Antimicrobial, antibiofilm, and anticancer properties against the pathogens <i>S. aureus</i> , <i>E. coli</i> , and <i>P. aeruginosa</i> . | |
| | | | Anticancer properties against normal lung (MRC-5) cells. | |
| | | | Compared to cells treated with ZnONPs, fibroblast L929 was less cytotoxic when exposed to PHY-ZnO NPs | |
| | | | UV-B-absorbing ZnO nanoparticles combined with the chemical shikorine | |
| | | | A new report for utilizing the cyanobacterial species | |
| | | | Anabaena variabilis in the nanotechnology field could provide future possible strategies for application in many fields. | |

have such remarkable mechanisms for fixing and absorbing atmospheric nitrogen and CO₂ and using them to grow in unfavorable climatic environments, like infertile soils and saline waters (Samadhiya et al. 2022). *Synechocystis*, *Spirulina*, *Anabaena*, and *Nostoc muscorum* are cyanobacteria that can serve as bio-factories to produce biofuel and bioplastic. They have the metabolic capacity to produce, among other copolymers, polyhydroxybutyrate (PHB) and polyhydroxyalkanoates (PHAs); two biopolymers that are both affordable and sustainable (Singh et al., 2017). Polyhydroxyalkanoates (PHA) is a biocompatible substance that is also being researched for use in the biomedical (Ansari et al. 2021) and biopharmaceutical (Elmowafy et al. 2019) industries. It is believed that synthesizing PHA from CO₂ will help developing a carbon-neutral plastic manufacturing method.

The material properties of biopolymeric PHB are similar to those of polypropylene; a common plastic made from petroleum (fossil fuels). PHB is biodegradable compared to normal plastics; therefore using it in place of standard plastics can help mitigate the negative ecological effects of excessive consumption of nonbiodegradable plastics and fossil fuels (Kiran et al., 2014; Klanchui et al. 2017).

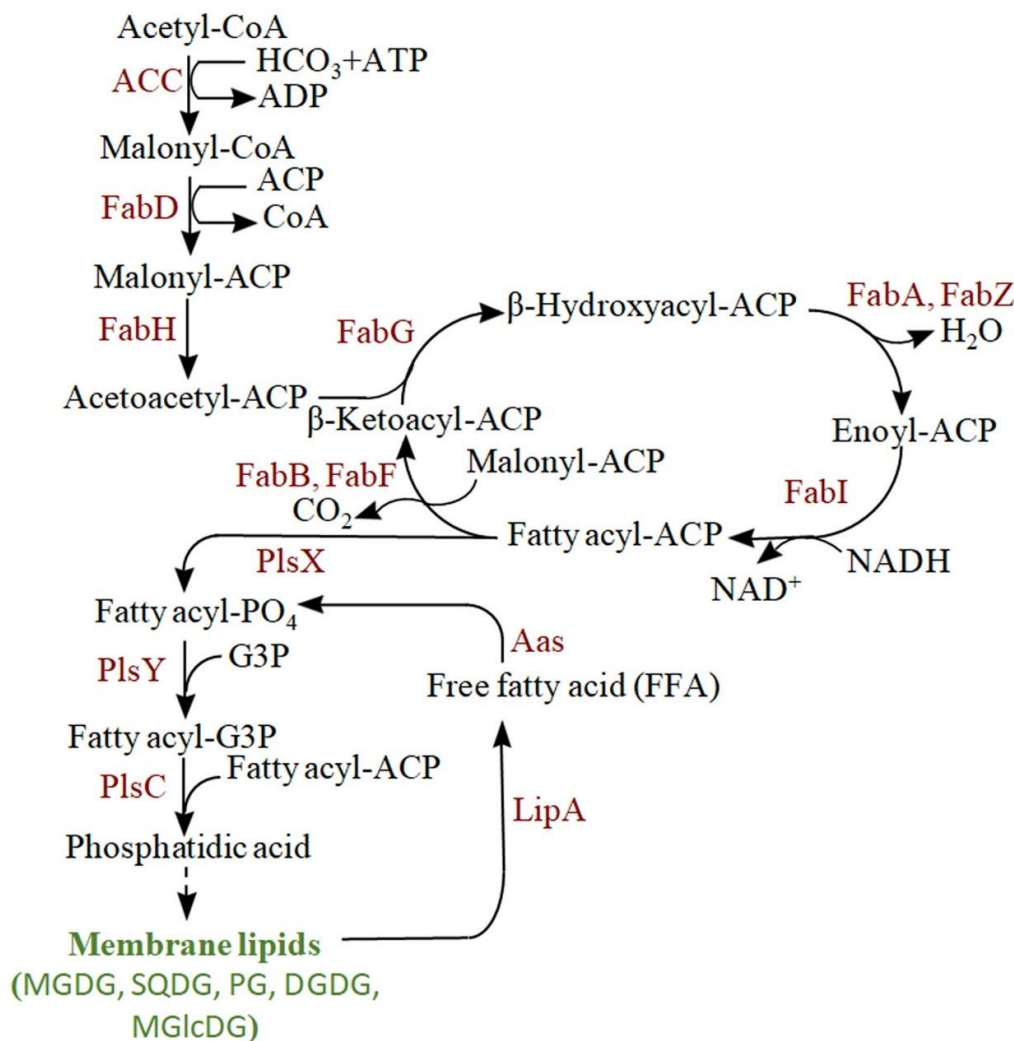


Fig. 3 Metabolism of lipid and fatty acid in cyanobacteria where MGLcDG: monoglucosyldiacylglycerol, MGDG: monogalactosyldiacylglycerol, DGDG: digalactosyldiacylglycerol, PlsC: putative 1-acyl-glycerol-P acyltransferase, PG: phosphatidylglycerol, SQDG: sulfoquinovosyldiacylglycerol, ACC: acetyl-coA carboxylase; ACP: acyl carrier protein. adopted with permission from (Velmurugan and Incharoensakdi 2022).



Bioremediation

Cyanobacteria are known for their remarkable adaptability to a variety of stress circumstances while also being quite resistant to hazardous substances from numerous sources (Rachedi et al. 2020). As a result, these photosynthetic bacteria are pertinent for a variety of bioremediation strategies, such as soil remediation, wastewater treatment, and the degradation of organic contaminants. As a result of anthropogenic activity, including urbanization, industrialization of the environment, as well as agricultural practices, water contamination constitutes a major environmental hazard. It has been demonstrated that brackish aquaculture effluent may be efficiently cleaned from ammonium using the marine *cyanobacterium* and *Synechococcus* sp. (Srimongkol et al. 2019). The usage of a cyanobacterial-bacterial consortium, which relies on the synergistic interaction between photosynthetic microorganisms and heterotrophic bacteria, is an intriguing method of biological wastewater treatment. It should be emphasized that these photosynthetic microbes are well-known for producing exopolysaccharides, which are essential for cyanobacteria to form symbiotic relationships with other organisms (Potnis et al. 2021). Cyanobacterial-bacterial aggregates were shown to be extremely efficient and successfully reduced the amounts of nitrogenous compounds, including nitrate (up to 80%), ammonium (up to 90%), and phosphorus compounds (up to 70%) in crude wastewater. These nitrogenous compounds included nitrate (up to 90%), and ammonium (up to 90%). The number of contaminants removed increased with the addition of an extra wastewater pretreatment stage including electrocoagulation and the use of an electrochemically treated supernatant as a substrate for the culture of microorganisms (Papadopoulos et al. 2020).

Cyanobacteria are a great choice for heavy metal removal due to a few characteristics. For instance, the characteristics of the cell wall, various transportation mechanisms, and the release of EPS (Javed et al. 2024). It has been observed that many cyanobacterial species can sequester heavy metal ions by biosorption, bioaccumulation, or, in many cases, a combination of the two. Certain species use EPS for biosorption to sequester heavy metal ions (Al-Amin et al. 2021). For example; *Anabaena doliolum* (Goswami, Syiem, & Pakshirajan, 2015), *Tolypothrixceytonica*, *Scenedesmus quadricauda* (Harris et al. 1990), *Cyanospira capsulate* and *Nostoc PCC7936* (De Philippis et al. 2003), *Cyanospiracapsulata* ATCC43193, *Cyanothece* (ET5, TI4, PE14, VI22, CE4) and, *Nostoc PCC7936* (Colica et al. 2010), *Gloeothece magna* (Mohamed, 2001), *Limnococcus* sp. (Sen et al. 2018), *Microcystis* sp. (Rai and Tripathi 2007), *Nostocmuscorum* (Roy et al. 2015). Various investigations discovered distinct functional groups in cyanobacterial EPS that are

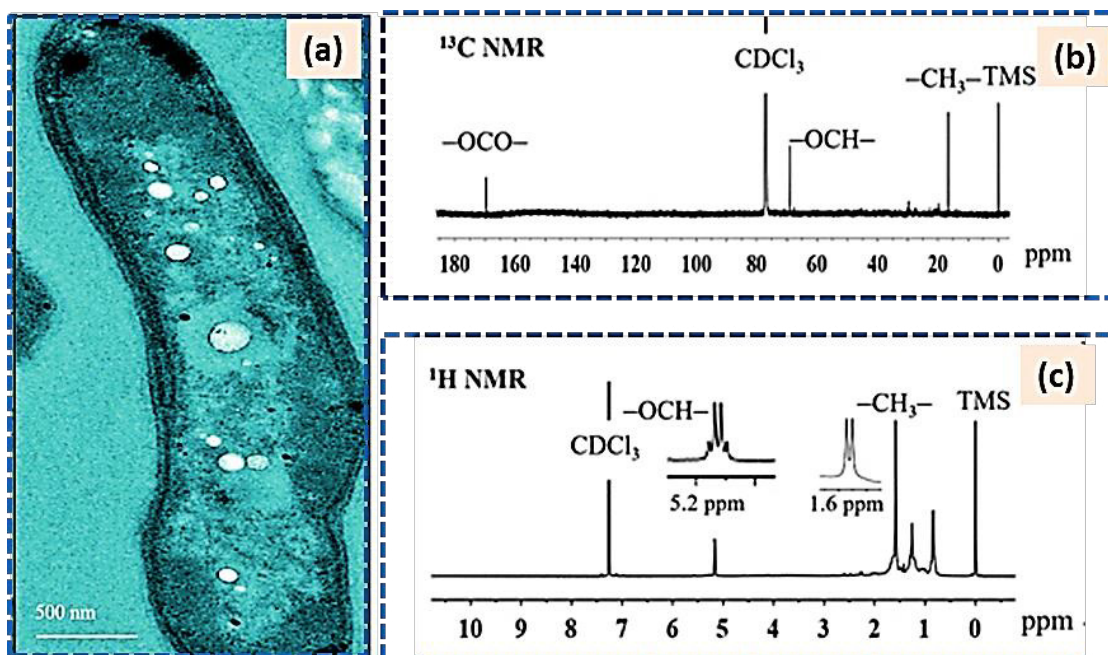


Fig. 4 TEM image (a), ^{13}C NMR (b), and ^1H (c) spectra of PLA homopolymer developed inside *S. elongatus* PCC7942. (Tan et al. 2022)



accountable for metal ion sorption. For example, Cu is bound to *Gloeotheca* sp. by the carboxyl and amide groups (De Philippis and Micheletti 2017). *Anabaena doliolum*'s carboxyl, sulfate, carbonyl, amide, and hydroxyl groups facilitate the biosorption of copper from water (Goswami et al. 2015). It was discovered that the hydroxyl and amide functional groups of *Synechocystis* sp. PCC6803 (Shen et al. 2021) and *Chlorella miniate* (Han et al. 2008) absorbed Cr(III) and Cr(VI). based on contact time, biosorption and bioaccumulation capacity, multi-metal removal efficiency, and efficiency of heavy metal removal, *Limnococcus* sp. (Sen et al. 2018), *Nostocmuscorum* (Roy et al. 2015), and *Synechococcus* sp. PCC 7942 (Rahman et al. 2011) are among the better prospects for heavy metal removal.

Production of alternative bioenergy

A substitute for the finite supply of fossil fuels in the future, hydrogen gas has been produced by cyanobacteria (Dutta et al. 2005; Milano et al. 2016). The benefits of using biological hydrogen as a fuel include its capacity to be recycled, efficiency, and lack of emissions of carbon dioxide during its manufacture and use. When heterocystous cyanobacteria containing nitrogenase are cultivated under nitrogen-limiting conditions, hydrogen is either produced as a byproduct of nitrogen fixation or by the reversible activity of hydrogenase enzymes in cyanobacteria (Mishra et al. 2019). Because of this, heterocystous cyanobacteria are more effective in producing hydrogen than non-heterocystous varieties (Pinzon-Gamez et al. 2005).

Bio-fertilizers

For their capacity to fix atmospheric nitrogen, heterocystous cyanobacteria and a few non-heterocystous cyanobacteria are well-known (Bhardwaj et al. 2024). Numerous tropical rice field soils' fertility has been largely credited to the presence of nitrogen-fixing cyanobacteria. it was estimated that cyanobacteria contributed 0.23 to 75.5 kg of nitrogen per hectare per year to the soils (Heimann and Cirés 2015). It has been tried and succeeded in injecting cyanobacteria into the soil to boost soil fertility. It was discovered that adding *Azolla* helped soil microorganisms, notably heterotrophic N₂ fixers, proliferate. According to recent reports, nitrogen-fixing cyanobacteria predominate in desert crusts all around the world. It was shown that the addition of *Azolla* promoted the growth of heterotrophic N₂ fixers and other soil microbes. Recent studies claim that cyanobacteria that fix nitrogen predominate in desert crusts all around the world (Garcia-Pichel and Pringault 2001)

Healthy food source

Cyanobacteria are considered one of the richest sources of vitamin B12 and contain more than 60% protein in addition to being high in beta-carotene, thiamine, and riboflavin. *Nostoc commune* has a high protein and fiber content, making it a valuable nutrient and physiological component of the human diet (Oregon, USA). Zeaxanthin, beta-carotene, canthaxanthin, and astaxanthin, among other cyanobacterial carotenoids, are excellent sources used in food supplements, colorants, food additives, and animal feed. These metabolites are being produced in greater quantities. The dietary supplements can be purchased as tablets, granules, or capsules. For instance, cyanobacteria like *Spirulina* produce the widely used vitamins alpha-carotene, riboflavin, vitamin B12, and thiamine. As a whole meal or dietary supplement, cyanobacteria are also known to contain minerals, amino acids, proteins, complex sugar, carbohydrates, phycocyanin, active enzymes, vital fatty acids, and chlorophyll (Markou et al. 2021).

Contrary to extracts used in pharmaceutical processes, varieties of nutritional supplements are often made from the biomass of cyanobacterial species (Marsan et al. 2018). For instance, ketocarotenoid (astaxanthin) is regarded as a potent antioxidant in comparison to vitamin A, vitamin C, and other carotenoids that play a critical role in preventing photooxidation-induced cell death in humans. Astaxanthin, a potent protease inhibitor produced by the bacterium *Haematococcus pluvialis*, has been used to treat several illnesses, including the human immunodeficiency virus disease (the virus that causes AIDS, the terminal stage of HIV disease) (Gregoire et al. 2022; Omatola et al. 2020).



Conclusions and future recommendations

This review has shown that cyanobacteria are potentially useful supplies for several biotechnology applications, particularly nanotechnology. Most of the commercially available chemicals were extracted from freshwater cyanobacteria, demonstrating the biotechnological potential of cyanobacteria. Because photosynthesis diverts the pathway for producing hydrogen, hydrogen productivity tends to decline at greater light intensities; therefore, the light regime needs to be carefully managed. Cyanobacteria have long been researched for their fascinating shape, diversity, and physiology, but groundbreaking work in recent decades has elevated these germs to a level where they are now favorably considered in biotechnology-related domains. Because of this, it is crucial to both comprehend and describe the diversity of cyanobacteria in as-yet unknown habitats as well as to profitably harness them for a range of industrial applications. Taxonomists, molecular biologists, biochemists, engineers, scientists with an interest in industry, and decision-makers in politics must all work together to achieve this. More studies are required on the best ways to use cyanobacteria in conjunction with environmentally friendly technological developments, as well as effective understandings of cyanobacteria applications at bigger scales. To investigate more advantageous uses of cyanobacteria in biotechnology, future research that is likewise promising is required.

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