

EXPLORING THE WORLD OF NANOGELS: PROPERTIES, APPLICATIONS AND FUTURE PROSPECTS

Bhargav E., Naresh Reddy Kami Reddy Midde* and Repollu Maddileti

Department of Pharmaceutics, Raghavendra Institute of Pharmaceutical Education and Research (RIPER)-
Autonomous, Anantapuramu -515721.

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*Corresponding Author

Naresh Reddy Kami Reddy

Midde

Department of Pharmaceutics,
Raghavendra Institute of
Pharmaceutical Education and
Research (RIPER)-Autonomous,
Anantapuramu -515721.

ABSTRACT

Nanogels are microscopic wonders, little hydrogel particles possessing remarkable qualities and an abundance of possible uses. Their unique properties make them a potential technology in many fields, such as medicine delivery, biotechnology, cosmetics, agriculture, and environmental science. They measure between 1 and 100 nanometers. Based on their unique specialties and their wide applications in various fields, this review focused on concept, characterization and advanced fabrication techniques, significance in nanotechnology, synthesis techniques, applications, challenges and future directions pertaining to Nanogels. This review also covered few highlights on case study and examples.

KEYWORDS: Nanogels, Nanotechnology, Synthesis techniques.

INTRODUCTION

Nanogels are real marvels of material science, and they stand out in the enormous field of nanotechnology, where little is beautiful and frequently transformative. With typical sizes between 1 and 100 nanometers, these nanoscale hydrogel particles have excited interest and curiosity across a wide range of scientific fields. The intriguing world of nanogels is explored in this introduction, which also explains their relevance, characteristics, and enormous promise in a variety of industries, including environmental science, drug delivery, biotechnology, cosmetics, and agriculture.^[1]

Concept of Nanogels

1. Understanding Nanogels

The definition of nanogels, which can be thought of as tiny hydrogel structures that are in turn three-dimensional networks of cross-linked polymers, is fundamental to this investigation. The extraordinary ability of these nanoscale gels to absorb and retain water or other molecules. They have unique characteristics that make them stand out in the field of nanomaterials due to their size and composition.^[2]

2. Significance and Context

The relevance of nanogels is found in their possible uses as well as their special qualities and adaptability. This little but powerful ones have gained importance in several scientific fields because of the following

A. Size and Scale

The small size of nanogels-typically at the nanoscale-allows them to interact at the cellular and even subcellular levels with biological processes. They can effectively enter tissues and cells thanks to this characteristic, which is advantageous for drug administration and diagnostics.

B. Composition and Structure

The cross-linked polymer networks that make up nanogels are often derived from biocompatible substances like chitosan or polyethylene glycol (PEG). Their stability is guaranteed by their structural composition, which also allows for personalization.

C. High Water Content

Water-rich nanogels have a remarkable capacity to both absorb and hold onto water. Because of this feature, which closely resembles the water content in biological systems, they are biocompatible.

D. Responsiveness and Control

The ability of nanogels to react to external stimuli, such as variations in pH, temperature, or ionic strength, makes them special. This responsiveness makes it possible to precisely control the release of the drug, which is an essential property in applications such as targeted drug delivery.^[3]

3. The Versatile Landscape of Nanogels

The versatility of nanogels unfolds as their applications span across a wide range of domains.

A. Drug Delivery and Targeted Therapy

The field of medication delivery is one in which nanogels show significant potential. They are skilled at controlling the release of medicines or therapeutic agents after they have been encapsulated. This characteristic is essential for tailored treatments, such as administering chemotherapy medications straight to the locations of tumors.

B. Gene Delivery and Gene Therapy

Nanogels are essential for gene delivery and gene therapy in the biotechnology industry. They open the door for advances in customized treatment by offering a secure and effective method of delivering genetic material to specific cells.

C. Imaging and Diagnostics

Nanogels high water content and tunable properties have a substantial impact on imaging and diagnostics. They can encapsulate imaging agents to enhance the contrast in various imaging modalities, allowing enhanced disease diagnosis and monitoring.

D. Cosmetics and Personal Care Products

Cosmetics and skincare items are among the consumer goods that use nanogels. They improve product performance and stability by encasing active substances, which benefits customers by producing better outcomes.

E. Agriculture and Crop Protection

Because they allow for the controlled release of fertilizers, growth regulators, and insecticides, nanogels have the potential to completely transform the agricultural industry. This method lessens its influence on the environment while simultaneously increasing agricultural yield.

F. Environmental Remediation

Nanogels are essential for eliminating heavy metals and contaminants from water sources in the environmental sciences. Because of their ability to absorb and neutralize dangerous compounds, they can provide answers to urgent.^[4]

Historical Perspective and Development of Nanogels

The amazing evaluation that nanogels-the wonders of nanotechnology and materials science have taken from conception to their wide range of current uses is truly astonishing. This story offers an examination of the historical backdrop and advancements in the field of nanogels, emphasizing significant turning points, discoveries, and the changes these hydrogel particles have undergone over time.^[5]

1. The Emergence of Nanogels

A relatively new discovery in the wide world of nanomaterials is nanogels. The domain of hydrogels, which have been researched and understood for many years, gave rise to the idea of nanogels. The extraordinary capacity of hydrogels-three-dimensional networks of hydrophilic polymers-to absorb and retain vast amounts of water, made them utilization in several fields. The applications in contact lenses, wound dressings, and other industrial and in other medicinal technologies were identified. On the other hand, nanogels are a relatively new technology, having been developed in the latter half of the 20th century.

2. The Pioneering Years

The synthesis and characterization of these novel materials was the subject of groundbreaking research during the early stages of the development of nanogels. During this time, several significant turning points occurred, as follows:

A. Early Synthesis Techniques

Researchers started looking into different ways to create nanogels. These techniques involved the use of sophisticated fabrication techniques, including microfluidics, as well as physical and chemical cross-linking.

B. Biocompatible Polymers

Materials such as polyethylene glycol (PEG) and chitosan have been used as a result of the hunt for appropriate biocompatible polymers. These polymers served as a precursor to biocompatible nanogels, an important advancement in the field of biomedical applications.

C. Initial Applications

During this time, comprehending the basic characteristics of nanogels and their possible uses took center stage. The foundation for today's wide range of applications was established by these early investigations.^[6]

3. Biomedical Applications and Drug Delivery

The use of nanogels for medication delivery is one of the most important advances in their development. The capacity of nanogels to encapsulate medications and release them in a regulated manner has demonstrated great promise in this field.

A. Chemotherapy and Targeted Drug Delivery

The discipline of cancer immediately became interested in nanogels. They were employed to specifically deliver chemotherapeutic medications to tumor locations by encapsulating them. This method of tailored medication delivery reduced systemic toxicity and improved the efficacy of cancer therapies.

B. Gene Delivery and Therapy

Furthermore, the advancement of gene delivery and gene therapy was greatly aided by nanogels. They opened up new avenues for the treatment of genetic illnesses by

offering a secure and effective method of delivering genetic material to target cells.^[7]

4. Characterization and advanced Fabrication techniques

More accurate and customized nanogel structures were produced by advancing characterization techniques and applying state-of-the-art production processes.

A. Characterization Methods

Researchers have been able to investigate nanogels with remarkable precision and get a better understanding of their size, appearance, and behavior thanks to the development of techniques including transmission electron microscopy (TEM), atomic force microscopy (AFM), and Dynamic Light Scattering (DLS).

B. Advanced Fabrication Techniques

Nanogels with exact characteristics, size, and structure might now be engineered thanks to the development of technology like electrospraying, 3D printing, and microfluidics.^[8]

5. Expanding Applications beyond Medicine

The uses of nanogels were not restricted to the medical domain. Their adaptability spanned across industries, inspiring advances across multiple domains.

A. Cosmetics and Personal Care Products

Nanogels are being used in skincare and cosmetic products sold to consumers. The stability and effectiveness of these products were enhanced by their capacity to encapsulate active components.

B. Agriculture and Environmental Science

In environmental science and agriculture, nanogels provide answers. They enhanced crop security and helped clean up the environment by permitting the controlled release of pesticides, fertilizers, and other contaminants.^[9]

The Significance of Nanogels in Nanotechnology:

Within the field of nanotechnology, nanogels—a subclass of nanomaterials—have a special and expanding importance. These 1-100 nanometer-sized hydrogel particles provide a wide range of characteristics and uses that are essential to many different industries. Understanding the importance of nanogels in nanotechnology and their potential to transform numerous fields is crucial in this context.^[10]

1. Size and Versatility

The significance of nanogels is largely determined by their nanoscale size. Due to its high surface area-to-volume ratio and compact size, there are greater chances for molecular and cellular interaction. They are useful in medical applications because of their ability to effectively enter tissues and cells.

2. Biocompatibility

Because they closely resemble biological systems' water content, nanogels are biocompatible. In biomedical applications, where materials must safely interact with living things, biocompatibility is an essential quality. For tissue engineering, drug delivery, and other medical uses, nanogels are a good fit.^[11]

3. Controlled Drug Delivery

The ability of nanogels to encapsulate medications and release them in a targeted and controlled manner is one of their most important contributions to nanotechnology. The realm of medicine has never seen anything like this regulated medication administration, which enables accurate dosing, decreased side effects, and greater therapeutic efficacy.^[12]

4. Environmental Solutions

Environmental nanotechnology is progressing with nanogels. They can be used to make ecosystems safer and cleaner by removing heavy metals and contaminants from water sources. Their ability to both absorb and immobilize dangerous substances is crucial for pollution treatment and control.^[13]

5. Advanced Fabrication Techniques

The precise engineering of characteristics and structure is made possible by sophisticated fabrication techniques like 3D printing and microfluidics, which are advantageous for nanogels. These methods enable nanogels to be customized for certain uses and requirements.^[14]

6. Smart Nanogels

A noteworthy breakthrough is the creation of smart nanogels. Even more accurate drug delivery is made possible by these nanogels' ability to react to particular biological stimuli, such as variations in pH or temperature. This characteristic makes targeted therapy and personalized medicine extremely important.^[15]

7. Diverse Applications

Biotechnology, cosmetics, agriculture, and environmental research are just a few of the disciplines in which nanogels find use. What makes them so useful in these fields is not only their adaptability and biocompatibility, but also their capacity to encapsulate and release chemicals.^[16]

Size and Morphology of Nanogels

As their name implies, nanogels are characterized by their minuscule size, usually spanning from one to one hundred nanometers in diameter. They are distinct from ordinary hydrogels, which are significantly larger due to their particular size. The dimensions of their nanoscale play a crucial role in shaping their form and characteristics.^[17]

1. Nanoscale Dimensions

Because of their nanoscale dimension, nanogels are unique. Their size gives them a particular edge by enabling them to function in a field where traditional materials cannot. Since they can easily penetrate tissues, cells, and even subcellular compartments, their small size makes them especially well-suited for applications that need contact with biological systems.

2. Morphology

Nanogels structure and form are referred to as their morphology. Cross-linked polymer chains form a three-dimensional network that defines nanogels. Depending on the fabrication process and particular design, this network produces a porous, gel-like material that might have a spherical, cylindrical, or more irregular shape.

The features of nanogels, especially their high water content and stimulus-responsiveness, are largely determined by their porous structure. One of the distinguishing characteristics of nanogels is their porosity, which permits the absorption and retention of water or other chemicals. Because nanogels can serve as carriers or absorbers of different chemicals, this property is very important in medicine delivery and environmental applications.

To sum up, the dimensions and structure of nanogels play a crucial role in characterizing their distinct characteristics and appropriateness for different uses. They are unique microscopic marvels with enormous potential in nanotechnology because of their nanoscale size, which makes them adaptable and effective in interacting with biological systems, and their three-dimensional, porous structure, which permits the absorption, retention, and controlled release of chemicals.^[18]

Composition and Structure of Nanogels

With their unique shapes and compositions, nanogels are an intriguing class of materials at the nanoscale. These tiny hydrogel particles are usually between 1 and 100 nanometers in size, and their distinct architecture makes them stand out in the field of materials science. The composition and structure of nanogels are examined in this talk, providing insight into the fundamental elements that enable their adaptability and usefulness in a range of settings.

Composition

To give them water-absorbing qualities, nanogels are made of a wide variety of polymers, most of which are hydrophilic. The polymer composition selected for a nanogel plays a key role in determining its responsiveness, biocompatibility, and application appropriateness. Often employed polymers in the creation of nanogels are Polyethylene Glycol, or PEG, is a polymer that is widely used because of its superior biocompatibility. It plays a crucial role in nanogels

intended for gene therapy, medication delivery, and other biological uses.

Polyacrylates: Nanogels are endowed with adjustable characteristics and structural stability by polymers based on acrylic acid. They are necessary to tailor drug release profiles into nanogels.

Chitosan is a polymer that is generated from chitin and is both biocompatible and biodegradable. It is frequently utilized in nanogels for purposes such as wound healing and medication delivery. Not only does the polymer selection affect the mechanical and chemical properties of nanogels, but it also affects their biocompatibility. They are appropriate for a variety of applications due to their versatility.^[19]

A three-dimensional network of cross-linked polymer chains defines the structure of nanogels. Nanogels are constructed using this network as the fundamental framework. The behavior and qualities of nanogels are largely determined by their structure. Among the crucial structural components are:

Cross-Linked Chains: Cross-linking polymer chains creates nanogels. The development of the three-dimensional network depends on these cross-links, which offer structural stability. The cross-links' type and density can change, which can affect the nanogel's porosity and reactivity.

Permeable and Expandable Matrix: The network of interconnected polymers in nanogels forms a permeable structure. Because of their porosity, they can take in and hold onto large volumes of water or other materials. A crucial component of medication delivery is the porous matrix, which permits the regulated release of chemicals that are encapsulated. The micelle nanogel formation depicted in Figure 1.

Size and Morphology: A nanogel's size and morphology are determined by the process used in its manufacture. Their shapes can be uneven, cylindrical, or spherical. They may be made to fit a specific application thanks to their adjustable size.

Nanogels are flexible and adaptive for a range of applications due to the combination of polymer content and structure. They have the ability to interact with biological systems, respond to external stimuli, and encapsulate and release chemicals.

The fundamental elements of nanogels composition and structure are what give them their extraordinary qualities and make them an invaluable tool in a variety of industries, from environmental research to medicine.^[20]

Swelling Behavior and Responsiveness of Nanogels

These tiny hydrogel particles, known as nanogels, are distinguished not only by their small size but also by their extraordinary capacity to display swelling behavior

and reactivity to outside stimuli. They are useful and adaptable for many different applications because of their special qualities, especially in medicine delivery and environmental cleanup.^[21]

Because of their porous, three-dimensional network, nanogels have the remarkable ability to absorb and hold large volumes of water or other chemicals. This is frequently called "swelling" behavior. The liquid that nanogels imbibe expands and swells their structure when they come into contact with it, whether it is in a solvent-rich environment or not. A number of variables, including the solvent type, cross-link density within the network, and nanogel composition, affect how much the swelling occurs.

This swelling tendency is important for a number of applications. Nanogels, for example, can be filled with therapeutic ingredients for medication delivery. The encapsulated medicine is released gradually and under control thanks to its ability to swell in the body's aqueous environment. This regulated release reduces adverse effects and guarantees long-lasting therapeutic benefits.^[22]

Responsiveness to External Stimuli

The phenomena of nanogel depicted in Figure 2. The capacity of nanogels to react to variations in temperature, ionic strength, pH, or other external stimuli is one of their special qualities. Often included in the structure of the nanogel, this responsiveness is especially important for developing "smart nanogels".^[23]

1. pH-Responsive Nanogels

In reaction to pH variations, nanogels can be designed to swell or deswell. For example, a nanogel might be engineered to expand in an acidic milieu (such as inside a tumor) and release its therapeutic payload precisely where it is needed, reducing the amount of medication that is exposed to healthy cells.^[24]

2. Temperature-Responsive Nanogels

Temperature-sensitive nanogels respond differently to changes in temperature in terms of swelling. This characteristic makes nanogels indispensable for uses such as hyperthermia therapy, in which medications are targeted to and released from specific body locations when subjected to high temperatures.^[25]

3. Ionic Strength-Responsive Nanogels

In addition, nanogels can adapt to variations in ionic strength by swelling or deswelling in response to changes in the concentration of salt in their immediate surroundings. Applications of this responsiveness include protein separation procedures and controlled medication delivery.

High levels of precision are possible in a variety of applications, such as medication delivery, tissue engineering, and environmental remediation, because of

nanogels' capacity to be specifically tailored to respond to environmental cues.

A key component of nanogels importance in nanotechnology is their capacity to swell and respond. Due to their ability to release compounds under controlled conditions, these characteristics are essential for medication delivery and other medical applications. Their flexibility in a variety of domains, including biotechnology, environmental research, and nanomedicine, where accuracy, effectiveness, and control are critical, is further ensured by their susceptibility to environmental signals. Because they are adaptable nanostructures, nanogels help close the gap between state-of-the-art nanotechnology and real-world applications for challenging problems.^[26]

Synthesis of Nanogels

Chemical Cross-Linking Methods

Chemical cross-linking is one of the most common ways used to create nanogels, which are adaptable and versatile nanostructures. Chemical cross-linking creates a three-dimensional network that encases water or other things by forming covalent links between polymer strands. Here, we discuss the techniques for chemical cross-linking that are utilized to create Nanogels.^[27]

1. Emulsion Polymerization

One popular technique for creating nanogels is emulsion polymerization. This procedure involves dispersing water-insoluble monomers, which are frequently compounds that resemble oil, in an aqueous phase that contains surfactants. After that, these monomers polymerize to create nanogels in the aqueous phase. In order to make covalent bonds between the polymer chains and construct the gel network, cross-linking agents are introduced. This process makes nanogel size, composition, and structure precisely controllable, which makes it appropriate for a wide range of uses.

2. Free Radical Polymerization

Another popular technique for creating nanogels is free-radical polymerization. It uses free radicals to start the monomer polymerization process. Free radicals are typically produced by mixing a monomer solution with an initiator and a cross-linker. These radicals help the polymer chains create covalent bonds with one another, which results in the production of nanogel. One may customize the properties of the nanogel by varying the monomers, cross-linkers, and initiators, which makes it a versatile approach for a range of applications.^[28]

3. Reverse Microemulsion Polymerization

Reactants are contained in nanoscale water droplets inside an oil phase in a specialized process called reverse microemulsion polymerization. The creation of highly homogenous nanogels with regulated sizes is encouraged by this enclosed environment. Water droplets are mixed with cross-linking chemicals, which cause polymer

chains to covalently connect and eventually form nanogels with special characteristics.^[29]

4. Cross-Linking in Solution

By cross-linking previously manufactured polymers in a solution, this technique creates nanogels. The production of covalent bonds and the generation of the nanogel structure are the results of mixing polymer chains with a cross-linking agent in a solvent. This method works well when a certain application calls for a certain polymer or copolymer.^[30]

5. Self-Assembly and Cross-Linking

Molecules or particles spontaneously organize into organized structures through the use of self-assembly processes. Cross-linking self-assembled structures can help to stabilize and solidify the nanogel network throughout the nanogel production process. This method's advantage is that it takes advantage of some molecules' innate ability to arrange themselves into nanoscale shapes.

With the help of these chemical cross-linking techniques, one can synthesize nanogels with a wide range of tools, allowing for customization of their size, shape, and characteristics. The method of choice is determined by the intended use and the particular qualities that the nanogels must have. With their precision, efficiency, and versatility in a range of applications, nanogels are set to become increasingly important as these techniques advance in the domains of biotechnology, drug delivery, and other areas.^[31]

Physical Cross-Linking Methods

Several methods can be used to create nanogels, which have special qualities and a wide range of uses. In contrast to chemical procedures that establish covalent bonds, physical cross-linking techniques rely on non-covalent interactions to form the three-dimensional network of nanogels. Simplicity and a lower chance of hazardous chemical residues are two of the benefits of these techniques. Here, we explained some physical cross-linking techniques that are frequently employed in the creation of Nanogels.^[32]

1. Ionic Gelation

One common technique for physical cross-linking is ionic gelation. Cross-links between polymer chains are created by the interaction of ions with opposing charges. Polymers that have ionizing functional groups in solution are typically employed. The nanogel structure is formed by the formation of links between the polymer chains by a cross-linker containing counterions of the opposite charge. This approach is used for many nanogel compositions due to its ease of use and adaptability in many applications.

2. Hydrogen Bonding

A key process in the creation of nanogels is hydrogen bonding. Functional groups like hydroxyl groups, which

can form hydrogen bonds, are found in polymers. These functional groups have the ability to create hydrogen bonds with other groups on the same polymer chain or on separate ones. The creation of a stable nanogel structure is the outcome of this dynamic and reversible interaction.^[33]

3. Physical Cross-Linking by Temperature

Certain nanogels are engineered to experience a reversible phase transition in reaction to variations in temperature. By choosing polymers with a lower critical solution temperature (LCST), physical cross-linking produced by temperature is accomplished. The creation of the nanogel structure results from the polymers being less soluble and aggregating as the temperature is raised over the LCST. This characteristic is useful in situations where temperature variations can be utilized to initiate the release of drugs, such as drug delivery.^[34]

4. Physical Cross-Linking by pH

Some nanogels are made to go through pH-responsive phase changes, which is comparable to temperature-induced cross-linking. It is possible to control the charge on polymers having ionizable functional groups by varying the pH of the surrounding solution. The polymer chains combine and create the nanogel structure as a result of this change in charge. pH-responsive nanogels find utility in medicine delivery and other fields where controlled release in response to pH variations is necessary.^[35]

5. Host-Guest Interaction

Guest molecules are encapsulated within the nanogel structure during host-guest interactions. The guest molecules are incorporated into the nanogel network to create reversible physical cross-links that stabilize it. One example of this approach is the way cyclodextrins interact with visiting molecules.

Physical cross-linking techniques have clear benefits, particularly in drug delivery applications where covalent interactions are not desired. The versatility of nanogels in various applications is further enhanced by their capacity to react to changes in environmental parameters such as pH and temperature. Nanogels are still a useful tool in many scientific and industrial fields because of these ongoing technologies' expansion and improvement.^[36]

Advanced Fabrication Techniques for Nanogels

Nanogels have become more well-known in a number of sectors, such as medication delivery, biology, and materials research, because of their distinctive qualities and nanoscale size. Precision and control over the tailoring of nanogels are largely dependent on sophisticated fabrication procedures. These methods enable scientists and engineers to produce nanogels with particular characteristics, dimensions, and configurations, allowing for a wide range of application possibilities. The following are a few of the sophisticated techniques for creating nanogels.

1. Microfluidics

The novel field of microfluidics manipulates fluids at the micrometer scale by means of minuscule chambers and channels. Microfluidics plays a key role in the development of regulated sizes, shapes, and characteristics of nanogels by enabling the precise mixing of reactants and polymers. When making monodisperse nanogels, which are crucial for drug delivery and nanomedicine, it is especially helpful.^[37]

2. Electrospaying

With the flexible method of electrospaying, polymer solutions are atomized into tiny droplets using an electric field. After that, these droplets can be cross-linked to create nanogels. Drug delivery and tissue engineering applications can benefit from electrospaying's ability to produce nanogels with narrow size distributions and excellent encapsulation efficiency.^[38]

3. 3D Printing

A rapidly developing technology, three-dimensional (3D) printing, has found its way into the production of nanogels. Researchers are able to precisely regulate the size and shape of the intricate nanogel structures they make by employing specialized 3D printers. With the use of this technique, nanogels may be created to replicate the characteristics of particular tissues and organs, which is useful in tissue engineering.^[39]

4. Supercritical fluids

Supercritical fluids are employed in the controlled manufacture of nanogels as both solvents and anti-solvents. Rapid expansion of supercritical solutions (RESS) or gas antisolvent (GAS) approaches can create nanogels by varying the pressure and temperature. Making nanogels for medical and environmental applications is a particularly helpful use of this technique.^[40]

5. Self-Assembly techniques

The process of self-assembly is based on the innate characteristics of particular molecules or particles, which enable them to spontaneously arrange into predetermined forms. When it comes to nanogels, self-assembly methods enable the creation of nanogels without the need for outside energy sources. The creation of Nanogels-which are utilized in biotechnology and medicine delivery-that imitate natural processes depends on this technique.

Thanks to these sophisticated production procedures, researchers may customize nanogels to precisely match the needs of particular applications. These methods are advancing the accuracy and adaptability of nanogels in the rapidly changing field of materials science and technology, whether it is by producing monodisperse nanogels for drug delivery, structurally complex nanogels for tissue engineering, or environmentally responsive nanogels for pollution control.^[41]

Advantages

High Water Content and Biocompatibility of Nanogels

They are the perfect option for applications in tissue engineering, drug delivery, and biomedicine because of their unique combination of nanogels. They are useful tools for tackling difficult problems in biotechnology and healthcare because of their capacity to effectively encapsulate and release water-soluble compounds while still being compatible with biological systems. In the realm of nanomaterials, nanogels represent the promise of accuracy and biocompatibility. A unique combination of high water content and biocompatibility characterizes nanogels, which are microscopic hydrogel particles with dimensions on the nanometer scale. Their wide range of applications and versatility, particularly in drug administration and biomedicine, can be attributed in large part to this combination.

High Water Content

The water absorption and retention capabilities of nanogels are well known. Their cross-linked polymer chains form a three-dimensional network that is intrinsically high in water content. Nanogels easily absorb water when they come into contact with aquatic environments, which cause their structure to inflate. The ability of hydrogels to encapsulate and release water-soluble compounds is largely attributed to their intrinsic swelling tendency. Because nanogels can be loaded with medications or therapeutic agents and release them in a regulated and sustained manner, this characteristic is especially useful in drug delivery. The high water content enhances the effectiveness of drug delivery systems by enabling a high payload of medications or biomolecules.^[42]

Biocompatibility

The ability of a material to be biocompatible is a prerequisite for using it in living things. The composition of nanogels is rich in water, which contributes to their high level of biocompatibility. Their water content is compatible with live tissues and cells since it is similar to that of biological systems. Furthermore, the biocompatibility of the polymer materials-such as chitosan or polyethylene glycol (PEG)-used to create nanogels is a deciding factor. This polymer's lengthy history of use in pharmaceutical and medical applications further supports nanogels' biocompatibility.

In the context of medication distribution, the combination of high water content and biocompatibility is especially beneficial. Nanogels can pass through tissues and cells in the body without endangering the organism or triggering an immune reaction. This characteristic is essential for the effective delivery of medications and therapeutic agents to tumors and other target areas without negatively impacting healthy tissues. Additionally, because of their high water content and biocompatibility, which are essential for interacting with biological systems and promoting tissue growth and

repair, nanogels are employed in tissue engineering, wound healing, and regenerative medicine.^[43]

Tunable Drug-Loading Capabilities of Nanogels

The key to nanogels' efficacy in drug delivery applications is their ability to load drugs in a customizable manner due to their distinct characteristics. This characteristic enables scientists and medical professionals to carefully regulate the dosage of medication encapsulated in nanogels, maximizing therapeutic efficacy and reducing adverse effects.

1. Size and Porosity

It is possible to modify the size of nanogels to fit particular medications. Due to their increased surface area in relation to volume, smaller nanogels have higher drug-loading efficiency, whereas bigger nanogels may carry more pharmaceuticals overall. The ability of nanogels to hold drugs is further improved by their porous shape. The drug-filled pores serve as reservoirs, allowing for gradual and regulated release.

2. Cross-Link Density

It is possible to modify the cross-link density in nanogels, which will impact the distance between polymer chains. A more compact structure with less room available for drug encapsulation is produced by a greater cross-link density. Larger drug molecules can be accommodated in a more porous framework with lower cross-link densities.

3. Hydrophilicity and Hydrophobicity

With the right polymer selection, nanogels may be made more hydrophilic or hydrophobic. Drugs that dissolve in water are more suitable for hydrophilic nanogels, whereas lipid-based medications are better suited for hydrophobic nanogels. Drug compatibility and loading efficiency can be improved by selecting hydrophilic or hydrophobic polymers.

4. Functional Groups

It is possible to build nanogels with certain functional groups that form chemical interactions with medicines. By creating links that keep pharmaceuticals within the nanogel network, these interactions can improve the capacity of the medication to be loaded.

Nanogels adjustable drug-loading properties are extremely important for personalized medicine, as they allow patients to get customized therapies depending on their unique medication needs. It also lessens negative effects and lowers the possibility of overdosing. For example, nanogels may be designed to precisely deliver larger quantities of chemotherapeutic medications to tumor areas while preserving healthy tissues, a technique used in cancer therapy.

Because of these qualities, nanogels are effective drug delivery agents that offer the accuracy required to maximize therapeutic results and improve patient

comfort. The adaptability and promise of nanogels in enhancing pharmacological interventions and healthcare are highlighted by their capacity to customize drug loading based on the individual medical condition being treated and the qualities of the medication.^[44]

Controlled Release Mechanisms of Nanogels:

A key component of drug delivery systems is controlled drug release, which guarantees that therapeutic drugs are delivered accurately, progressively, and steadily to maximize efficacy and minimize adverse effects. Because of their special qualities and versatility, nanogels provide a number of controlled release mechanisms that are essential for drug delivery applications.

1. Swelling and Deswelling

Temperature and pH are two environmental variables that have a significant impact on nanogels. Nanogels absorb water and expand, releasing the medications they contain when exposed to circumstances that cause swelling. On the other hand, when deswelling occurs, they shrink and hold onto the medication. Nanogels are very useful for targeted therapy because of their ability to swell and deswell in response to certain physiological stimuli, allowing for regulated drug release.

2. Diffusion-Controlled Release

Pharmaceuticals can be released from nanogels via a diffusion-controlled process in which the pharmaceuticals move through the porous nanogel network. The concentration gradient, the density of the nanogel network, and the size of the drug molecules all affect how quickly the drug is released from the body. Nanogels are able to accurately regulate the kinetics of drug release by manipulating these factors.

3. Stimuli-Responsive Release

Certain nanogels are made to react to outside stimuli, such as temperature or pH variations. For example, medications can be released via pH-responsive nanogels in reaction to changes in acidity, which are frequently linked to disease states like cancer. When subjected to high temperatures, as can be done with hyperthermia treatment, temperature-responsive nanogels release medication.

4. Triggered Release

It is possible to construct nanogels with certain triggers, like enzymes, or external energy sources, like ultrasound or light. The payload of the nanogels is released upon the application of these triggers. Since some enzymes may be present in illness areas, enzyme-triggered nanogels are particularly useful in the context of targeted treatment.

5. Time-release profiles

Therapeutic demands can be met by customizing the cross-linking density, content, and structure of nanogels to achieve desired drug release patterns. Nanogels can be

made to release medications quickly, steadily, or gradually, guaranteeing that prescriptions are taken in accordance with the prescribed schedule.

Drug distribution can be done with a high degree of accuracy thanks to nanogels' controlled release mechanisms. This is especially crucial in personalized medicine, where medication release may be customized to meet the demands of each patient, and in applications like cancer treatment, where focused drug delivery reduces negative effects on healthy tissues. Nanogels are essential instruments for maximizing therapeutic results in the field of drug administration because of their versatility and reactivity.^[45]

Applications of Nanogels

Nanogels in Drug Delivery and targeted therapy

When it comes to drug delivery, nanogels have become extremely useful instruments since they allow for focused, precise, and regulated treatment. They are the perfect carriers for therapeutic drugs because of their special qualities, which include high water content, biocompatibility, and stimuli-responsive behavior—especially when used in targeted therapy.

1. Controlled Drug Delivery

Long-term, regulated medication release is made possible by nanogels. Precise medication delivery is made possible by their characteristic of swelling and deswelling in response to physiological signals such as pH and temperature. In addition to reducing adverse effects, this regulated release profile guarantees that medication is given at therapeutic doses.

2. Targeted Drug Delivery

Drugs are delivered exactly where they are required because of the ability of nanogels to target particular locations in the body. This is especially important for diseases like cancer, where targeted therapy can improve treatment outcomes while protecting healthy tissues. Drug release may be regulated at the appropriate location and time by engineering nanogels to react to certain disease indicators or environmental cues.

3. Personalized Medicine

The use of nanogels in medicine may be individualized. Nanogels may be tailored to each patient's specific needs by adjusting target-specific mechanisms and medication release patterns. This degree of accuracy is very useful for maximizing treatment results and reducing side effects.

4. Overcoming Biological Barriers

Drug effectiveness can be circumvented by nanogels by overcoming biological barriers. Due to their tiny size, they are able to enter tissues and cells and deliver medications to intracellular destinations that would otherwise be difficult to reach.

5. Improved Pharmacokinetics

By improving drug solubility and bioavailability, hydrophobic medications that would normally be difficult to give may now be delivered thanks to nanogels. This guarantees that the therapeutic action of medications is maximized and enhances their general pharmacokinetics.

6. Combination therapy

Combination treatment is made easier by the ability of nanogels to transport several medications at once. This is especially helpful for complicated illnesses when a multi-drug approach is more beneficial than a single-agent approach. Because they offer a degree of control and accuracy that was previously unachievable, nanogels have the potential to completely transform the delivery of pharmaceuticals. Their significance in developing healthcare and enhancing patient outcomes is highlighted by their roles in personalized medicine, targeted therapy, and overcoming biological obstacles.^[46]

Gene delivery and gene therapy

Nanogels hold great potential as gene delivery and gene therapy carriers. They are ideal for these uses because of their high water content, biocompatibility, and capacity to preserve genetic material. They may safely transfer DNA or RNA into cells by effectively encasing and shielding them. Furthermore, by designing nanogels to react to certain biological stimuli, targeted gene delivery for the treatment of hereditary illnesses or regulation of gene expression is made possible. In order to maximize on-target effects and guarantee the intended therapeutic benefits, accuracy is essential in gene therapy. By improving the transport and therapeutic effectiveness of genetic materials, nanogels have the potential to completely transform gene therapy.^[47]

Imaging and diagnostic applications

Nanogels have demonstrated potential in diagnostic and imaging applications. Nanogels can function as contrast agents for MRIs, CT scans, and optical imaging by enclosing imaging agents, such as fluorescent dyes or contrast agents, inside their structure. They are appropriate for *in vivo* imaging because of their tiny size and biocompatibility. Furthermore, nanogels may be engineered to react to certain biological stimuli, allowing for targeted imaging of particular organs or illnesses. This accuracy lowers the need for intrusive treatments while improving diagnostic accuracy. Nanogels have a lot of promise to advance non-invasive diagnostics and imaging in medicine.^[48]

Challenges and Future Directions

Biocompatibility and toxicity concerns

Because of their water-rich composition and the biocompatible polymers used in their manufacture, nanogels have a high level of biocompatibility. They lessen the possibility of immunological reactions or negative side effects since they closely mimic the aqueous environment of biological systems. But worries

regarding possible toxicity are still there, especially when it comes to impurities, the release of compounds that are contained, and the materials utilized to fabricate nanogels. To allay these worries and guarantee that nanogels fulfill safety requirements for use in medical and environmental applications, rigorous testing and quality control are necessary. The goal of ongoing research is to improve nanogel safety and biocompatibility, increasing its value across a range of industries.^[49]

Regulatory aspects and commercialization challenges

Despite their potential, nanogels encounter difficulties with regard to safety assessment and categorization under regulations. These cutting-edge materials can take longer to get approved for commercial or medical usage by regulatory authorities since they frequently find it difficult to adjust to them. Another difficulty with standardizing is the variety of nanogel formulations available.

The challenges associated with commercialization are manufacturing scalability, cost-effectiveness, and competition from pre-existing drug delivery systems. Commercialization may also be hampered by concerns about intellectual property and the requirement for long-term clinical validation. In order to traverse the complicated environment of nanogel-based goods and therapies, researchers, industry, and regulatory bodies must work together to overcome these hurdles.^[50]

Future prospects for nanogels

Nanogels have very bright futures in a number of different sectors. They might revolutionize medication delivery, gene therapy, and diagnostics in medicine by providing precise, focused treatments with fewer adverse effects. Nanogels have the potential to be extremely important in tissue engineering and regenerative medicine, advancing the creation of sophisticated biomaterials.

Nanogels' special qualities might be advantageous for environmental applications like pollution management and water purification. We foresee a wider commercial presence as nanogel research advances, resolving regulatory obstacles and improving scalability. Because of their capacity to respond to a variety of stimuli and their potential for tailored therapy, nanogels are poised to play a significant role in the development of healthcare and technology in the future. They will help solve difficult problems and enhance people's quality of life.^[51]

Case Studies and Examples

By showcasing particular uses and encouraging tales Nanogels have demonstrated impressive results in a variety of applications, especially in the fields of biotechnology and medicine. When it comes to medication distribution, nanogels are excellent at precisely delivering therapeutic ingredients while reducing adverse effects. For example, research that was

published in the "Journal of Controlled Release" showed how pH-responsive nanogels may be used to successfully transport anticancer medications to tumor locations in a targeted manner, improving therapeutic efficacy and lowering systemic toxicity.

Nanogels have advanced significantly in the field of gene therapy. They have been employed by researchers as genetic material carriers in an effort to deliver genes precisely and effectively. A study published in "Biomaterials" demonstrated how well cationic nanogels work to transport siRNA for gene silencing, highlighting the possibility of using them to treat genetic problems. Imaging applications have also shown success using nanogels.

An article published in "ACS Applied Materials & Interfaces" described the creation of imaging agent-loaded nanogels for improved contrast in magnetic resonance imaging applications. They are useful instruments in non-invasive diagnostic imaging because of their tiny size and capacity to target certain tissues. Tissue engineering is starting to use nanogels for biotechnological purposes. Nanogels have been used by researchers as scaffolds for tissue regeneration and cell development.

In an investigation that was published in "Biomacromolecules," growth factor-loaded nanogels improved tissue regeneration and repair. These achievements highlight the adaptability and promise of nanogels in solving challenging problems in biotechnology, medicine, and diagnostics. Nanogels have the potential to significantly impact targeted therapy, customized medicine, and advanced biomaterials as research and development proceed, hence further influencing the field of creative applications in the years to come.^[52]

Research breakthroughs and their impact

Recent advances in nanogel research have had a profound effect on a number of industries, opening up new avenues and extending existing uses. The creation of stimuli-responsive nanogels, which enable precise and regulated drug delivery, is one noteworthy innovation.

In a ground-breaking work that was published in "Advanced Drug Delivery Reviews," scientists created nanogels that react to particular stimuli, such as temperature or pH changes. This responsiveness minimizes off-target effects by enabling precise medication release at specific places inside the body. This discovery has significant implications for personalized medicine, as maximizing treatment results requires customized medication delivery.

A further innovation is the combination of sophisticated imaging methods with nanogels. A study that was published in "Nano Letters" showed how contrast agents may be successfully added to nanogels for improved

imaging modalities, including optical and magnetic resonance imaging. This innovation improves diagnostic imaging precision and provides a more lucid understanding of physiological functions and illnesses.

A seminal paper published in "Nature Communications" demonstrated the creation of nanogels as effective delivery systems for CRISPR gene-editing instruments in the field of gene therapy. The genetic material was shielded by the nanogels and was more easily delivered to the intended cells. This discovery has enormous potential for improving gene therapy and curing hereditary illnesses.

These discoveries have an influence that goes beyond the lab; they have an impact on clinical procedures and the direction of healthcare. Drug delivery may now be approached with unprecedented accuracy thanks to

stimuli-responsive nanogels. By combining nanogels with cutting-edge imaging technologies, diagnostic capacities are improved, enabling better illness diagnosis and monitoring. The effective application of nanogels in gene therapy creates new opportunities for molecularly treating hereditary illnesses.

The commercialization of medicines and technologies based on nanogels is made possible by these developments. We should expect more scientific advances that will continue to influence the field of nanogel applications, ranging from gene editing and targeted medication delivery to sophisticated diagnostics. These developments are not only altering the science of nanotechnology but also have the potential to completely change how patients are treated and how treatments are administered.^[53]

FIGURES

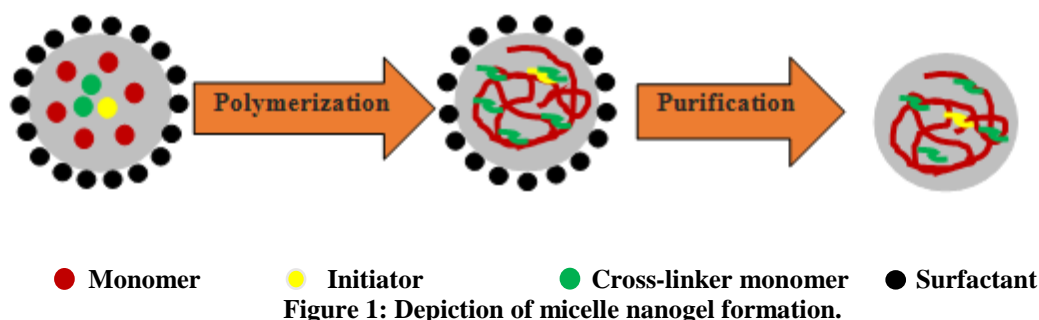


Figure 1: Depiction of micelle nanogel formation.

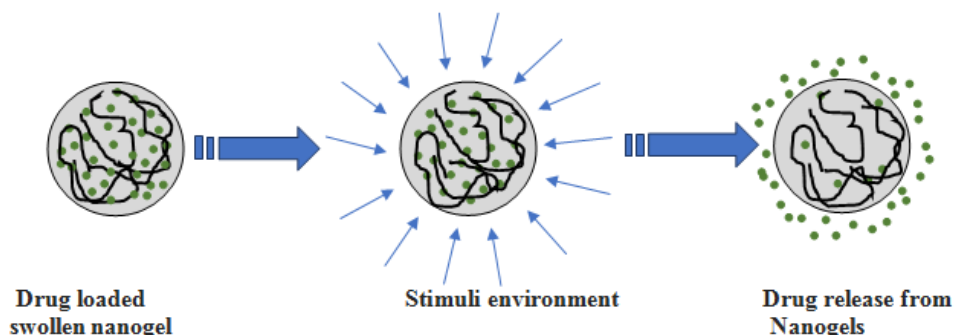


Figure 2: Depiction of various phenomena of nanogels.

CONCLUSION

Finally, nanogels demonstrate unmatched adaptability and promise, establishing them as game-changing players across a range of scientific fields. They can perform very well in targeted medication administration, gene therapy, diagnostics, and other fields thanks to their special qualities, which include high water content, stimulus responsiveness, and biocompatibility. Recent developments highlight their accuracy and versatility in various applications, providing answers that were previously difficult to obtain. The numerous benefits that

nanogels offer are further highlighted by their achievements in tissue engineering, imaging, and regenerative medicine.

The potential of nanogels to transform customized treatments and tackle intricate healthcare issues is becoming more and more apparent as research on them progresses. These nanostructures not only represent a major advancement in nanotechnology but also have the potential to completely change the way that technology and medicine are developed in the future by offering

specialized solutions that will improve patient care and treatment outcomes.

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