**A Review of Hydrogels their Classifications, and Applications**

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**Abstract**

A hydrogel is an insoluble network of polymers in three dimensions that has the ability to absorb biological fluids. Such a polymer network is produced by chemical and physical cross-linking processes. Covalent forces create chemical hydrogels, whereas weak secondary forces create physical hydrogels. Hydrogels can be made from a wide range of synthetic and natural polymers. The swelling, mechanical, and biological capabilities of hydrogels are its most important features; these attributes all affect the hydrogel's morphology and structure. Hydrogel finds its application in several fields such as wound dressings, tissue engineering, contact lenses, adsorbents, sensors, and medicine due to its ability to absorb water and structural similarity to extracellular matrix (ECM). This review covers hydrogels, their varieties, and their uses

.**1 Introduction**

**1.1 hydrogels**

Hydrogels have been defined differently by researchers over time. For example, they are large-scale hydrogel networks that retain a significant volume of water within their structure despite being insoluble in it. The absorbance and water retention properties of a hydrogel are attributed to the hydrophilic functional groups that are linked to the polymeric network. Conversely, the polymeric chains' resistance to dissolution can be attributed to the cross-links that exist between them. A hydrogel is characterized by the presence of two or more components in a three-dimensional polymeric chain network. Networks of polymeric chains, or hydrophilic gels, are typically colloidal gels with water acting as a dispersion medium.1-2 The most frequent definition of hydrogels is a cross-linked polymeric chain that swells when exposed to water. And are produced by a straightforward reaction involving one or more monomers. Its ability to stay three-dimensional throughout the swelling phase is due to crosslinking.3. Hydrogels have garnered significant interest in the last fifty years because of their outstanding potential in a wide range of applications.4. Hydrogels are able to hold huge amounts of water and fluids, including biological fluids that may imitate biological tissues, and as a result, their flexibility is extremely comparable to that of genuine tissue. This property has sparked a great deal of interest in creating novel devices by adjusting their tunable physicochemical properties. Synthetic hydrogels, which have unique architectures and can produce variable functioning and degradation with further modification, have largely superseded natural hydrogels in recent times. Depending on the characteristics of the parts they can contain different amounts of water in equilibrium depending on the characteristics of the components used in the polymeric network and the density of the network joints.5.

Hydrogel synthesis can be accomplished through various chemical techniques, including one-step methods such as polymerization and cross-linking of multifunctional monomers, as well as multiple-step methods that entail the synthesis of polymeric chains with highly reactive functional groups, which are then cross-linked using an appropriate cross-linking agent. A polymer engineer is able to create polymeric networks with customized qualities such as mechanical capabilities, biological and chemical reaction to stimuli, biodegradation, and molecular-scale control over structure, including cross-linking density.6

**1.2 Natural Gums based Hydrogels**

Because of their exceptional qualities, affordability, structural variety, and accessibility, natural gum polysaccharides have drawn the attention of researchers in a variety of fields, including the water, food, energy, medicine, and biotechnology industries. Natural gums, also known as polysaccharides7, are derived from a variety of tree species and have remarkable qualities such as being renewable, biodegradable, biocompatible, non-toxic, and easily chemically modifiable.7. When compared to synthetic origin, natural gum-based hydrogels or polysaccharides offer some advantageous features. Since the uncontrolled use of hydrogels manufactured of synthetic polymers has resulted in health, ecological, and environmental issues, they have seen notable progress as a fresh alternative in recent years.8, 9, As a result, there is a high demand for environmentally friendly materials. These hydrophilic polymeric networks have greater strength and elasticity and are insoluble in water.1, 10 They exhibit remarkable reactivity to changes in temperature, pH, electric field strength, solvent composition, pressure, and solvent composition.12, 13, These hydrated polymeric networks are becoming increasingly important in practice and are useful as biomimetic, intelligent, and intelligent materials. They find use as actuators and sensors, and they are frequently being researched as self-oscillating gels.14. Hydrogels known as "smart networks" react physiochemically significantly to even minute alterations in their environment. These changes are reversible; they can return to their initial state if the trigger is removed.15.

**1.3 Classification of Hydrogels**

 Depending on the source, they might have an artificial or natural origin. Natural polymers include gums, agarose-formed hydrogels, starch, cellulose, glucomannan, pectin, hemicellulose, and polysaccharides like alginate and proteins like collagen and gelatine. Chemical polymerization techniques are typically used to create synthetic polymers such as hydrogel-formed polyethylene glycol (PEG), hydrogel-formed polyvinyl alcohol (PVA), hydrogel-formed polyacrylic acid (PAA), and hydrogel-formed polyacrylamide (PAM).2.

Depending on the synthesis or composition of polymers:

(I) The fundamental structural and functional unit of homopolymer hydrogels is made up of a single kind of monomer within the polymeric network. Depending on the kind of monomer and the polymerization process, their skeletons could be cross-linked.

(II) Copolymeric hydrogels are derived from a variety of monomeric units with at least one hydrophilic component. The polymeric network chains can be arranged in a random, block or alternating configuration. 17

(III) Multipolymer interpenetrating polymeric network (IPN) can be synthesized using two separate, cross-linked components of natural or synthetic polymers, confined in a network form. In the case of Semi-interpenetrating hydrogel, one polymer is cross-linked and the other polymeric component is non-cross-linked.18 One straight polymeric chain enters another crosslinked network, and they interact without any chemical bonding.19

Based on physical and chemical composition: (I) Non-crystalline (Amorphous) (II) Semicrystalline, a composite of amorphous and crystalline phases. (III) Crystalline.2

Based on cross-linked networks: Chemically cross-linked networks have permanent bonding involving covalent interaction while physical networks have transient junctions involving entanglements of polymeric chains involving hydrogen bonds, polar or ionic, hydrophobic type of physical interactions.2

(IV) Based on electrical charge: (I) Neutral (non-ionic), (II) Ionic (including anionic or cationic), (III) Amphoteric having both acidic and basic groups, (IV) Zwitterionic (polybetaines) possess both cationic and anionic functionality in each repeating unit.2

**1.4 Preparation of hydrogels**

Hydrogels can be produced employing natural, synthetic, and composites of both natural and synthetic polymeric materials. Hydrogels have been created by cross-linking polymer chains through chemical alteration, external cross-linking agents, exposure to high energy radiation, and polymerization grafting. In the hydrogel formation chemical cross-linking involves the formation of new covalent bonds between polymeric chains in the hydrogel, where as physical cross-linking involves physical interactions between polymer chains.53 Both chemical and physical methods have their own advantages and disadvantages related to them. Conventional and controlled radical polymerization techniques result in hydrogels with various morphologies, sizes and compositions including hollow core-shell particles.54, 55 The most widely used mechanism is free radical polymerization to prepare hydrogels.56, 57 Usually in hydrogel preparation, the gel reactants react with a crosslinker (s) to generate 3D-crosslinked networks in the presence of radical initiators like potassium persulfate or ammonium persulfate. Free radical polymerization occurs in three main steps: initiation, propagation, and termination. In the initiation step, free radicals (R●) are produced when an initiator dissociates and then reacts with other molecules (M) to produce the first radicals M●. In the propagation step, highly reactive free radicals rapidly react with molecules of monomer resulting in the formation of macroradicals. Usually, termination occurs by the combination or disproportionation reaction of free radicals. Various shapes of hydrogels like bulk, sphere, and films can be obtained by selecting the appropriate preparation process, raw material, and polymerization condition.47

Bulk Hydrogels: Usually, they are smoothly obtained by solution or homogeneous polymerization wherein all the reactants i.e., the monomer (or polymer), initiator, and cross-linker are soluble in the medium. The resultant hydrogel generally takes up the shape of the container in which it has been polymerized and yields a relatively homogeneous hydrogel.59 However, due to the slow diffusion of solute to the adsorption sites within hydrogel it will take a long time to reach equilibrium during water treatment.60 Usually, bulk hydrogel is cut into small sized pieces manually or by using a food blender to produce small size hydrogel beads for better adsorption efficiency. On the other hand, occasionally a cutting or grinding step could produce hydrogel particles with a fractured morphology and polydispersity.61, 62

Spherical Hydrogels:  Spherical hydrogel does not require grinding or cutting and thus avoids further morphology destruction and energy consumption. Hydrogel bead is an example of spherical hydrogel with a millimetre diameter. Usually, the synthesis of spherical hydrogels involves dropping the monomer or polymer suspension using a syringe into a solution. Thus, the size of the resultant hydrogel beads typically depends on the syringe's diameter. Chitosan, a natural polysaccharide which is biodegradable, nontoxic, odourless, biocompatible and Hydrogel beads are often prepared using biopolymer. When it comes into contact with potassium and sodium cations, it can get crosslinked.63-67

Hydrogel Films: Hydrogel composite films appear to hold a lot of potential for practical use. They are simple to make and show robust and repeatable self-healing behaviour in the aqueous medium. Numerous hydrogels have been used as an effective ion-exchange film to purify water. Recently, the direct synthesis of nanofiber hydrogel film has been achieved using electrospinning technology.38 Some extra components could be added to the hydrogel film either by grafting after polymerization or by combining additive with hydrogel precursor before polymerization to create a specific hydrogel composite with the required qualities.69 Hydrogel film is typically utilized as an active membrane in sensing applications, mainly to give a more hydrophilic surface that is less prone to contamination.65

**1.5 Characterization of Hydrogels**

Numerous characterization approaches have been utilized for knowing the hydrogel’s physical and chemical properties. The physical properties of polymeric hydrogels are determined by the volume fraction, effective molecular weight of the polymeric chain in between two crosslinking junctions and by the density of crosslinking.20 Hydrogels have many properties, such as absorption capacity, permeability, swelling behaviour, optical, surface, and mechanical properties. The nature of the polymer chains and the crosslinking present in the network structures play a significant function in the result of the properties of the hydrogel. All these properties are responsible for making hydrogel a promising material for a wide range of applications.21

Fourier Transform Infrared Spectroscopy FTIR analysis: This method provides reliable crosslinking data and gives a notion of the hydrogels' morphology.

Atomic Force Microscopy (AFM): This technique helps to examine hydrogels' surface morphology. It uses a multimode atomic force microscope.

Network Pore Size: Various techniques, for instance, mercury porosimetry, quasi-elastic laser light scattering, equilibrium swelling, electron microscopy, and rubber elasticity measurements experiments are employed to find out the network pore size of hydrogel. This is an important technique for hydrogel characterisation.

 X-ray Diffraction: X-ray diffraction analysis enables one to understand the crystalline and amorphous nature of hydrogel, whether the crystallinity is maintained or was distorted while synthesizing.

Swelling Behaviour: To study the potential use as a hydrogel, specific swelling data studies are employed and it has been successfully studied by numerous researchers.

Crosslinking and Mechanical Strength: The crosslinking density inside the network structure of hydrogel determines its mechanical strength. Generally, with increasing crosslinker concentration the mechanical strength of the hydrogel also increases.

Rheology: It depends on the kind of interactions (entanglement, association, and crosslinks) present in the polymeric network among polymer chains.

All these characterization methods provide important information about the desired crosslinking results, formation of hydrogel. Which can be useful further for various applications.3

**2 Applications of Hydrogel**

The salient features of hydrogels are biodegradable, hydrophilic in character, biocompatible, less toxic, highly flexible like tissues and easily modifiable. They have good transport properties and the capacity to adapt to changes in the environment, such as those in pH, temperature, or metabolite concentration. Owing to their extraordinary properties, hydrogels are said to have novel applications in a number of fields like drug delivery, wound dressing, agriculture, tissue engineering, water purification, hygiene applications, etc.3,7

Because hydrogels exhibit characteristics that are comparable to those of human soft tissue, they are widely used in biomedical fields, including drug delivery.22-25, gene vectors, tissue engineering26, 27, and biosensors28, 29. Hydrogels meet both material and biological requirements because they have unique characteristics like desired functionality, reversibility, and biocompatibility. They are frequently employed for cell-laden, tissue regeneration, drug delivery, and biosensors.

Soft Contact Lenses: It remains one of the most popular uses for hydrogels because of their biocompatibility and mechanical properties.

 By dissolving the lens's water, hydrogels can be adjusted to match the curvature of the entire eye, allowing atmospheric oxygen to reach the cornea.30 Polyhydroxyethylmethacrylate (PHEMA) was the first ever established synthetic hydrogel as a favourable and great candidate for the manufacture of contact lenses by Wichterle and Lim (1960).3

Tissue Regeneration and Tissue Engineering: The loss or chronic failure of any organ function due to some severe disease or accident necessitates the demand for tissue and organ transplantation. It has become more difficult because there are fewer donors available and because of societal, legal, and other norms.31 Tissue engineering has raised hopes for creating a perfect living replacement that mimics the ways in which living tissues perform in the human body.3 Scaffolds act as 3-D artificial templates in which the rebuilding of targeted tissue is cultured to grow. The extremely porosity of hydrogels enables the diffusion of cells during migration, transfer of nutrients and excludes unwanted products outside of cellular membranes.32 Hydrogels, both natural and synthetic, are utilized as scaffolds in numerous tissue engineering applications, such as the restoration of blood arteries, skin, heart valves, cartilage, and tendons.33 They have been used in a number of biomedical applications, including fillers for scar cosmetic repair7, bladder34, cartilage35, orthopaedic applications36,

Skin37 and bone38. Polysaccharides-based hydrogels that exhibit biocompatibility with tissues which increases their significance in tissue engineering and biomedical applications.

Wound healing: Injured skin is covered to avoid bleeding and to protect the wound from environmental infections. Wound dressings are non-toxic, antiseptic, permeable to oxygen, preserve wound moisture, cause minimal damage, eliminate excess exudates and thus fasten the healing process while directly interacting with the wound. A great advantage of gum-based hydrogels in wound dressings is that they can easily be applied or removed without interfering with the wound beds.39, 40 Compared to traditional bandages, pads, or gauzes, the mechanical characteristics of hydrogels increase their elasticity and flexibility to adapt to wounds and provide patients with immediate pain relief. They act as a coolant to localized wounds in case of burn and also reduce the pain and recover from resultant damage.41, 42 The non-adhesive nature and hydrophilic surface of hydrogels do not allow it to attach with cells therefore causes less pain and discomfort to the patient. Hydrogel transparency has a benefit over traditional bandages as it causes less discomfort when peeling it off. Various hydrogels for wound dressings are available, like amorphous gels, gel-impregnated gauzes, plasters or sheets. The development of hydrogel formulations to address different aspects of wound healing and management such as easy dressing, reduction in infection is attaining new heights.43, 3

Drug Delivery: Hydrogels' porous structure can act as a matrix for the loading or distribution of pharmaceuticals while also shielding them from harsh environments. Hydrogel targets specific sites like the colon as a drug delivery agent and releases drugs or other nutrients timely. In addition to this hydrogel interacts very less with the drug and other loaded solutes hence sustained and prolonged release occurs in a larger fraction comparative to conventional drug delivery systems.7 Because of their special ability to retain large volumes of water, hydrogels are valuable in drug delivery applications that regulate the release of solute over a predetermined length of time. This trait is known as hydrophilicity. Many biomaterials that function through two mechanisms have been investigated for this aim. (1) By adjusting the crosslinker dosage and keeping an eye on the proportion of hydrophilic to hydrophobic monomers, a controlled release of the medication can be accomplished. (2) Hydrogel releases a large fraction of active drug molecules (protein and peptides) because its interaction with the drug is very less. Drug delivery that is targeted and controlled would help with healing and lessen unwanted side effects. Drug release from hydrogel is expressed by a number of processes, including diffusion, chemical control, deswelling, and environmentally responsive release. 3

Agricultural Applications: One significant step in achieving sustainable development and growth in agriculture is water management. Superabsorbent polymeric hydrogels (SPH) derived from natural polysaccharides have gained significance in agriculture due to their remarkable capacity to hold and retain large amounts of water. In dry and semi-arid regions, SPH can be added to the soil to prolong longer moisture retention, enabling crops to tolerate arid weather. In dry and semi-arid soil, hydrogels act as "mini liquid tanks," releasing water into the soil along with the targeted amount of loaded nutrients. Modification in hydrogel properties as required gives fertile physical properties of soil.45 The usage of natural polysaccharides based SPH is flourishing owing to their biodegradability, durability, high water holding ability, avoid loss of nutrients, nontoxic, and their sustainability compared to synthetic polymer based hydrogels.46 Hydrogels have been used for the prevention of soil erosion over a decade by reduction in soil erosion, increasing water holding capacity, enhancing permeability of finely textured soils, enhance water infiltration among fine-textured agricultural soils. The water-soluble polyacrylamide (PAM) hydrogels form a thin film covering soil surface and are very efficient in preventing soil erosion. This film protects the soil surface from washing away during irrigation and retains the optimum water content within the soil system, so that irrigation water can permeate easily.3

Hydrogel as an adsorbent:  As we all know emerging contaminantsincluding pharmaceuticals, pesticides, industrial chemicals, metal ions, surfactants, and personal care products have elevated worldwide concern for their noteworthy hazard to marine ecosystems and human health. Many of them have no regulatory standards on the effects of chronic exposure due to the lack of information.70 These contaminants are stable under a variety of circumstances such as aerobic digestion, heat and light thus they have the potential to build up and harm ecosystems. Therefore, the adsorption method has been greatly adopted to treat emerging contaminates as it is really efficient and affordable.71

The use of hydrogel as an adsorbent in pollution management applications is becoming more and more popular. Both the hydrogel adsorbents and the type of adsorbate have a significant impact on the adsorption process. The Freundlich and Langmuir models provide a good interpretation for the adsorption data of emerging contaminants on hydrogels; the kinetic model is often pseudo-second-order. Because of the several interactions that occur between the adsorbate and the adsorbent, such as hydrophobic interaction, hydrogen bonding, ionic or electrostatic interaction, and π–π interaction, hydrogel adsorbents have a great affinity for pollutants. This fluctuates depending on several factors, including pH, the ionic strength of the solution, the chemical makeup of the adsorbent and adsorbate, and more.

The surface of the adsorbent will have an ionic charge when the pH of the solution differs from the hydrogel adsorbents' isoelectric point. Ionic adsorbates undergo simultaneous protonation and deprotonation at varying pH levels, leading to electrostatic interactions between them. Hence, pH is an important factor responsible for the adsorption mechanism.47, 3 Many solid and liquid phase removal trials remain tracked for the elimination of pollutants from liquid such as coagulation, biochemical precipitation, adsorption, photodegradation, ion exchange, flocculation, electrochemical treatment, and membrane percolation.48-51 Of all these many methods, adsorption is seen to be superior due to its high efficiency, low effort, and ease of use. Hydrogels are thought to be special for adsorption-based water refining because of their high absorption capacity, low crystallinity, abundance of functional groups, and porous structure. Due to the several significant functional groups that polysaccharides (Gum) include in their structure, hydrogels based on them and graft copolymers have been thoroughly investigated as adsorbents for the removal of contaminants from aqueous environments, including heavy metal ions and organic dyes. The principle advantage of using natural gum-based hydrogels as adsorbents is their biocompatibility. Their structure can be easily tailored according to the nature of the pollutant.52 Although hydrogels are found to be superior candidates for the elimination of several aqueous contaminants, including heavy metals, dyes and other emerging contaminates, selective adsorption of contaminates is hardly explored. Therefore, research attempts are required to prepare hydrogels with desired properties, sensitivity, and selectivity for a specific contaminant. Many hydrogels have been developed with desirable strength and adsorption capacity, but their chemical and biological stability is always ignored which needs to be considered for the sustainability and economic viability of wastewater treatment.47

Hydrogel as a sensor of heavy metal ions in environmental and biological samples: Nonbiodegradable heavy metal ions' widespread existence in water is potentially threatening to the ecosystem and living organisms. Hydrogels have been functionalized with many biomolecules, including DNA, to form stimuli-responsive sensors and materials72,73 However, for sensing applications, most of them rely on hydrogel phase transition or volume change. Because of their high sensor loading capacity, excellent biocompatibility, and extremely low optical background, hydrogels are perfect for optical sensor immobilization. Moreover, hydrogel backbone properties such as charge and hydrophobicity can be modified by mixing with different monomers, allowing further control of sensor performance.73

As a transducer material, the stimuli-responsive hydrogel can be utilized to convert a recognition unit's reaction into a physical signal that can be detected by, e.g., quantifying the change in optical length with an optical fibre, observing the resulting change in swelling pressure under isochoric conditions or by measuring the diffracted wavelength of a polymerized crystalline colloidal array.

Host-guest interactions in sensing applications have been proven to be a powerful tool. Usually, macrocyclic polyethers, i.e., crown ethers have been proved to be promising candidates in combination with hydrogel facilitated by the generation of highly selective and reversible host-guest complexes with specific alkali and heavy metal cations. Notably, the colorimetric sensing approach has garnered a lot of interest due to its direct visual perception, affordability, speed, and ease of use. Most colorimetric sensors are dispersed in sample solution for full contact with the target chemical to increase sensitivity; however, an uneven and unstable dispersion of the sensors might occasionally result in unsteady detection findings. Chemo sensors can be constructed on solid substrates to circumvent this issue. The choice of a solid substrate is crucial since it has a significant impact on the sensor's sensitivity. In sensor studies, a colorimetric chemo sensor can be formed by the design of molecules that change their colour in sample solution due to an alteration in their molecular structure in the presence of target ions.74 In aqueous conditions, the designed sensor molecule's solubility is crucial. The fact that the sensor cannot dissolve directly in water, although pollutant species, such as anion or cation, are soluble in water, is one of the most significant problems in this research. Additionally, the soluble sensor is meant to be used just once. To deal with such issues, researchers have opted to polymerize molecules that have sensing properties. According to this perspective, the sensor must be easily extracted from the sample solution and insoluble to be utilized again. Hydrogel is the best illustration of a solid support or polymer that is readily extracted from the sample solution through filtration. They are important for various sensing applications as they can be synthesized with good yield, have swelling in water, are reusable and stimuli responsive.74,75

**3 Conclusion**

The presented review demonstrates the literature concerning classification of hydrogels, their properties, and applications. Hydrogels can be integrated into systems and changed into different configurations due to their biocompatibility, sensitivity to external stimuli, and physical and chemical structure. Hydrogel-based chemical and biosensors have advanced significantly in the last several years across a wide range of application areas. Hydrogels have proven beneficial in a variety of disciplines, including energy, environmental remediation, humidity sensing, medicine, soft robotics, and health monitoring, due to their versatile composition, innate properties, and ability to adjust numerous physicochemical parameters.

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