Synthesis and Swelling Behavior of Sodium Alginate / Poly (vinyl alcohol) Hydrogels

Sodyum Aljinat/ Polivinil Alkol Hidrojellerinin Sentezi ve Şişme Davranışları

Lachakkal Rudrappa SHIVAKUMARA¹ and Thippaiah DEMAPPA^{1*}

¹Department of Post Graduate Studies and Research in Polymer Science, University of Mysore, Sir.M.Visvesvaraya Post-Graduate Center, Tubinakere, Mandya, Karnataka, India- 571402.

*Corresponding author: Dr. T. Demappa. Email: tdemappa2003@yahoo.co.in

ABSTRACT

Objectives: Hydrogels are macromolecular networks able to absorb and release water/biological fluids in a reverse phase manner, in response to specific environmental stimuli. Such stimuli –sensitive behavior makes hydrogels interesting for the design of smart devices applicable to a variety of technological fields. They are able to absorb and retain 10-20% and upto 1000 times of water or biological fluids than their dry weight. The aim of this study is to extend the work in drug delivery in stomach at pH 2-2.2.

Methods: The authors are synthesized sodium alginate/poly (vinyl alcohol) hydrogels. These hydrogels are characterized by FTIR, SEM, swelling properties of hydrogels in different pH, in salts, different temperature, different acids and bases.

Results: The authors studied and reported that the swelling effects or variations such as effect of salts, acids, bases, temperature and pH. The effect of cross-linking agent glutaraldehyde shows that the 8 mL glutaraldehyde has the highest swelling rate as compare to that of 10 mL and 12 mL.

Conclusions: In this work authors studied the swelling degree in different acids and bases. We are concluded that the degree of swelling is decreased with increase in the concentration of the glutaraldehyde and also depending on the concentration of the acids. The swelling degree of PVA/SA hydrogels are

gradually increased with increase in concentration of acids (low p H). The swelling of hydrogels decreases with increase of pH (> 7) or at high alkaline. In the effect of salt solutions the swelling behavior is found to be in the order: $K^+ > Na^+ > Ca^{2+} > Mg^{2+}$.

Key words: swelling, P^H , cross linking agent, buffer solution, Sodium alginate, PVA

ÖZET

Amaçlar: Hidrojeller, belirli çevresel uyaranlara yanıt olarak su / biyolojik sıvıları ters fazda emebilen ve serbest bırakabilen makromoleküler ağlardır. Bu tür uyaranlara duyarlı davranış, çeşitli teknolojik alanlara uygulanabilen akıllı cihazların tasarımı için hidrojelleri ilginç kılar. Kuru ağırlığından% 10-20 ve 1000 veya daha fazla su veya biyolojik sıvıları emebilir ve tutabilirler. Bu çalışmanın amacı, ilacın 2-2,2.

Yöntemler: Yazarlar sentezlenmiş sodyum aljinat / poli (vinil alkol) hidrojelleridir. Bu hidrojeller FTIR, SEM, farklı pH'larda hidrojellerin şişme özellikleri, tuzlar, farklı sıcaklık, farklı asitler ve bazlar ile karakterize edilir.

Bulgular: Yazarlar, tuzların, asitlerin, bazların, sıcaklığın ve pH'ın etkisi gibi şişme etkilerini veya varyasyonlarını incelediler ve rapor ettiler. Çapraz bağlama maddesi glutaraldehitin etkisi, 8 mL glutaraldehidin, 10 mL ve 12 mL'ninkiyile karşılaştırıldığında en yüksek şişme oranına sahip olduğunu göstermektedir.

Sonuçlar: Bu çalışmada yazarlar farklı asit ve bazlarda şişme derecesini incelemişlerdir. Glutaraldehit konsantrasyonundaki artışla ve aynı zamanda asit konsantrasyonuna bağlı olarak şişlik derecesinin azaldığı sonucuna varılmıştır. PVA / SA hidrojellerinin şişme derecesi, asit konsantrasyonundaki artışla (düşük pH) kademeli olarak artmaktadır. Hidrojellerin şişmesi pH'ın (> 7) artmasıyla veya yüksek alkali ile azalır. Tuz çözeltilerinin etkisinde şişme davranışının sırasıyla: $K^+ > Na^+ > Ca^{2+} > Mg^{2+}$ olduğu bulunmuştur.

Anahtar kelimeler: şişme, PH, çapraz bağlama maddesi, tampon çözelti, sodyum aljinat, PVA

INTRODUCTION

Hydrogels have been used in various chemical and biomedical applications in ophthalmology as contact lenses, surgical sutures, as well as in numerous other areas like agricultural applications.

Sodium alginate is an anionic co-polymer composed of 1, 4-linked β -D mannuronic (M- blocks) and α - L- guluronic acid (G- blocks), interspersed with the regions of alternating structure. Gel formation and three –dimensional network structures were occurs when divalent ions (Ca^{2+} , Ba^{2+} , Fe^{2+} , Si^{2+} etc.) or trivalent ions (Al^{3+} , etc) cross-linking with G-blocks in the polymer chain. Such binding zones between G-blocks are often referred to as “egg boxes”. These cross- linked ions that stabilize alginate chains forming a gel structure, with more freely movable chains that bind and entrap large quantities of water or biological fluids. The gel formation (gelification) processes is characterized by the eviction of water.¹ The softer, more fragile and lower porosity gels are made of M-rich alginate groups. This is due to the lower binding strength between the polymer chains and to the higher flexibilities of the molecules. The gel formation process is highly dependent on diffusion of gel formation ions into polymer network. Visco- elasticity and transmittance of alginate structures are highly affected by the M/G ratio. Alginic acid and its salts of sodium and calcium are being used in the medical, pharmaceutical, cosmetic and food industry because of its non-toxic and biocompatibility.² The main advantages of hydrogels are that they possess a degree of flexibility very similar to that of natural tissues, due to their significant water content. The stimuli-sensitive behavior makes hydrogels interesting for the design of smart devices applicable to a variety of technological fields.

Hydrogels are in general, materials composed of three - dimensional hydrophilic polymer networks and water that fills the free spaces inside this network. Hydrogels are able to absorb and retain 10-20% upto 1000 times of water or biological fluids than their dry weight. Hydrogels respond reversibly to slight changes of the properties of surrounding media; hence they are called as

“intelligent materials.” This ability is due to the hydrogels found many applications in industry and pharmacy, for example as controlled drug delivery systems.³ The swelling and dehydration behavior is one of the most importances among other significant properties of hydrogels.⁴ Several applications of hydrogels such as drug delivery systems (drug slow release), a wound dressing, dental applications, transdermal implants, injectable polymers, contact lenses, super absorbents, and environmental sensitive hydrogels.⁵⁻⁹ Hydrogels are interacting with aqueous solutions and swelled to certain equilibrium and retain a significant proportion of water within their structure.¹⁰ Sodium alginate (SA) is a hydrophilic polysaccharide, natural polymer, composed of mannuronic acid unit. This compound has been used for a long time in various industries, such as agriculture, foods, medicines, textiles, cosmetics, printing, etc. SA is also used as a thickener, stabilizers, emulsifier or micro encapsulation. SA is also used in slow release drug delivery system, and fertilizer.¹¹ SA has a molecular structure similar with collagen, therefore it can make skin smooth, elastic and can recover wound, and it can be used as a natural alternative product for health care and cosmetics.¹²⁻¹⁴ SA is biological origin, has good characteristics, such as biocompatible, or biodegradable and gel forming ability. Poly (vinyl alcohol) (PVA), is hydrophilic polymer, has a great interest to be used as a biomaterial because it has good biocompatible properties. It has chemical stability, high durability and high degree of swelling in water or biological fluids. PVA is non-toxic to viable cells, non- carcinogenic, has high biocompatibility, and has consistency similar to soft tissue, film forming with high mechanical strength and long –term temperature stability. Its 3-dimensional network enable to facilitate diffusional exchange of nutrients and waste products with surrounding environment, it is used in various biomedical, pharmaceutical, biotechnology and other industrial fields.¹⁵⁻²⁰ However, even though PVA is a biomaterial, but brittle in nature; therefore it needs to be blended with other polymer or by copolymerization, eg. with sodium alginate, to obtain a better property that can be used to encapsulate or entrap or to immobilize an enzyme or drug in micron or submicron (nano) size, to keep the constancy of its activity or to prevent activity decreases drastically. Hence make

it work more effective and efficient compared to when it is in free condition.²¹⁻²³ The cross linked alginate hydrogels have been used as a controlled release medium for drugs,²⁴⁻²⁸ Pesticides,²⁹ superabsorbent filament fibers³⁰ and flocculants.³¹

In the presence of an aqueous solution, the polymer chains absorb water and the association, dissociation and binding of various ions to polymer chains causes the hydrogel to swell. The shrinking and swelling properties of hydrogels are currently being exploited in a number of applications including control of micro fluidic flow,³² muscle-like actuators,³³⁻³⁴ filtration/separation³⁵ and drug delivery.³⁶⁻³⁷ The structure and properties of hydrogels are similar to many biological tissues such as cartilage and the corneal stroma in the eye.³⁸⁻³⁹ Hydrogels are accomplished of undergoing large reversible deformation in response to changes in several environmental factors.⁴⁰ For example, hydrogel size is sensitive to pH, temperature, concentration of salts, and electric fields.

EXPERIMENT

MATERIALS AND METHODS

Materials

Sodium alginate, Poly (vinyl alcohol) molecular weight 1,25,000 and glutaraldehyde (25 %) was purchased from S.d. fine chem limited, Mumbai- 30, India. Hydrochloric acid, perchloric acid, sodium hydroxide and acetic acid were purchased from Reachem Laboratory chemicals private ltd, Chennai-600058, India. Calcium chloride, Magnesium chloride, sodium chloride, potassium chloride were purchased from E-Merk limited Mumbai -400018, India, and double distilled water were used throughout the experiment.

Methods

Preparation of Sodium alginate/ Poly (vinyl alcohol) hydrogels

7 g of sodium alginate dissolved in 100 mL of water by constant stirring. 9 g of PVA dissolved in the same solution by stirring about 3 hr at 80°C to 85°C and 12 mL of glutaraldehyde (25 %) is added to the same solution, this solution is kept under 80°C for 3 hr. After 3 hr the obtained hydrogel is washed with distilled water and ethanol, to remove the excess monomer, cross –linking agent. After 2-3 times of washing the hydrogel are dried under 40°C in an oven.

Likewise synthesized the different SA/PVA hydrogels by varying volume of glutaraldehyde (10 and 8 ml) are used for swelling studies.

CHARACTERISATION

Fourier-transform infrared spectroscopy

The FTIR spectra of the sodium alginate, PVA and the cross-linked hydrogel samples have been recorded in the range of 4000 to 500 cm^{-1} to provide the proof of hydrogels (Figure.1).

Surface morphology of hydrogels

The surface morphology of sodium alginate, PVA and glutaraldehyde cross-linking hydrogels were investigated by using scanning electron microscopy (SEM zeiss, LS15) (Figure.2).

Swelling Behavior of hydrogels

In the swelling behavior of hydrogels, authors also studied swelling ratio of the hydrogel sample was measured at different temperatures in different solvents by using gravimetric method. The pre weighed dry hydrogel samples were immersed in excessive different solutions and kept undisturbed by 24 hr, 48 hr and 72 hr at different temperatures like room temperature, 30°C, 37°C and 40°C until the constant values is obtained. Degree of swelling rate can be calculated by the equation:

$$\% \text{ DS} = (W_2 - W_1) / W_1 \times 100 \dots \dots \dots (1)$$

% DS is the degree of swelling expressed in percentage, W_1 and W_2 are the masses of sample before and after swelling respectively (Figure.3 to 5) (Table1).

Swelling at various pHs

We are prepared the solution to acidic, basic and neutral pHs by diluting of phosphate buffers (pH 3.2, pH 7, pH 2 and pH 10) solutions at room temperature, 30°C, 37°C and 40°C temperatures. The pH values are checked by using pH meter. The dried hydrogel samples were used for the swelling measurement according to eqn (1) (Figure.6).

Swelling in salt solutions

Swelling capacity of hydrogels was determined in the different salt solutions (KCl, NaCl, CaCl₂, and MgCl₂) and also with various concentrations like 0.4, 0.6, 0.8, 1, and 1.2 N according to the above method (Figure.7).

RESULTS

The authors reported the swelling effects of hydrogels on salts, acids, base, temperature and pH. 8 mL of glutaraldehyde shows the highest swelling rate as compared to that of 10 mL and 12 mL. Figure.1 shows that FTIR spectra it deals with the molecular interaction between sodium alginate and PVA. Surface morphology of hydrogels was studied by using scanning electron microscopy (SEM zeiss, LS15) in figure 2, figure 3 to 5 and table 1 represent swelling behaviour of hydrogels at different temperature i.e; 30°C, 37°C, 40°C and room temperature with different salts. Figure 5 represent behaviour of hydrogels at different pH.

DISCUSSION

Fourier transformed infrared spectral analysis

Figure 1 represents the FTIR spectra, characterized the molecular interaction of sodium alginate and Poly (vinyl alcohol) hydrogels. The FTIR spectra of SA shows the characteristic absorption peak at 3270.47 cm⁻¹ is for hydroxyl groups (-OH). The asymmetric and symmetric stretching vibration of carboxylic group (COO-) is found to be at 1597.36 and 1412 cm⁻¹ respectively.⁴¹⁻⁴² The peak at 2925.38 cm⁻¹ represents for C-H alkyl stretching bond.⁴³ An absorption peak around 2919.05 cm⁻¹ shows the characteristic spectra of PVA. This peak is arising from the C-H stretching at 1570 to 1420 cm⁻¹ assigned for CH₂ (vinyl group), the sharp absorption peak at 1150-1050 cm⁻¹ is used for indication of PVA.⁴⁴ In addition, it was found that the peak at 1549-1453 cm⁻¹ is stretching band for CH₂ group. This band is also found in pure PVA and cross linked SA/PVA hydrogels. The decreasing in wave number of carbonyl peak from 1652.14 to 1635.24 cm⁻¹ is for the cross-linking of SA/PVA hydrogels. Cross – linking of glutaraldehyde is takes place at 2863 to 2750 cm⁻¹.

Scanning electron microscopy (SEM) images

Scanning electron microscopy (SEM) describes the surface morphology of SA/PVA hydrogels in figure 2. According to the SEM images (Figure 2a), the pure sodium alginate shows a very smooth surface which is about devoid of any surface feature⁴⁴. In figure 2b SEM images of pure PVA shows a very smooth, uniform and non-porous surface structure, which may be attributed to the crystallization of PVA.⁴⁵ However, the addition of SA into PVA hydrogel in different portions provides very tiny pores at surface, these pores decreases with increase of glutaraldehyde concentration (Fig 2c to 2e).

Effect of pH on Swelling of Hydrogels

The sensitivity of the hydrogels was measured from pH 2 - pH 10. No additional ions (through buffer solutions) were added to medium for setting pH because absorbency of an absorbent is strongly affected by ionic strength. Therefore stock HCl (pH 1.0) and NaOH (pH 10.0) solutions were diluted with distilled water to reach preferred acidic or basic pHs respectively. In table 2, figure 6, the swelling capacity of hydrogel at pH 2 can be accredited high repulsion of anion-anion carboxylic (COO⁻) groups. At basic conditions (pH $\geq$ 7), the most of the carboxylate groups are protonated and the low swelling values of hydrogels can be attributed the presence of non-ionic hydrophilic -OH, and - COOH groups in the PVA and alginate back bones respectively. The swelling capacity is decreased with further increase of pH (pH10 or pH > 7). Again the swelling loss is due to the counter ions, i.e Na⁺, that shield the charge of the carboxylate anions and prevent efficient anion- anion repulsion. As a result, a remarkable decrease in equilibrium swelling is observed.

pH Dependence of Swelling rate of water

The swelling rate was decreased with increasing from pH 2-10; it was evident from the data given. The negatively charged ionic back bone of PVA and SA are more expanded because of protonation of carboxylic groups is negatively charged. This expanded form makes for easy diffusion of water molecules into the hydrogel network. On the converse, the hydroxyl groups are mostly in the

protonated form, and show less polar character on $\text{pH} \geq 7$. And therefore, results in the polymer with lower affinity to water. Thus, hydrogels are less expanded on $\text{pH} \geq 7$. The increased in swelling ratio are responsible for theory of electrostatic repulsion between carboxylic ions in the polymer chains and ionic present in the pH solution and the ionic osmotic pressure generated from mobile counter ions to charged ion in the network. Thus, dilute the charge density of the hydrogel, because of PVA is not ionic in character (Table 2). Figure.6 show the Swelling studies of phosphate buffered pH solutions, the glutaraldehyde cross-linked PVA/SA samples show gradual increase (in swelling degree in the pH 2 and 3.2) followed by decreased in swelling degree in the pH 7 and 10 at different temperatures (25°C , 30°C , 37°C). In this studies 8% glutaraldehyde cross linked hydrogel having high swelling degree as comparing to the 10% and 12% glutaraldehyde cross- linked hydrogels.

Effect of Temperature on Swelling of Hydrogels

It is obvious from the Figure 6 that the temperature leading to hydrogel with the highest absorbency is found to be around 37°C . The swelling capacity of hydrogel was decreased with increasing temperature above the 37°C . The increase in swelling rate values is dependent on kinetic energy of the polysaccharide chains which led to lower soluble content of the hydrogel as well as increasing the concentration of glutaraldehyde diffusion rate of SA and PVA back bone. The higher reaction temperature proves the results from higher reactant movement and effective collision. The temperature about 37°C , the possible "thermal cross linking" reaction to polysaccharide backbones may act as a major role to lead low-swelling hydrogels. In addition, the swelling loss may be related the increasing of cross linked bond formation of completion of the ester and ether formations by further reaction to the possible mono-ester species with another polysaccharides chain (scheme1).

Effect of Salt solution to Hydrogels

Hydrogels are considered as poly electrolytes suggest that their porosity should decrease as ionic strength increases. SA/PVA hydrogels with various chloride salt solutions are appreciably reduced in swelling compared to that of measured

in de-ionized water. This results from a charge screening effect of the additional cations causing an anion-anion electrostatic repulsion, which leads to decreased osmotic pressure difference between the polymer network and the external solution. As a given ionic strength, Mg^{2+} and Ca^{2+} contribute more charge than mono valent cations like Na^{+} and K^{+} and induce a bigger drop in intermolecular repulsion and increased interaction between molecules, which in turn, cause to a large extent of the hydrogel collapse. In addition, Mg^{2+} and Ca^{2+} can chelate carboxylic group, leading to a compact network and causing further shrinking from the hydrogel, on the other hand, we also find that the smaller the radius of atom of some valent mono atomic cation, the more the water absorption capacity if different cations was $K^{+} > Na^{+} > Ca^{2+} > Mg^{2+}$.⁴⁶ (Table 3) & (Figure 7)

CONCLUSION

In this work authors studied the swelling degree in different acids and bases. By this study we are concluded that the degree of swelling is decreased with increase in the concentration of the glutaraldehyde and also depending on the concentration of the acids. Here swelling degree of PVA/SA hydrogels has gradually increased with increase in the concentration of acids. The aldehyde groups are covalently bonded with the COO^{-} and OH^{-} groups of PVA/SA and consequently the swelling degree is significantly reduced. In different concentration of different salts the swelling rates are found to be decreased with increasing concentration of salts. The swelling rate is in the order of $K^{+} > Na^{+} > Ca^{2+} > Mg^{2+}$.

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Further extension of the work

This swelling study is to be extended for biomedical and agricultural applications such as drug delivery and controlled release fertilizers.⁴⁶

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Table 1. Data's of Swelling rate of 8mL glutaraldehyde, 10mL glutaraldehyde and 12mL glutaraldehyde crosslinked hydrogels at 37°C.

Con.c	8 ml glutaraldehyde			10 ml glutaraldehyde			12ml gltaraldehyde		
	24hr	48hr	72hr	24hr	48hr	72hr	24hr	48hr	72hr
1N CH ₃ COOH	54.46	107.56	109.85	45.78	63.39	66.32	14.02	32.15	33.85
2N CH ₃ COOH	252.3	288.74	348.05	51.23	69.34	70.15	14.71	32.79	34.21
4N CH ₃ COOH	260.74	297.36	348.05	60.19	72.43	74.12	14.97	33.56	34.69
6N CH ₃ COOH	266.08	300.42	303.35	63.48	74.12	77.06	15.09	33.81	34.8
1N HCl	70.16	229.4	240.17	61.19	67.93	71.72	12.43	23.79	25.77
2N HCl	495.5	662.89	771.3	88.49	112.83	115.98	19.19	28.07	32.08
4N HCl	673.93	918.08	938.37	126.25	133.29	137.58	28.18	35.93	40.34
6N HCl	679.83	937.44	973.77	129.18	133.7	139.0	55.44	69.88	71.86
1N HClO ₄	124.47	169.92	173.65	132.12	133.5	137.56	11.9	10.69	11.79
2N HClO ₄	179.23	209.78	239.01	166.09	207.7	209.05	17.74	22.18	27.19
4N HClO ₄	890.19	957.01	976.08	260.01	268.59	277.18	19.21	23.29	31.07
6N HClO ₄	892.27	968.14	973.0	263.72	268.72	278.29	23.41	32.17	34.02
1N NaOH	47.26	47.02	46.24	32.87	29.4	27.64	34.97	22.83	19.31
2N NaOH	50.75	50.13	49.08	130.31	105.43	101.92	35.21	29.01	29.09
4N NaOH	53.24	53.02	52.86	134.04	107.47	104.36	39.93	38.14	36.69
6N NaOH	55.23	54.71	53.01	135.6	104.25	102.1	41.74	37.25	36.07

Table 2. Data's of Swelling studies of hydrogels in different pH solution at 37°C.

	8ml glutaraldehyde			10ml glutaraldehyde			12ml glutaraldehyde		
	24hr	48hr	72hr	24hr	48hr	72hr	24hr	48hr	72hr
pH 2	120.2	121.56	122.01	75.65	76.92	77.15	75.19	76.08	77.33
pH 3.2	109.2	113.2	116.07	72.01	74.1	76.12	73.24	74.19	75.28
pH 7	70.37	77.78	83.34	67.92	68.12	69.12	66.27	67.19	68.92
pH 10	69.24	71.34	73.45	65.26	66.74	67.05	65.48	66.47	67.19

Table 3. Data's of swelling studies of hydrogels in different salt solutions.

Con.c	12 hr				24hr			
	KCl	NaCl	CaCl ₂	MgCl ₂	KCl	NaCl	CaCl ₂	MgCl ₂
0.4	370.42	340.16	310.72	288.02	377.72	345.45	314.7	293.6
0.6	230.17	200.36	190.64	165	235.46	203.8	193.4	167.9
0.8	200.23	185.34	140.39	117.79	205.0	188.1	142.9	119.2
1.0	120.14	80.08	70.06	67.58	124.19	82.5	72.01	69.1
1.2	75.09	60.63	40.46	35.96	78.20	62.04	42.02	37.4

Figure 1. FTIR spectra of pure SA, PVA, 12% glutaraldehyde cross linked, 10% glutaraldehyde cross- linked and 8% glutaraldehyde cross-linked hydrogels.

Figure 2. SEM images of (a) Pure sodium alginate, (b) Pure PVA (c) 8 ml glutaraldehyde cross linked SA/PVA hydrogel (d) 10 ml glutaraldehyde cross linked SA/PVA hydrogels (e) 12 ml glutaraldehyde cross linked SA/PVA hydrogels.

Figure 3. Swelling rate of 8 ml glutaraldehyde crosslinking hydrogels.

Figure 4. Swelling rate of 10 ml glutaraldehyde crosslinked hydrogels.

Figure 5. Swelling rate of 12 ml glutaraldehyde crosslinked hydrogels.

Figure 6. Swelling studies of hydrogels at different pH of phosphate buffer solutions.

Figure 7. Swelling studies of different concentrated salt solutions at different time.

Glutaraldehyde

(a) Glutaraldehyde in acid medium

SA

SA

(b) Crosslinked Sodium Alginate(SA)

(c) Crosslinked Poly Vinylalcohol (PVA)

Scheme.1

Scheme.1