Hemocompatibility and swelling studies of poly(2-hydroxyethyl methacrylate-co-itaconic acid-co-poly(ethylene glycol) dimethacrylate) hydrogels

Sava N. Dobić, Jovana S. Jovašević, Marija D. Vojisavljević, Simonida Lj. Tomić
University of Belgrade, Faculty of Technology and Metallurgy, Belgrade, Serbia

Abstract
In this study a novel series of hydrogels, based on 2-hydroxyethyl methacrylate (HEMA), itaconic acid (IA) and poly(ethylene glycol) dimethacrylates (PEGDMA) (of varying molecular weight and concentration) were prepared by free radical cross-linking copolymerization. Preliminary hemocompatibility characterization of hydrogels obtained by hemolytic activity assay indicated good compatibility with blood. Preliminary biocompatibility characterization of P(HEMA/IA/PEGDMA) hydrogels, done by the cytotoxicity assays using the HeLa cell line revealed that the cell viability of all samples was in the range of 97−100%, with no significant decrease in cell viability with the change of PEGDMA molecular weight and concentration. Swelling studies were conducted for all P(HEMA/IA/PEGDMA) samples in a physiological pH and temperature range and network parameters were determined. Swelling studies showed pH sensitive behaviour, typical for anionic hydrogels, and temperature dependent swelling. The effects of concentration of PEGDMA component on hydrogel swelling properties depend on the PEGDMA molecular weight. The samples with 550PEGDMA show different swelling capacities when 550PEGDMA content is changed, whereas for P(HEMA/IA/875PEGDMA) samples there was practically no difference in equilibrium degree of swelling, %w, with varying 875PEGDMA content, which trend is the same as in the case of %w versus pH dependences. It was concluded that P(HEMA/IA/PEGDMA) hydrogels show good potential to be used as biomedical materials.

Keywords: hydrogel; 2-hydroxyethyl methacrylate; itaconic acid; poly(ethylene glycol) dimethacrylate; hemocompatibility; pH-sensitive and temperature dependent swelling; network parameters.

The development of new polymeric biomaterials for medical and pharmaceutical purposes is of great interest to life-care science and engineering. Hydrogels are polymeric biomaterials which do not dissolve but swell considerably in contact with aqueous and physiological media. They provide a swollen three-dimensional matrix with a predetermined amount of water or physiological fluids, resembling to some degree the environment of the native tissue [1−6]. Special types of stimuli-responsive hydrogels have been investigated for the development of “smart” materials in various fields. The term “stimuli-responsive” implies that marked changes of hydrogel volume can be induced by an external stimulus such as changes of the pH-value, ionic strength, temperature or pressure, light, or electrical and magnetic fields [7]. For example, thermosensitive systems in aqueous media are generally aimed at changing the hydrophilic character of functional groups into a hydrophobic one, or vice versa [1,7], with the change of temperature, whilst pH sensitive systems have basic or acid groups which react to the change of pH in aqueous media [1,2]. Smart hydrogels are of special interest in controlled drug release applications because of their soft tissue biocompatibility, simple loading and dispersing of the drugs in the matrix, and a high degree of control achieved by the selection of the physical and chemical properties of the polymer network and the response by volume variation to some external stimuli [5−7].

The homo- and copolymer hydrogels of 2-hydroxyethyl methacrylate (HEMA) have found extensive applications in the field of biomedicine because of their good chemical stability, high biocompatibility and physicochemical properties similar to those of living tissues [8,9]. Numerous studies have been conducted to modify PHEMA with the aim of improving swelling, mechanical properties and of eliciting better physiological responses to design stimuli-responsive hydrogels [10−17].

Due to the fact that IA is obtained from the non-petrochemical resources it is of a great interest for polymeric biomaterials. IA is obtained by fermentation from renewable resources such as carbohydrate mater-
rials containing sucrose and glucose (molasses and hydrolyzed starch) [18–20].

Different properties of poly(2-hydroxyethyl methacrylate/itaconic acid) hydrogels were investigated in the last decade by few authors. Caykara et al. determined the solubility parameters of pure PHEMA and P(HEMA/IA) hydrogels in 20 solvents, with various solubility parameters, in swelling experiments [21]. Tomić et al. investigated the swelling behavior and network parameters of hydrogels based on HEMA/IA and different types of poly(alkylene glycol) (meth)acrylates [22]. The same authors investigated swelling and thermodynamic properties of PHEMA and copolymeric P(HEMA/IA) hydrogels with varying IA content in a wide pH and temperature range. The copolymers showed interesting pH and temperature sensitivity [23]. These authors also investigated the drug release profiles from P(HEMA/IA) hydrogels in order to investigate transport phenomena [24]. Metal ion adsorption on P(HEMA/IA) hydrogels was investigated for uranyl ions by Inam et al. and Ozsurek et al. [25,26]. Hamdy et al. studied the potential use of P(HEMA/IA) hydrogels in immobilization of Citrullus vulgaris urease. Immobilized urease maintained a higher relative activity than free urease at both lower and higher pH levels [27]. Cytotoxicity test and hemolytic activity were investigated by Tomić et al. [28] for poly(alkylene glycol) (meth)acrylates/HEMA/IA) hydrogels. The in vitro study of biocompatibility showed no evidence of cell toxicity nor any considerable hemolytic activity. Also, all hydrogels showed satisfactory bioadhesive properties.

Poly(ethylene glycol) (PEG) is one of the most widely used synthetic materials for biomedical applications. Its biocompatibility, flexibility, and “stealth” properties make it ideal for use in drug delivery applications. Over the past few decades, poly(ethylene glycol) (PEG) based hydrogels have been extensively used as matrices in controlled drug delivery systems, as well as cell delivery vehicles for promoting tissue regeneration [29–32]. The versatility of the PEG macromer chemistry [33], together with its excellent biocompatibility, has spurred the development of numerous hydrogel systems for biomedical applications. Many of these studies have produced encouraging pre-clinical and clinical results.

Poly(ethylene glycol) dimethacrylate (PEGDMA), obtained by substituting PEG terminal hydroxyl groups with methacrylates, are versatile building-blocks for the preparation of “smart” biomaterials. Hydrogels containing PEGDMA crosslinker are highly tunable [34–38]. The mechanical properties of these hydrogels can be controlled by varying the molecular weight or concentration of PEGDMA. The mesh size and swelling ratio can be controlled similarly, and the mechanical and biochemical properties can be varied independent of one another.

The benefits of controlled drug delivery are essential for pharmaceutical applications as drug administration may be improved by using a delivery system designed for continuously maintaining the plasma levels of the active molecule in a therapeutically desirable range. Furthermore, drugs can be released in a precise and prolonged manner without repeated administration, thus improving patient comfort. Other benefits of controlled drug delivery are localized delivery to a particular site in the body and preservation of active agents that have short lifetimes in the body.

In this study, novel copolymeric hydrogels based on 2-hydroxyethyl methacrylate, itaconic acid (IA, Fluka), and different poly(ethylene glycol) dimethacrylates (550PEGDMA and 875PEGDMA) were prepared by free radical cross-linking copolymerization. All samples were characterized via hemocompatibility and swelling studies in order to investigate the influence of molecular weight and concentration of PEGDMA on hydrogel properties.

EXPERIMENTAL

Materials

The components used in this study (Scheme 1) were 2-hydroxyethyl methacrylate (HEMA, Aldrich), itaconic acid (IA, Fluka), and poly(ethylene glycol) dimethacrylate (PEGDMA, M, 550 and 875; Aldrich). HEMA and PEGDMA components were purified by vacuum distillation before polymerization. All polymerizations were performed in water/ethanol mixture as solvent, using ethyleneglycol dimethacrylate (EGDMA, Aldrich), as cross-linking agent, potassium persulfate (KPS, Fluka), as initiator, and N,N',N'-tetramethylethylene diamine (TEMED, Aldrich), as activator. Buffer solutions of different pH values were prepared using hydrochloric acid (La Chema), potassium chloride (Fluka), potassium monoo and dihydrogenphosphate (Fluka) and sodium hydroxide (Fluka). Demineralized water was used for all polymerizations and the preparation of the buffer solutions.

Synthesis of hydrogels

The P(HEMA/IA/PEGDMA) copolymeric hydrogels were prepared by free radical crosslinking copolymerization. HEMA/IA copolymers were chosen because small contents of IA introduce pH sensitive behavior of copolymers and do not change good chemical stability, high biocompatibility and physicochemical properties similar to those of living tissues of PHEMA homopolymer. Hydrogels containing PEGDMA cross-linker are highly tunable [34–38]. The mechanical properties and the mesh size and swelling ratio can be controlled by varying the molecular weight or concentration of
The reactants were dissolved in water/ethanol mixture (Table 1). The PEGDMA mole fractions were 5, 10 and 15 while the HEMA/IA ratio was kept constant. The samples were designated according to monomers used and the PEGDMA molecular weight and mole fraction as P(HEMA/IA/550(or 875) PEGDMA-5, -10 or -15). The initiator, activator and crosslinker were added to the monomer feed mixture. The polymerization was carried out at 50 °C for 24 h. The reaction mixture was degassed prior to polymerization and placed between two glass plates sealed with a rubber spacer (2 mm thick). After the reaction, the gels were cut into discs and immersed in water for a week to remove unreacted chemicals. The water was changed daily. The discs were dried to xerogels (1 mm thick and 5 mm in diameter). The amount of uncross-linked IA was determined by titration of extract against NaOH (0.05 mol/l) to phenolphthalein end point. On the other hand, the amount of uncrosslinked HEMA and PEGDMA were determined using a UV spectroscopy. In both cases, results indicate that the conversion during cross-linking reaction was nearly complete. The yields of P(HEMA/IA/PEGDMA) copolymeric hydrogels of various compositions were above 99%. All the gels were transparent flexible discs with a limited fluid absorption capability.

**Hemolytic assay**

The hemolytic assay was determined in terms of hemolytic activity of the hydrogels by the direct and indirect contact methods, according to ISO 10 993–4 (1992) [39]. In the direct method, the hydrogel discs were immersed in 5 ml of a physiological solution (PS) to which 0.25 ml of whole rat blood had been added. The PS and distilled water were used as the negative and the positive control, respectively. Then the contents of the tubes were gently mixed and incubated in a water bath at 37 °C for 1 h. Subsequently, the absorbance of the supernatant liquid in each tube was determined at 545 nm using a Pharmacia LKB Ultrospec Plus UV/Vis spectrophotometer and the percentage of hemolysis was calculated. The mean hemolysis value of 5% or less variation from two tests was considered acceptable. In the indirect contact method 5 ml of an isotonic aqueous extract from a hydrogel disc was used with 0.25 ml of a 10% suspension of rat erythrocytes. To prepare the isotonic aqueous extracts, pieces of each disc were kept for 72 h at 37 °C in 100 ml of sterilized water.

Table 1. Feed compositions for the P(HEMA/IA/PEGDMA) hydrogels (same conditions for 550PEGDMA and 875PEGDMA component)

<table>
<thead>
<tr>
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<tr>
<td>HEMA (mol%)</td>
<td>90</td>
<td>85</td>
<td>80</td>
</tr>
<tr>
<td>IA (mol%)</td>
<td>5</td>
<td>5</td>
<td>5</td>
</tr>
<tr>
<td>PEGDMA (mol%)</td>
<td>5</td>
<td>10</td>
<td>15</td>
</tr>
<tr>
<td>initiator (mol%)</td>
<td>0.25</td>
<td>0.25</td>
<td>0.25</td>
</tr>
<tr>
<td>HEMA+IA+PEGDMA (wt.%)</td>
<td>10</td>
<td>10</td>
<td>10</td>
</tr>
<tr>
<td>Demineralized water (wt.%)</td>
<td>45</td>
<td>45</td>
<td>45</td>
</tr>
<tr>
<td>Ethyl alcohol (wt.%)</td>
<td>45</td>
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</table>

Scheme 1. Chemical structures of components used for hydrogel synthesis.
Swelling studies were performed to investigate the swelling properties of P(HEMA/IA/PEGDMA) hydrogels. The changes of P(HEMA/IA/PEGDMA) equilibrium swelling degrees with pH, in a physiologically important pH range (pH 2.20–7.40) at 37 °C, and temperature, in the temperature range from 25 to 55 °C, in buffer of pH 7.40, are presented in Figures 3 and 4. Biocompatibility testing of polymeric biomaterials is an important step in the development of systems for biomedical applications [42]. Hemocompatibility testing is an imperative for medical devices intended for direct or indirect blood exposure. The hemocompatibility of P(HEMA/IA/PEGDMA) hydrogels was evaluated by their testing for hemolysis. In the in vitro testing conditions P(HEMA/IA/PEGDMA) samples in contact with blood showed a mean hemolysis value less than 1.0% in the direct contact assay, and even less than 0.5% in the indirect contact assay. According to the obtained results, these hydrogels are not considered as hemolytic (Figure 1). From our earlier investigations, it is known that incorporation of small amounts of itaconic acid in hydrogels (5 mol%), obtained from natural renewable sources, improves the hemocompatibility of PHEMA-based biomaterial [43]. Also, PEG-based polymer (PEGDMA) is beneficial for favorable hemocompatible behavior [42]. In accordance with the results of hemolytic activity testing all hydrogels exerted favorable hemolytic activity. From Figures 1a and 1b it can be seen that hydrogels with low PEGDMA content (PEGDMA-5) show lower degree of hemolysis, namely better hemocompatibility. The influence of PEGDMA content is more pronounced than the influence of PEGDMA molecular weight. The dry (PEGDMA-15) gel was associated with the presence of the additional C=O group at 1730 cm–1 in the spectrum of P(HEMA/IA/S550PEGDMA-15) gel was associated with the presence of the additional C=O group from IA. On the other hand, several bands appeared in the fingerprint region for ethylene glycol units, originating from PEGDMA component, between 1600 and 1000 cm–1. These peaks were assigned to the –CH2 scissoring band of ethylene glycol units at 1480 cm–1 and the antisymmetric and symmetric stretching bands (–O–R) of ethylene glycol units at 1160 cm–1, respectively. Other characteristic bands represent C–C and C–H vibrations of –CH2 and –CH3 groups. Therefore, the FTIR spectroscopy results confirmed the incorporation of poly(ethylene glycol) dimethacrylates and itaconic acid in hydrogel.\n
Swelling studies

Swelling studies were performed to investigate the influence of the hydrogel composition and external stimulus signal (change of pH and temperature) on the swelling properties of P(HEMA/IA/PEGDMA) hydrogels. The changes of P(HEMA/IA/PEGDMA) equilibrium swelling degrees with pH, in a physiologically important pH range (pH 2.20–7.40) at 37 °C, and temperature, in the temperature range from 25 to 55 °C, in buffer of pH 7.40, are presented in Figures 3 and 4. It is evident from Figure 3 (a and b) that all hydrogels show pH-sensitive behavior, due to IA carboxyl group which gives the hydrogel anionic character and pH sensitivity. The equilibrium degree of swelling, qe, versus pH dependences shows similar trend for all samples. At low pH values (lower than both pK1 and

\[ q_e = \frac{(m_f - m_0)}{m_0} \]
Figure 1. Hemolytic activity of P(HEMA/IA/PEGDMA) hydrogels with different concentrations of 550PEGDMA (a) and 875PEGDMA (b).

Figure 2. FTIR spectra of components: HEMA, IA, PEGDMA, and P(HEMA/IA/550PEGDMA-15) gel.
pK\textsubscript{a2} values \(= pK\textsubscript{a1} = 3.85, pK\textsubscript{a2} = 5.45\), the swelling degrees are low and slightly depend on PEGDMA content, \textit{i.e.}, the swelling degree is low and almost similar for all the samples. The low \(q_e\) values are primary due to the intermolecular physical cross-linking, via hydrogen bond formation, between carboxylic groups in IA, as well as hydroxyl groups of HEMA, with ether groups in PEGDMA residues.

As the pH value of the surrounding medium rises above pK\textsubscript{a} values of both carboxylic groups, \(\text{−COOH}\) groups are transformed into more hydrophilic carboxylate anions, which undergo electrostatic repulsion of same charges on the network chains. Furthermore, due to ionization of \(\text{−COOH}\) groups the hydrogen bonds between network chains are broken and for these two reasons the swelling is substantially increased. In pH range above 3.85 the difference in swelling for samples with different 550PEGDMA content is more pronounced (Figure 3a), \textit{i.e.}, the sample with highest content of 550PEGDMA (P(HEMA/IA/550PEGDMA-15) swells less than the samples P(HEMA/IA/550PEGDMA-5 and P(HEMA/IA/550PEGDMA-10, which is not the case with P(HEMA/IA/875PEGDMA) hydrogels (Figure 3b), where all samples show similar swelling properties.

The similar situation is in the case of \(q_e\) \textit{versus} temperature dependences (Figures 4a and b), which all show the same trend regarding the shape of the curves. It is also evident that P(HEMA/IA/550PEGDMA-5) and P(HEMA/IA/550PEGDMA-10) hydrogels show similar \(q_e\) values, which are higher than that for P(HEMA/IA/550PEGDMA-15) sample, while \(q_e\) values for P(HEMA/IA/875PEGDMA) do not depend much on 875PEGDMA.

Figure 3. pH sensitive swelling behaviour of P(HEMA/IA/PEGDMA) hydrogels with 550PEGDMA (a) and with 875PEGDMA (b).
content. In the temperature range 37–50 °C there is a small decrease in swelling with the increase of 875PEGDMA content. The PHEMA/IA/PEGDMA) hydrogels are not temperature-sensitive, they have no LCST, but they show temperature dependent swelling. It is known that hydrogels based on HEMA homopolymer or its copolymers with IA have no LCST in the investigated temperature range [16,44].

The difference in swelling capacity with the change of 550PEGDMA content can probably be explained by the following argument. By introducing PEGDMA chains in the hydrogel structure two opposing effects take place: PEGDMA chains are hydrophilic and highly elastic, so they will induce higher swelling, but at the same time they act as a cross-linking agent, which will reduce the swelling degree. From the results obtained it is obvious that the chain length and content of PEGDMA is the main factor which determines which of those effects will prevail. This effect is clearly visible only for the samples containing 550PEGDMA because shorter 550PEGDMA chains are less flexible so there is a critical concentration (15 mol%) where the cross-linking effect outweighs the chain flexibility effect. The longer 875PEGDMA chains are more flexible and therefore hydrogel swelling behaviour is not much influenced with 875PEGDMA contents used in this work.

Therefore, the change of environmental conditions tunes hydrogel swelling. These pH responsive and tem-

Figure 4. Temperature dependent swelling behaviour of P(HEMA/IA/PEGDMA) hydrogels with 550PEGDMA (a) and with 875PEGDMA (b).
perature dependent hydrogel properties along with appropriate swelling degrees values could be beneficial for their use as drug delivery systems. The pH responsive hydrogels play a significant role in controlled drug delivery systems [45]. These delivery systems exhibit substantial changes of swelling degrees in various physiological media. Due to low swelling in acidic and higher swelling in neutral and basic media, they can protect protein drugs from denaturation in acidic media and deliver drugs in media with higher pH values.

**Network parameters**

Properties of hydrogels with weak acid moieties depend on the network structure, which is controlled by the feed composition. The most important network parameters are the molar mass of the polymer chain between two neighboring cross-linking points, $M_c$, the effective crosslinking density, $\nu_c$, and pore size, $\xi$. In order to determine $M_c$ for hydrogels containing diprotic itaconic acid, the following equation is used [46]:

$$V, X^2 \phi_r^2 \left( \frac{2K_{a1}K_{a2} + 10^{-aa}K_{a1}}{2(10^{-aa})^2 + 10^{-aa}K_{a1} + K_{a1}K_{a2}} \right)^2 = \left[ \ln(1 - \phi) + \phi + \chi \phi^2 \right] + \frac{V, \rho}{\overline{\rho}} \phi_3^{2/3} \phi_r^{1/3}$$

(2)

where $M_c$ is the molar mass of the polymer chain between two neighboring crosslinking points, $K_{a1}$ and $K_{a2}$ are the first and second dissociation constants of a diprotic acid, $X$ is the weight fraction of ionizable polymer in the system, $l$ is ionic strength of the swelling medium, $\phi_r$ is the polymer volume fraction in the swollen gel, $\phi$ is the polymer volume fraction in the relaxed state, $V, l$ is the molar volume of water, $\rho$ is the polymer density, $\overline{\rho}$ is the average molar volume of polymer repeating units, and $\chi$ is the Flory polymer–solvent interaction parameter. The effective crosslinking density, $\nu_c$, was calculated as $\nu_c = \rho / M_c$.

The pore size, $\xi$, which is a term that describes the available space for solute transport within the polymer network, is also an important parameter in analyzing crosslinked polymers and calculated according to Eq. (3), which is described in more detail by Canal and Peppas [47]:

$$\xi = \phi_r^{1/3} \left( \frac{2C_{\overline{\rho}}}{M_c} \right)^{1/2}$$

(3)

Here, $M_c$ is the molecular weight of the repeating unit; $l$ the C–C bond length (1.54 $\times$ 10$^{-10}$ m for vinyl polymers); and $C_{\overline{\rho}}$, the Flory characteristic ratio, $\overline{\rho}$ is the average of the molar mass of repeating unit [48].

According to the potential biomedical application in dermatology and drug delivery systems, the calculations were done for the results obtained at pH 7.40, and 37 °C (Table 2).

It can be seen that network parameters for P(HEMA/IA/PEGDMA) hydrogels depend on PEGDMA content in hydrogels and PEG chain length (550 and 875) in the similar way as in the case of $q_e$ vs pH and $q_e$ vs temperature dependencies. The values of the molar mass of the polymer chain between two neighboring cross-linking points ($M_c$) are in the range of 4715–7634 g/mol for P(HEMA/IA/550PEGDMA) samples and from 6392 to 8159 for P(HEMA/IA/875PEGDMA) samples. The pore size values of P(HEMA/IA/550PEGDMA) and P(HEMA/IA/875PEGDMA) hydrogels are in the range of 6.7–9.5 nm and 6.85–9.35, respectively, i.e. they all are in the nonporous regime.

### Table 2. Network parameters of P(HEMA/IA/PEGDMA) hydrogels at pH of 7.40, and 37 °C

<table>
<thead>
<tr>
<th>Sample</th>
<th>$M_c$ / g mol$^{-1}$</th>
<th>$\nu_c$ / mol dm$^{-3}$</th>
<th>$\xi$ / nm</th>
</tr>
</thead>
<tbody>
<tr>
<td>P(HEMA/IA/550PEGDMA-5)</td>
<td>7634</td>
<td>0.424</td>
<td>9.52</td>
</tr>
<tr>
<td>P(HEMA/IA/550PEGDMA-10)</td>
<td>7945</td>
<td>0.441</td>
<td>8.56</td>
</tr>
<tr>
<td>P(HEMA/IA/550PEGDMA-15)</td>
<td>4715</td>
<td>0.718</td>
<td>6.70</td>
</tr>
<tr>
<td>P(HEMA/IA/875PEGDMA-5)</td>
<td>8159</td>
<td>0.425</td>
<td>9.35</td>
</tr>
<tr>
<td>P(HEMA/IA/875PEGDMA-10)</td>
<td>7752</td>
<td>0.463</td>
<td>8.18</td>
</tr>
<tr>
<td>P(HEMA/IA/875PEGDMA-15)</td>
<td>6392</td>
<td>0.451</td>
<td>6.85</td>
</tr>
</tbody>
</table>

CONCLUSION

Free-radical crosslinking/copolymerization of 2-hydroxyethyl methacrylate with IA and PEGDMA, with varying molecular weight and content, resulted in the formation of cross-linked P(HEMA/IA/PEGDMA) hydrogels. The aim of our study was to demonstrate the influence of poly(ethylene glycol) chain length and content on the properties of P(HEMA/IA/PEGDMA) hydrogels. All hydrogels show a fair level of blood compatibility, as confirmed by *in vitro* experiments of percentage hemolysis, and can be described as hydrogels with beneficial hemocompatible behaviour. Swelling studies showed typical pH-sensitive swelling behaviour of anionic hydrogels, as well as the temperature dependent swelling. Itaconic acid is responsible for pH sensitive behavior and short, flexible poly(ethylene glycol) dimethacrylate...
chains inside the network play a role of elastic springs but at the same time act as crosslinking agent. The swelling of P(HEMA/IA/550PEGDMA) hydrogels, monitored as a function of pH and temperature, is influenced by the content of 550PEGDMA component. By introducing lower concentration of 550PEGDMA chains there is a critical concentration of 15 mol% where the cross-linking effect outweighs the chain flexibility effect so they swell less than those with lower 550PEGDMA content (5 and 10 mol%). In contrast, P(HEMA/IA/875PEGDMA) hydrogels show very small change in swelling with the change of 875PEGDMA content. The longer 875PEGDMA chains are more flexible and the crosslinking effect is balanced with the chain flexibility effect, so the swelling behavior is similar for all samples in the concentration range used in this study. The obtained results indicate that pH–responsive and temperature dependent P(HEMA/IA/PEGDMA) hydrogels show good potential for biomedical applications.

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REFERENCES


IZVOD

PROUČAVANJE HEMOKOMPATIBILNOSTI I BUBRENJA HIDROGELOVA POLI(2-HIDROKSIEITIL METAKRILAT-KO-
-ITAKONSKA KISELINA-KO-POLI(ETILEN GLIKOL) DIMETAKRILAT)

Sava N. Dobić, Jovana S. Jovašević, Marija D. Vojisavljević, Simonida Lj. Tomić

Univerzitet u Beogradu, Tehnološko–metalurški fakultet, Beograd, Srbija

(Naučni rad)

U radu je izvedena sinteza dva nova tipa kopolimernih hidrogelova na bazi 2-
-hidroksietil (met)akrilata, itakonske kiseline i poli(etilen glikol) dimetakrilata, s
ciljem primene ovih polimernih sistema u biomedicinske svrhe. Testirana je
biokompatibilnost preko probe hemokompatibilnosti. Hemolitička aktivnost svih
hidrogelova je bila u dozvoljenim granicama, prihvatljivim za biomedicinsku pri-
menu. Studije bubrenja hidrogelova, izvedene u opsegu fizioloških pH i tempera-
turnih vrednosti, pokazale su da bubrenje zavisi od pH i temperature. Sintetisani
hidrogelovi su pokazali sličan trend zavisnosti stepena bubrenja od pH i tem-
perature. Sadržaj 550PEGDMA u hidrogelu utiče na stepen bubrenja, dok kon-
centracija 875PEGDMA komponente vrlo malo utiče na stepen bubrenja. Hidrogel
sa 15 mol% 550PEGDMA manje bubri od uzoraka sa manjim sadržajem ove
komponente (5 i 10 mol%), što ukazuje da pri toj koncentraciji prevladava umre-
žavajuće dejstvo ove komponente. Duži PEG lanci u 875PEGDMA su fleksibilniji, pa
je efekat umrežavanja uravnotežen sa efektom fleksibilnosti lanca, što ima za
posledicu slično bubrenje za sve uzorke u opsegu koncentracija koje su korišćene
u ovom radu.

Ključne reči: 2-Hidroksištil metakrilat • Itakonska kiselina • Poli(etilen glikol) di-
metakrilat • Hidrogelovi • Hemokompa-
tibilnost • pH-Osetljivo i temperaturno zavisno bubrenje • Parametri mreže