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Applied Surface Science xxx (2018) xxx-xxx



Contents lists available at ScienceDirect

Applied Surface Science



journal homepage: www.elsevier.com/locate/apsusc

Full Length Article

Metal oxide curcumin incorporated polymer patches for wound healing

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ARTICLE INFO

Article history: Received 11 October 2017 Received in revised form 27 December 2017 Accepted 16 January 2018 Available online xxx

Keywords: Sodium alginate Poly vinyl alcohol Curcumin Titanium dioxide nanoparticles Wound healing

1. Introduction

ABSTRACT

Sodium alginate and Poly vinyl alcohol (PVA) based scaffolds are very well used for wound healing purpose. Sodium alginate is a natural polymer which is used for many biomedical applications. PVA is a biocompatible polymer used in many biomedical applications. Incorporation of Titanium dioxide (TiO₂) nanoparticles in the scaffold increases the wound healing property of the scaffold. TiO₂ nanoparticles are biocompatible. They enhance the wound healing activity by photo catalytic property. Curcumin is used for the treatment of inflammation. It is the main active compound of turmeric Curcumin is used for treatment of wound and inflammation. It had antimicrobial and antioxidant property.

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Incorporation of metal oxide nanoparticles inside polymer scaffolds has been of great interest nowadays. These scaffolds are used in the wound healing activity. This incorporated nanoparticles increases the antibacterial efficiency of the scaffold [1]. Hemostasis, inflammation, proliferation and remodelling are the four steps of wound healing. Infections creates lots of problem in wound care management. They cause exudate formation. It will delay wound healing. The infectious microorganisms will enter the body through the wounds. Then it forms colonies. The internal portion of the body will also be invaded by the microorganisms. This condition will lead to lethal infections. The best way to avoid such serious conditions is by using wound healing scaffolds with antibacterial property [2–5].

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https://doi.org/10.1016/j.apsusc.2018.01.143 0169-4332/© 2018 Elsevier B.V. All rights reserved. Titanium dioxide nanoparticles (TiO₂ NPs) are biocompatible materials. There is no toxicity with TiO₂ nanoparticles both *in vitro* and *in vivo*. It can be used in biomedical applications for various applications like drug delivery, antibacterial coatings *etc*. It has antibacterial, antifungal and anticancer properties. It is an inert material *in vivo*. TiO₂ is a very important material that can be used in life science. The reactivity of titanium dioxide nanoparticles is very high. The TiO₂ is widely used in paint, pharmaceutical, and cosmetics industries [6,7].

Alginates are used for man biomedical applications. They are biocompatible polymers. They are non-toxic polymers. They are used for many applications like drug delivery, drug targeting, wound healing *etc.* Alginates cannot be processed other than spherical forms because of its rigid and fragile nature. They are biocompatible and biodegradable in nature. Alginates are unbranched binary copolymers. The structure of alginate is 1-4 glycosidicallylinked b-Dmannuronic acid (M) and its C-5 epimer a-L-guluronic acid (G). It is obtained from marine brown algae. Sodium alginate is used as gelling agent, as stabilizers and as blood expander in pharmaceutical industry. It forms a viscous solution when dissolved

Please cite this article in press as: T.S. R., et al., Metal oxide curcumin incorporated polymer patches for wound healing, Appl. Surf. Sci. (2018), https://doi.org/10.1016/j.apsusc.2018.01.143

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in water [8–12]. Poly Vinyl alcohol (PVA) is used in biomaterial application for a long time. It is s synthetic polymer. It is non-toxic, biocompatible. Poly vinyl alcohol belongs to the group of poly hydroxy polymer. Poly vinyl alcohol is water soluble. It forms fibre, it is resistant to chemicals and it is biodegradable. The interaction between sodium alginate and poly vinyl alcohol is through hydrogen bonds. Due to these advantages poly vinyl alcohol is used in medical, cosmetic, food, pharmaceutical and packaging applica-tions.

In this work we have prepared SA/PVA composite scaffolds by gel casting method. TiO_2 nanoparticles were prepared and characterized. Before casting TiO_2 nanoparticles and curcumin were added to the SA/PVA composite. The resultant scaffold is characterized and analysed for antibacterial property.

2. Experimental procedure

2.1. Synthesis of titanium dioxide nanoparticles

Titanium tetra isopropoxide was added to HNO_3 and agitated for 2 h. The pH was maintained as 3. The suspension was centrifuged and washed with H_2O . TiO₂ nanoparticles were dried for 1 h at 100 °C. The resulting powder was then calcinated 450 °C for 3 h [13].

2.2. Synthesis of patches

The solutions of Sodium alginate (2%) and Polyvinyl alcohol (16%) are prepared by dissolving them separately in 100 ml of distilled water. PVA solution is stirred and simultaneously heated up to 80–90 °C to prevent from gel formation. The solution of Sodium alginate is made to spin in the spinner. But, the ratio 1:1 of SA and PVA combination could obtain a fine fibre. Thus 0.5 g and 4 g of SA and PVA are dissolved and stirred in 25 ml of H₂O separately and 5 ml from each solution is taken and allowed to spin in an Electro spinner about 10 min with 750 rpm.

5 ml solution of SA and PVA are mixed together and allowed to spin for few minutes. In this polymer, 0.1 g of TiO₂ nanoparticles are added slowly to form a composite fibre of SA/PVA/TiO₂. Then in a solution of Dichloromethane (5 ml) containing 0.5 g of curcumin is dissolved and stirred. From this 0.5 ml of the solution is added to the composite containing SA/PVA/TiO₂.

2.3. Cross linking of sodium alginate/PVA/TiO₂ patch

The sodium alginate/PVA/TiO₂ patches were treated with 2% glutyraldehyde vapours for 48 h and then dipped in 1% calcium chloride solution for 1 h.

2.4. In vitro evaluation of antibacterial and antifungal activity

The antibacterial activity of titanium dioxide nanoparticles were done against *Bacillus subtilis, Klebsiella pneumonia, Staphylococcus aureus, Escherichia coli* and antifungal activity against *Candida albicans and Aspergillus niger* using agar diffusion method [14]. The antibacterial activity of scaffolds were analysed against *Bacillus subtilis, Klebsiella pneumonia.* The patches are placed on the agar plates. Nutrient Agar (NA) plates were inoculated with test organisms. The plates were evenly spread out. Then wells were prepared in the plates with a cork borer. The well was loaded with 10 µg of Streptomycin dissolved in 1 ml of DMSO was used as a Positive control for antibacterial activity. The plates were incubated for 24 h at 37 °C. The development of inhibition zone around the well was measured and recorded.

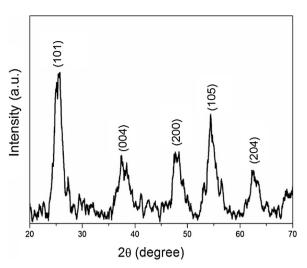


Fig. 1. XRD of TiO₂ Nanoparticles.

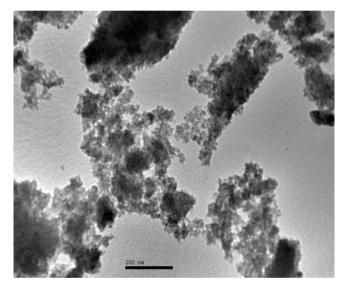


Fig. 2. TEM of TiO₂ Nanoparticles (Insert DLS).

3. Results and discussion

The XRD image of TiO₂ nanoparticles annealed at 450 °C is given in Fig. 1. The synthesised sample is confirmed as anatase phase by XRD image. The peaks which appears at 2θ values of 25.20°, 37.80°, 48.04°, 53.890° and 62.68° correlates to the planes (101), (004), (200), (105) and (204). This confirms that the nanoparticles formed is titanium dioxide anatase phase.

The titanium dioxide nanoparticles nanoparticles synthesised are very small and spherical in shape. The size of the formed particles is found to be around 12 nm. This is confirmed by TEM analysis, Fig. 2. The particles are agglomerated.

Fig. 3 gives the FTIR image of TiO_2 nanoparticles. The band at 3500 cm⁻¹ is due to the hydroxyl group stretching vibrations. The vibration due to Ti–O–Ti is visible at 1139 cm⁻¹. The bending vibration of H₂O and Ti–OH is seen at 1628 cm⁻¹ and 1784 cm⁻¹. Vibration due to Ti–O bond is seen at 1383 cm⁻¹. The vibrations of Ti–O–Ti band and O–Ti–O band are present at 490 cm⁻¹ and 400–1250 cm⁻¹ respectively [15].

All the six modes of Raman spectrum of anatase TiO_2 are shown in Fig. 4. Three of these modes are Raman active and three modes are infrared active. One mode is inactive in both infrared and Raman spectra. The band at $147 \,\mathrm{cm}^{-1}$ confirms that the nanoparticles

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