A Review of Antiviral Polymer Nano Composites (PNC) in Biomedical Applications; Coronavirus Pandemic Situation

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Abstract

Since December 2019, scientists and engineers have tried to minimize and control Covid-19 spreading by different ways. These ways usually focused on cutting the circlet of Covid-19 infection by both direct and non-direct ways. As a polymer composites widely used in every day products, a short review of Polymer Nano Composites (PNCs) for antiviral activity against Covid-19 has been achieved. Firstly, virus's structure, especially Covid-19 has been discussed. Then, different PNCs applications in biomedical that work against coronavirus have been presented and discussed. Also, comprehensive review of antiviral activity mechanisms have been reported with selected examples of PNCs and applications.

Keywords: Biomaterials; Covid-19; Nano Composites; Polymers; Antiviral; Health Care

Abbreviations: PNCs: Polymer Nano Composites; PPE: Personal Protection Equipment; PES: Polyethersulfone; PMMA: Polymethylmethacrylate; PP: Polypropylene; ACE2: Angiotensin-Converting Enzyme 2; ROS: Reactive Oxygen Species; PDMS: Polydimethylsiloxane; PEI: Polyethylenimine

Introduction

In recent years, Polymer Nano Composites (PNC) field have been developed for many biomedical applications as antiviral, antibacterial, and antifungal agents, and have been used in many products in our daily life [1]. Additionally, the antiviral activity of PNC has reached in the top of the list research of polymer, nanotechnology, and materials science since Covid-19 was discovered in December 2019 [2,3]. Moreover, there were more than 76 million confirmed cases with more than 1.6 million deaths because Covid-19 until Friday 20 December 2020, as it is clear on Johns Hopkins University website, Johns Hopkins University, 2020 [4]. This has high impact on all life sectors, such as economic, education, and social. As PNCs have been used widely in every day products in health care system, for example the personal protection equipment (PPE), the improvement in antiviral activity of these composites could be helpful for both heath system and public society against Covid-19. For example, PNCs based on copper oxide nano particles have high ability to work as an antiviral agent in PPE [5]. In this review, the probable role of PNC as an antiviral agent against coronavirus has been highlighted in terms of the more contact products in human life, such as drug, coating, cloths and house products.

Polymer Nano Composite (PNC) against Coronavirus

PNC has high opportunity to work against coronavirus because of its scope activity that located in nanoscale measurements, as it shown in Figure 1A [6]. Also, the

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ingredients and structure of coronavirus belong to nano dimensions, which encourage PNC to be a highly recommended factor in this case, as it is clear also from Figure 1B [7]. Generally, the coronavirus spreads by different ways, without contact or with contact as directly or indirectly, as it is clear in Figure 2 [8]. For carrier drops of viruses that less than 5 microns, can spread during the air from infected to non-infected person. In this case, there are two main ways to stopping this spreading, by making social distance or wearing a mask, which mainly could improve it by PNC layer. For carrier drops of viruses that more than 5 microns, can collect on the product surfaces to spread the virus indirectly in a contact situation, or on human body parts such as hands to spread the virus directly in contact case too. To minimize the spreading in these situations, especially in indirectly, in contact case, the products have to be coated with coating based on PNC.





In addition to that, there are different ways and applications of PNC to protect human against viruses, such as drug carrier, drug delivery system, nanoparticles, biopolymers and protective equipment's, as it shown in Figure 3 [8]. Figure 4 represent materials kinds and nano materials based, respectively, which are used as antiviral agents [9]. As PNC have wide important properties such as high mass and easy production, cheap, high ability of sterilization and disinfection comparing with others kinds of materials, they have been used in many biomedical applications, as it was shown in Figure 5 [5]. The antiviral PNC have been prepared, produced and investigated with different filled nanoparticles such as silver, titanium dioxide, and zinc oxide nanoparticles for both human cells, and life products coating applications [1, 21]. This activity works mainly on the principles of making the barrier between the viruses host cells and other cells of the body or repelling the virus host drops with high repulsive forces on the products and surfaces. These forces have been confirmed to play a significant factor in different virus's activities in vivo and on the product surfaces, which might raise thoughts to apply that in these days to minimize coronavirus infections, and for better management of economic and social life activates [10].





An important application also that PNC based on polyethersulfone (PES), polymethylmethacrylate (PMMA), and polypropylene (PP) have been used in plasmapheresis process to collect blood plasma from recovered patients who infected by Covid-19 and gives it to another infected patients [5]. Another important application is using PNC as an antiviral agent such as antiviral coating, antiviral drug and medical devices like personal protective equipment (PPE) that could help in coronavirus pandemic situation, where the necessary time to kill the viruses has been found within ten minutes of virus exposure in average [5].



Figure 6 illustrates some products that were coated by Cu2O/polyurethane as a PNC for its antiviral activity against coronavirus [11]. Figure 7 summarizes using PNC for therapy of Covid-19 by different ways [12]. The First approach may be done by adding nano additives into PPE to prevent catching virus by the respiratory system. Another way could

be by applying nanomaterials into the drug delivery system to the pulmonary system by inhalators. Also, using PNC to preventing cellular binding of virus by targeting angiotensinconverting enzyme 2 (ACE₂) receptors or viral S protein in virus membrane.



Mechanisms of Antiviral

There are three mechanisms of antiviral activities for artificial materials, which are depending on heat, ions or small molecules techniques, as it is clear in Figure 8 [13]. For example, PNC based on cupper nano particles work as antiviral surfaces by heat generation technique because of its high heat connectivity properties, as it was shown in Figure 9 [14]. Figure 10 illustrates ions technique for an antiviral activity of PNC [8]. In which some PNC materials might produce an ions by light that work as reactive oxygen species (ROS), which damages of the membrane, proteins and DBA/ RNA of viruses. Small molecules such as nano particles or some polymer chains could make a barrier between the host and virus to prevent Covid-19 spreading, as it is clear in Figure 11, which represents an antiviral activity by small molecules (nano) technique [15].







Figure 10: Photodynamic inhibition of viruses by reactive oxygen species (ROS) induced damage of the membrane, proteins and DBA/RNA [8].

There are different ingredients of PNC that work as antiviral agent [16]. For example, PNC coating may consist of pharmaceutically admissible salts of chlorhexidine, and a lubricating agent that does not significantly be adsorbed to the anti-infective agent. Also, some PNC structures have a porous plastic layers and antiviral agents, which could be applied into vents, and filters applications. Another PNC has high hydrophilic or self-cleaning ability to repel viruses away from surfaces. A successful example to use PNC as a coating layer for antiviral applications is polycationic PEI, as it shown in Figure 12 in the schematic representation of its antiviral mechanism against virus [8]. For PNC based on copper nano particles, the antiviral activity might be simply started by adsorption of microorganisms such as coronavirus on the PNC surface, as it was shown in Figure 13 [19]. Then, diffusion of water through the PNC layer to start the corrosion process by oxygen and copper ions. Consequently, copper ions damage the microorganism cell membrane, to damage RNA also.



Figure 12: Schematic representation of antiviral mechanism of virus inactivation in polycationic PEI derivative coatings [8].



copper nano particles [17].

PNC in Drug System

One of the most important biomedical applications of PNC is in the drug industries and drug delivery systems especially as an antiviral drug, for example heparin polysaccharides against HPV16 [2]. Martinez LS, et al. [2] studied the antiviral ability of polysaccharides polymers with oligomers of sulfated-glycomimetic in vivo for different kinds of viruses such as HPV16, Influenza a Virus (143 IAV), and Virus (HSV) [2].

As we can see in Figure 14 that illustrates the schematic model proposed of the inhibition mechanism for antiviral PNC by competing with virus to prevent it enters the cells. The results shown that these polymers could prevent spreading the viruses inside the human tissue environment by providing a barrier and covering layers around the active sites on cells wall, which are targeted by viruses [2]. These results might open the light on the possibility of those types of PNC to apply against Covid-19 spreading, at least to manage and control the high critical cases that have infected by a high percentage of coronavirus. Figure 15 shows different forms of carriers based on PNC that work as an antiviral agent [18].



Figure 14: A schematic model represents the inhibition mechanism for antiviral PNC by competing with virus to prevent it enters the cells [2].



PNC in Daily Products

PNC has been used in a wide range of daily routine applications especially that are in contact with human directly, such as cloths, and fabric industries, and products. The most popular antiviral nanoparticles loaded in polymer including silver, copper, zinc oxide, titanium dioxide, and many metals oxides [1]. Antiviral based-cotton fabrics have been coated by polydimethylsiloxane (PDMS) which was loaded with Couper oxide (CuO) nanoparticles by using different cross linkers [1]. Figure 16 shows the contact angle of a water drop on cotton fabric with and without the coating layer of PDMS/ CuO. The high contact angle of coated fabric might give a chance to decrease spreading Covid-19, especially for people who work in high environment of viruses, such as hospitals and healthcare centres.



Figure 16: Pictures of the contact angle of a water drop on cotton fabric; a) without, b) with the coating layer of PDMS/CuO [1].

It has been discovered that polycations polymers that has hydrophobic ability and insoluble in water, such as N, N-dodecyl methyl-polyethylenimine (PEI), can kill bacteria and viruses by high polycationic chains forces [10]. These forces destroy the lipid layer membrane that protects the viruses, which might works probably with Covid-19. For example, it can apply to coat the walls, doors, windows and all high contact surfaces to make these surfaces have selfantiviral ability. Another important example of PNC coating that has high ability of antiviral activity was illustrated by Figure 17 [19]. As it has been discovered that The PVA based Cu-Gr (copper-graphene) nanocomposite can be used to generate a highly transparent antiviral coating of the cell phone screen [19].



Additionally, one of the important applications of antiviral ability of PNC is to apply its layer in PPE, such as the mask fabrication to increase its ability to non-infectious and preventing viruses spreading. As an example, adding fourth layer to three layers mask structure that shown in Figure 18, to produce four layers mask structure that shown in Figure 19 [6,20,21]. An example of PNC that could use is based on nylon 6/poly acrylonitrile or polyethylene/polypropylene nanofibers layer that have been prepared by electrospinning-netting technique, as it were shown in Figure 20 [6].



INNER LAYER Non-toxic, non-irritant, soft and comfortable non-woven fabric materials OUTER LAYER Non-woven fabric materials coating with antiviral materials (Optional)

Figure 19: Illustration showing three layers mask structure [20].

BARRIER LAYER

Melt-blown materia

Made or coated by

antiviral materials



Figure 20: Illustration of an example of PNC based on nylon 6/poly acrylonitrile or polyethylene/polypropylene nanofibers layer [6].

Conclusion

Polymer Nano Composites (PNC) could be a significant factor in the developing process of antiviral drug toward Covid-19, such as polysaccharides nanocomposites. Another PNC might play a crucial solder in the cutting of coronavirus circle spreading's by introduce a barrier between productproduct and product-human infections. This can be a simple and limited products like spoons and door knobs, or even wide range application uses such as entirely wall coating of house walls. Consequently, PNC could save spending the money and energy, management the health system, decreasing the lockdown periods and social distancing as well.

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