Review Article

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Electrospun cellulose acetate nanofibers and Au@AgNPs for antimicrobial activity - A mini review

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Abstract: Au@Ag nanoparticles decorated on cellulose paper could be worthwhile biomedical applications. Electrospinning technique is broadly employed for fabrication of nano and micro size fibers with a variety of biopolymers adding cellulose acetate nanofibers. Evolutions in cellulose research demonstrate that it is an anticipating material for the biomedical application. Nanofibers acquired by electrospinning technique were utilized in various biomedical applications. In this report, electrospinning of cellulose acetate, the solvent choice for cellulose acetate e-spun nanofabrication and decoration of AgNPs including shape and size for antimicrobial activity are argued.

Keywords: electrospinning technique; cellulose acetate; solvent; silver nanoparticles; antimicrobial activity

1 Introduction

Before the starting into electrospun cellulose acetate nanofibers, it would be productive to elaborate on the fundamentals of electrospinning technique. As the origin of this century, investigators all over the world have been reluctance at a century old procedure [1, 2] presently known as electrospinning. Likely unidentified to most investigators for most of the last century, electrospinning is able to develop uninterrupted fibers from the submicron diameter down to the nanometer diameter. It was not until the mid-1990s with concern in the area of nanotechnol-
Figure 1: a) Simple procedure of electrospinning technique b) different types of spinnerate c) different types of collectors
solution is released from the tip of the Taylor cone. The fired polymer fluid jet changes an unbalance and extension procedure, which permits the jet to turn very long and thin. The solvent vaporizes, allowing for a charged polymeric fabric.

Furthermore, particular changes to the fundamental experimental set-up have been established for developing the level of nano-fibers and for adjusting their possessions corresponding to demands of distinct uses. In fact, in horizon of prominent level output and of uses those obviously demonstrate important infomercial potency. The uniaxial placement of nano size fibers amongst the prescribed lay out is presented to be significant for various uses, therefore getting a significant requirement to be brought into importance in level high of the e-spun industry potentiality. In the area of tissue re-formation employed to muscles, bone and cartilage meniscus, and neural cells, different reports demonstrate that cell cultures on uniaxially aligned nano size fabrics scaffolds advantageous lengthen the nanofiber lengthwise axis [9]. The high volumes are particularly suitable for biomedical uses such as tissue engineering, design of wound healing, drug delivery, and for textile uses. In the theory of tissue engineering, e-spinning is especially desirable for the manufacturing of three dimensional composites afterward the concluding results accommodate almost of demands in conditions of substantial porousness (determined as the number of three dimensional, unsatisfied changes in the scaffolds, large surface area to volume ratio, and adjustable mechanical dimensions [10]. For wound curing or wound burn, e-spun, interconnected hooley constructions are advantageous for exuding fluidize, restraining evacuation and helping oxygen permeation, and additionally, the flexibleness of the e-spinning procedure permits the accession of agents in the scaffolds, for health check discussion and antimicrobial uses. The concept to choose and right mix solution factors and line up their ratio to grants one to efficacy orient the attributes of e-spinning of nanofibers to get novel and in demand structures and practicalities.

2 Cellulose acetate (CA) electrospun nanofibers

CA is a easily obtainable substantial derived from cellulose at low cost and it is employed in the broad scope in sheets for food technology and medical applications because of its less perniciousness, good hydrolytic stability, and environmentally well-disposed attributes [11]. In addition, development in cellulose research demonstrates that it is an promising biological substance for tissue engineering, stem cell research, and regenerative medicine [12]. Bacterial cellulose is easily examined for bone regeneration. Thus, it doesn’t propose the quality to assure the fabrics on the nano size or micro size, which bounds its pertinency in tissue engineering. Moreover, it is examined that, CA is easy in electrospinning as compared to cellulose. CA nanofibers can be easily converted from cellulose acetate to cellulose by alkaline hydrolysis; researching the many attributes of cellulose acetate in the nano size governing is a pertaining proposal. Within many, e-spinning has conferred a completely novel appearance to the uses of CA fibers or nanofibers. Previously reported by worldwide industriousness investigation, the global mart of CA is assigned to about 1.05 M metric tons by 2017 (http://www.strategyr.com/Cellulose_Acetate_Market_Report.asp) and the main worldwide individuals regard Celanese Corporation, Daicel Corporation, Primester, Eastman Chemical Company amongst many. CA demonstrates a broad concept [13] evidencing its value in polymers investigation. The usual level of subtle of 2.45-2.5 discusses better solubility (in various kind of solvents) and meltability [14].

In previous report, it has directed extinct the environmental characteristic of CA situated fabrics evaluated in status of their biodegradability as a purpose of the synergy amongst different user consequences [15]. Lately, Group of Sousa have reported a dynamic work, concentrated on the subglass mobility in CA beneath geomorphic various constitutes [16]. Cellulose acetate NFs has relatively more modulus and enough angularity and tensile strength [17]. Conjoining with functional groups enables other concerning concept to cellulose acetate such as, surface modification with heavy metal ions [18]. Electrospun cellulose acetate nanofibers have experience a retentive journey and quaternary period conveys a exertions throughout the range of different fields. Herein, we have briefly discussed electrospinning of CA nanofibers for antimicrobial activity. Moreover, the effective and better handled purpose of e-spinning method should be modified for the structural fibers [19, 20]. Synthesized fibers have developed many-sided uses in biomedical field [21–29]. Cellulose acetate has been studied due to its more beneficial characteristics which are; decomposable, compatible with biological tissue, not soluble in H2O etc. CA can be possibly employed for injure bandage and broadly employed for different form of possible exertions i-e germicide fibrous sheets, nanocomposites for biomedical uses etc [15, 30, 31].
Table 1: Parameters for electrospinning of cellulose acetate nanofibers

| Solution % | Solvent | Solvent ratio | Voltage kV | Distance cm | Feed rate | Reference |
|------------|---------|---------------|------------|-------------|-----------|-----------|
| 17         | acetone/N,N-dimethylacetamide | 2:1 | 17.5 | 15 cm | 0.6 mL/h | [48] |
| 20         | acetone/N,N-dimethylacetamide | 2:1 | 8 | N/A | 12 L/min | [49] |
| 13, 19, 25 | acetone/N,N-dimethylacetamide | 3:1:5 | 18 | 16 | 1 mL/h | [50, 51] |
| 6, 8, 10, 12 | 4,4-diphenylmethane | 1:2 | 20 | 17 | 0.5 mL/h | [52] |
| 18         | Acetone/ N,N-dimethylformamide | 3:2 | 12 | 15 | N/A | [53] |
| 15         | Acetone/ N,N-dimethylformamide | 3:2 | 18 | 16 | 1 mL/h | [54] |
| 17         | acetone/N,N-dimethylacetamide | 2:1 | 17 | 15 | 1 mL/h | [55] |
| 10         | acetone/Polyethylene oxide | N/A | N/A | N/A | N/A | [56] |
| 20         | N,N-dimethyl acetamide / dimethyl sulfoxide | N/A | N/A | N/A | N/A | [57] |
| 20         | N,N-dimethyl formamide | - | 25 | 20 | 0.5 mm | [58] |
| 20         | acetone/N,N-dimethylacetamide | 2:1 | 16 | 16 | 0.02 mL/min | [59] |
| 13         | acetone/N,N-dimethylacetamide | 2:1 | 25 | 25 | 3 mL/h | [60] |
| 15         | acetone/N,N-dimethylacetamide | 2:1 | 14.25 | 25 | 1 mL/h | [61] |
| 17         | acetone/N,N-dimethylacetamide | 2:1 | 25 | 25 | 1 mL/h | [62] |
| 12.5       | acetone/N,N-dimethylacetamide | 2:1 | 20 | 20 | 1 mL/h | [63] |
| 12         | dichloromethane/methanol | 4:1 | 15 | 10 | 1 mL/h | [64] |
| 15         | acetone | - | - | 15 | 3 or 5 mL/h | [65] |
| 9, 11, 13, 15 | Acetone/water | 80:20, 90:10 | 12-15 | 20-22 | 2 to 4 mL/h | [66] |

2.1 Choice of solvent for cellulose acetate e-spun nanofabrication

In 1934 [32], Formhals tried the e-spinning of cellulose acetate nanofibers by utilizing acetone as the solvent. CA nanofibers have been synthesized by using different conditions Table 1. Here we are describing selection of solvent for CA electrospinning. The solvent choice is important and has been conventionally grounded on contest and occurrence, outcomes of solvability methods set by physical chemical database [33]. The phenomena of different solvent schemes on the morphology and size of nanofibers have been examined [34–39]. The single solvent system contained of acetone, chloroform, N, N-dimethylformamide, dichloromethane, formic acid, and methanol (MeOH) and pyridine. Acetone-dimethylacetamide (DMAC), chloroform-MeOH, and DCM–MeOH were amongst the combined solvent systems. The deformation viscosities, surface tension, and conductibility of these solvent systems have been assessed to be vital solution arguments in producing smooth nanofibers. However, three solvent system of acetone/DMF/trifluoro-ethanol has also been assessed for e-spinning CA [40]. The property version of a mixed solvent system of acetic acid/water has been determined to prescribe the size allocation of the electrospinning of cellulose acetate [41]. CA based electrospun fibers in existence of poly(ethylene glycol), poly(ethylene oxide) and hydroxyapatite have been evaluated applying DMAC/acetone, DMF, DMF/dioxane and acetic acid/acetone systems [42]. An important remark was sent on by Haas and co-workers concerning the function of solvent excitableness in restraining the appeared coordinate of cellulose acetate fiber [33]. The Hansen's method of solvability tells us to choose atomic binary systems, improvement for production of cellulose e-spun nanofibers systems with adaptable level of fibers merger deserves limited remark in the circumstance of eco-friendly nanotech. Good packaging property was acquired with two less volatile mixture solvent. Ribbon like structure was observed for cellulose nanofibers produced from binary solvent system and achieved cylindrical shape [43]. Few studies are acquirable on the consequence of close arguments and e-spinning arguments on the quality of CA e-spun nanofibers. Humidity has influence on size of CA fibers, average size of diameter increased by increasing humidity. Besides, temperature
prescribes the solvent drying up rate and viciousness of the mixture. Humidity and temperature were discovered to have fundamental effect on the size of cellulose acetate nanofibers [44]. The phenomenon of various factors considering field strength, space between tip to collector, feed rate of solution and property on the structure of CA e-spin nanofibers are previously reported [45]. Some determined parameters can be modified by using Box-Behnken pattern method for CA electropsun fibers [30]. Structure and fibers diameter were determined to be affected by High power, space between needle tip to collector and feed rate of the polymer solution made in acetic acid/water mixture solvent. Concern for CA electrospinning fibers as blends and pure has been increasing in last decay. CA has been blend electorspun with natural polymers and nanomaterials such as AgNPs etc. Studying to cellulose, cellulose CA has better dissolvableness therefore, different solvents system are available [30, 46, 47].

2.2 Electrospun cellulose acetate nanofibers with AgNPs for antimicrobial activity

The destructive effects of antimicrobial are already manifesting themselves across the globe. Several disinfectant substances have been formulated for healing and keeping illnesses in public health medicine and antigen in biomedical diligence [11, 12, 67, 68]. A very important aspect of nanomaterials action on bacterial cells, which must be considered when discussing the inhibitory effect of it tested on different types of microorganisms. Bacteria are commonly classified into Gram positive and Gram negative, depending on their cell walls. Gram positive bacteria have an extra thick peptidoglycan layer on the outer surface, while Gram negative bacteria have an outer membrane behind which a thin peptidoglycan layer exists. This drastic difference in the nature of the cell boundaries is a great challenge for having a general antibacterial material against the bacteria [69]. Biological active substances, considering sutures, implants (such as vascular prostheses, prosthetic heart valves, ureteral stents, and hernia meshes), and composites, gestate the adventure of infections at operative position, those are basic interest of hospital borrowed infections. The synthesis of nanofibers containing metal NPs is well explored due to vantages required with mixing the functional attributes of metal nanoparticles with the broadly relevant ownership of nano size fibers. Metal carrying e-spin nano size fibers have tugged concern as a new shape of antimicrobial matter. Ag is a broadly employed and distinguished wide spectrum antimicrobial substance which is good versus bacteria, fungi, and viruses but is nontoxic to mammalian cell. In addition, sequence of the high level, particular overhead side and thinness of e-spin nano size fibers with the antimicrobial effect of metal NPs outcomes in an excellent and various disinfectant substance. Materials based on cellulose like CA are a particularly good choice for preparing these materials, due to their excellent performance characteristics; CA, for example, exhibits good toughness, high biocompatibility, and relatively low cost. Moreover, the cellulose explores demonstrates, a predicting promising biological substance for tissue engineering, stem cell research, and regenerative medicine [70]. Cellulose produced by bacteria is merely examined for bone reclamation [71–73]. Therefore, it doesn't provide the quality to assure the fibers on the nano and micro size, it confines applicability of cellulose nano or micro size fibers for tissue engineering. CA is easy to electrospin. Besides, CA nanofibers can be easily deacetylated to cellulose by alkaline hydrolysis [74]. Furthermore, a number of antimicrobial agents or antibiotics can be comprised of nanofibers for the prevention of microbial infection [75, 76].

In this part, we will briefly discuss the effect of cellulose nanofibers with AgNPs on antimicrobials. Cellulose acetate nanocomposite was determined to have able of inhibiting the growth of infective micro-organism [77, 78]. Numbers of reports are available on cellulosic based material [34–36, 38, 41, 49–52, 79–85]. Antimicrobial agents of cellulose acetate such as AgNPs and ZnNPs have been reported recently [30]. It has been reported that the synthesis of e-spin CANFs surface coated with AgNPs [86]. Apart, interceded in place chemical reaction of metal salts for the synthesis of NPs coated cellulose acetate nanofibers [87, 88]. Before elevating CA/AgNPs, it would be fruitful to discuss basic of silver nanoparticles. Within many types of bactericide stuffs, investigators are progressively becoming to nano-size materials because of specific sequence of physio-chemical concepts. Those nano-size materials have been studied in different areas, like drug delivery and imaging [89–95]. Specifically, investigators have studied the attributes of nanomaterials like Ag, Au, Zn, and Cu [96–99]. From these nanomaterials, AgNPs have evidenced fantabulous bactericide effect [100]. Moreover, toxicity mechanism of AgNPs has been previously studied. Up to date the report is available on biomedical application of AgNPs [101]. Incorporation of silver nanoparticles cellulose substratum to carry out bactericide effect has also been recognized [102].

Furthermore, the high dose of AgNPs also elicits wellness and binomical involvements. The termination of AgNPs in H2O creates the stuff to easy break up into ions few days, and toxic conditions for cell enhance with the
amount of Ag⁺ in the distribution. Some study demonstrated AgNPs are supposed of exposing harm for mammalian cells [96, 103, 104]. Still, although there being many infomercial products are available which flick AgNPs to control bacteria to handle cuts, scrapes, and burns, it is significant to formulate novel stuff which exhibit less perniciousness and more biocompatibility to develop bactericide profile which should be secure for mammalian cell and the ecosystem. Moreover, Ag also has been studied with Au for improving its properties such as antimicrobial effect and non toxicity for mammalian cell. Though Ag@AuNPs core and shell nanoparticles have been synthesized this can show better results for biomedical applications. However, Only AgNPs are depicted harmful for mammalian cell in high dose; combination of noble metals would be a better choice for biomedical applications such as Ag@Au and Au@Pt [105, 106].

Additionally, Silver nanoparticles decorated in polymeric substrata can empower the complexes with convenient optical attributes and increased bactericide effects [107–110]. That stuff has been employed in broad biomedical applications [111–113]. Due to bactericide action of Ag-based materials have been demonstrated powerfully believe in their property such as size and shape [114]. Cellulose based material is more attractive for Ag-Based nanocomposite due to its surface functional groups (OH) which is electron rich characteristic. Though, it has effective colloidal steadiness in aqueous solution. Incorporation of nanoparticles can be restrained by the H-bond system that is organized by the hydroxyl groups in the morphology of cellulose. Aforementioned, as compared with cellulose, cellulose acetate can be easily electrospun, it can be regenerated as cellulose nanofibers by alkaline hydrolysis. Recently cellulose acetate is used for electrospun and then AgNPs incorporated by chemical reduction method. Cellulosic NFs were produced by alkaline hydrolysis from cellulose acetate nanofibers. Cellulose acetate nanofibers were synthesized by e-spun method, various concentration of CA solution was synthesized in solution of 2:1.5 N,N-dimethylacetamide DMAc and acetone (v/v) and cellulose acetate and cellulose nanofibers structure were found smooth at 25% solution of cellulose acetate Figure 2A [50, 51]. Figure 2B is showing cellulose regenerated electrospun nanofibers. After deacetylation, there was a little change in size of nanofibers. Regenerated cellulose nanofibers were decorated by AgNPs this can be seen in Figure 2C. Cellulose nanofibers with AgNPs have bactericide effects and the physiochemical properties of AgNPs have an impression on antibacterial activity Figure 5B. Moreover, AgNPs with excessive amount has better results than lower concentration. Cellulose acetate is definitely a good choice for electrospun nanofibers for biomedical applications by combining with nanomaterials. Electrospun cellulose acetate nanofibers in combination with Ag@Au core and shell could be a good choice for biomedical applications. Recently different size of Ag@Au NPs core shell decorated on cellulose paper has been prepared by Tsai et al for antibacterial activity. It was postulated that Au NPs works as core and AgNPs as shell [115]. Cellulose paper directly has been directly used for Au@AgNPs decoration Figure 3. It is presented that Au-Ag100/1 and Au-Ag1000/1 NP-coated cellulose paper can inhibit E. coli, with 15 nm Au-Ag100/1 NP demonstrated the firmest antibacterial activity. Figure 4 showing antibacterial effect of different size Au@AuNPs/cellulose. Moreover, it can be suggested that the silver coating is creditworthy for the bactericide activity, nearly because of the action of silver ions from the coat of the Au@AgNPs.

### 3 Silver nanoparticles (AgNPs) properties

Silver nanoparticles breaking consequence on the bacterial membrane could be powerfully associated with

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**Figure 2**: SEM Images of a) Cellulose acetate nanofibers b) Regenerated cellulose nanofibers c) AgNPs incorporated cellulose nanofibers [51]
their physical and chemical properties, such as size and shape [116, 117]. The primary requirement for AgNPs to cause the membrane impairment is the capability to act with the bacterial membrane. Size of the nanoparticles is significant expressions in influencing their action with membranes. Small size NPs would amend interaction with the bacterial cells as studied to the same dose of their bigger size NPs. 1-10 nm in diameter were determined to be the most efficient in inhibiting to the bacterial cell due to their discriminatory directly action with the cell membrane [118]. Additionally, the smallest AgNPs demonstrated the most eminent antimicrobial effeteness and rapid bacterial inhibiting activity [119]. Electrospun Cellulose acetate nanofibers decorated with different size of AgNPs were studied. Figure 5a showing TEM images of AgNPs decorated cellulose nanofibers and inhibition effete of different size AgNPs on E. coli can be seen in Figure 5b. 14 mm and 11.9 mm zone inhibition activity has been found from two different sizes AgNPs.

Polyvinylpyrrolidone decorated silver nanoparticles with unlike forms proposes a stats between the shape of AgNPs for antimicrobial properties. Two dimensional Ag nanoplates possess higher germicide toward S. aureus and E. coli when examined to one dimensional Ag nanorods and round AgNPs zero dimensional morphology. It was claimed that Ag nanoplates contain the larger surface area, which could provide higher interaction with bacterial cell [120]. Moreover, the shape of nanoparticles also can affect Ag atoms dispersion [121]. Silver ions are responsible for killing or inhibiting microbes. The oxidation states of AgNPs assuming when desig of antimicrobial AgNPs. It was less toxic towards E. coli as compared to the Ag⁺ ions in an aerobic place. The lack of Ag oxidation was responsible for less antimicrobial activity, suggesting the value of silver ion in the antimicrobic activity of Ag-
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Figure 5: A) demonstrating TEM images of a) regenerated cellulose nanofibers b&c), regenerated cellulose nanofibers with AgNPs decoration (42.33 nm and 16.69 nm); B) demonstrating antibacterial activity of AgNPs against E. coli d) regenerated cellulose nanofibers e&f) regenerated cellulose nanofibers with AgNPs decoration (42.33 nm and 16.69 nm) [50]

NPs [122]. Partially oxidized AgNPs can have somehow antimicrobial effect, but zero valent AgNPs may not. Aforementioned, suggesting the quantity of chemical-sorbed silver ion on the cover of silver nanoparticles would influence the maintained antimicrobial activity [123]. Silver ions possible inhibition mechanism for microbes are Ag⁺ controls Adenosine triphosphate (ATP) production through attaching to the ATP synthesis enzyme in the cell wall, it enters into the cell wall and attaches to DNA which leads to the DNA alteration or these ions with the nucleus of the cell.

4 Conclusions

For the application of cellulose acetate nanofibers, electrospinning technique has presented a completely modern orientation. CA has been considered for electrospun nanofibers because of decomposable, biocompatible, and not soluble in water. In addition, the selection of solvent, decently mix solution factors and settle their ratio which effectively reserves the attributes of nanofibers synthesized by e-spinning technique. Moreover, for the cellulose acetate, acetone/dimethylacetamide found better solvent during electrospinning. Aforementioned, CA nanofibers can be possibly employed for antimicrobial activity and wound dressing. However, cellulose acetate nanofibers can easily be deacetylated by alkaline hydrolysis. Cellulose nanofibers were decorated with different concentration and size of AgNPs for antimicrobial effect. A nanoparticle with smaller size possesses higher antimicrobial activity because it can directly interact with the cell. Combination of metal with electrospun cellulose acetate nanofibers could make a great contribution to biomedical application. Au@Ag bimetallic nonmaterial with electrospun nanofibers could be used for biomedical applications.

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