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Impact of Bionanopolymers in Wound Healing Materials and Their Mechanisms

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Abstract: Many sorts of injuries were dealt with by focusing on different parts of the cure process with the technological development of over 3000 items. This review recounts the history of dressings from its very beginnings to the present state, as well as the benefits and limits of dressing materials. Nanomaterial wound therapy offers enormous potential to cure and prevent injury infections with a variety of advantages compared to standard methods of treatment. In this connection, researchers may perform comprehensive investigations of wound healing applications using the physiochemical characteristics of nanomaterials. However, concerns related to the use of nanomaterials to speed up the effectiveness of current therapies remain unsolved. The current study emphasises new approaches to recent and detailed revolutionary wound-healing techniques and infection controls employing nanomaterials, nanoparticles, nano-composites and scaffolds. The effectiveness of nanoparticles as transporters of medicinal substances related to wound-healing applications was also investigated. Finally, nanomaterial scaffolds and their premises were outlined for further investigations. We think that the detailed analytical analysis, future insights and possible issues outlined in this review will give researchers with up-to-date reference to the usage and creative ways of nanomedicine that may help improve the application of wound healing.

Keywords: Wound healing, Nanocomposite, Nanoparticles, Dressing material

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Introduction

Wound disruption refers to the disruption of the epithelial skin or mucosal linings as a result of physical or thermal injury. Depending upon the length and nature of the healing method, the wound is categorised as acute and chronic. An acute injury is abrupt harm to the skin or surgery.

The treatment usually heals in the prescribed and predicted 8-12 weeks, depending on the size, depth and degree of damage to the skin's epidermal and dermal layers (Kamal *et al.*, 1986). Chronic injuries, on the other hand, fail, and cannot be repaired properly and promptly in the

normal healing phases. Chronic injury generally results from ulcer of the decubitis, leg ulcers and burns (Iliuk *et al.*, 2010). Healing wound is a dynamic and complicated tissue regeneration and development process in separate phases-- (i) clotting and haemostasis (immediately after injury); and (ii) inflammatory (just after damage to the tissue) during which the swelling occurs. These stages occur in a well-connected cascade to overlap (Konwarh *et al.*, 2010). The progression of these stages is largely determined by the nature and pathological conditions associated with the wound, as well as the type of dressing used. With technical advancement, various types of wound dressing material for all types of wounds are now conceivable (Singh *et al.*, 2011). It is nonetheless crucial for a particular wound to choose a chemical to encourage faster cures. This review has reinforced the many kinds of wound dressing materials and their function in the treatment process (Sahoo *et al.*, 2011).

Factors impacting the treatment of wounds:

Cytokine promotes healing via a number of methods, including promotion of the production of components of the cell membranes, avoidance of dehydration, enhanced inflammation and growth of granular tissues (Madaan *et al.*, 2013). Various local and systemic variables impact these pathways. The feature of the wound is directly influenced by local conditions that include hypothermia, pain, infection, radiations, and tissue oxygen pressure, in which the general health or sickness of the individual affects the capacity for the individual to cure. Furthermore, bad diet, age, protein, vitamins and shortage of minerals can further delay healing periods (Chaudhury *et al.*, 2014).

Features of a good dressing for wound:

Depending on the type of wound the appropriate dressing material should be used. Select a dressing to enhance epidermal migration should be found on its ability (a) stimulate angiogenesis and synthesis of connective tissue, (b) facilitate tissue-to-environmental exchange of gas, (c) maintain a

sufficient tissue temperature to promote flow into the wound bed and epidermal migration, (d) Bacterial and non-wound protection, (e) ready detachable after therapy, (f) using leukocyte migration and enzyme build-up debridements, and (g) must be not-toxic, non-allergic and sterile (Picheth *et al.*, 2014).

Wound Dressings:

Wound, whether it is a little cut or a large incision, careful care is necessary. Wound treatment is part of the operation. Dressing was designed to come into touch with the wound other than the dressing bandage. The wet to dry dressing has historically been frequently utilised in cases of wounds that require debridging. Wounds were covered with strips of linen soaked in oil or plaster-coated grease about 1600 BC. The treatment of mesopotamian wounds was based on clay tablets from approximately 2500BCE (Jayasundera *et al.*, 2014). Wounds were cleansed by water or milk before to treatment with honey or resin. In the afternoon, wine and vinegar wounds were used for the treatment with sweet honey, olive oil and wine, followed by ancient Greek Hippocrates about 460–370 B.C. (Karak *et al.*, 2015). The wool was boiled in water or wine as a bandage. In the 19th century antibiotics were a major advance in controlling and minimising illnesses. The Modern wound wardrobe (Nasrollahzadeh *et al.*, 2015) arrives in the 20th century.

If the wound is closed down with dressing, it continually exposes proteinases, chemotactics, complements and growth factors lost in open wound. At the beginning of the 20th century, then, the production of occlusive clothing began to shield the wound and provide water for the aquatic environment (Amoozgar *et al.*, 2016). This garment helps to swiftly re-epithelise, synthesise collagens, stimulate angiogenesis to the injured bed via hypoxia and alter the pH of the wound bed to decrease injury. The tissue was used as cotton gases in 1891. By the half of the 1900s wounds were thought to be healed faster while they remained dry and uncuts, although closed wounds were more rapidly treated than open ones (Das

and Baker, 2016). The first current wound dressing with substantial moisture and fluid absorption characteristics was launched around the mid-1980s (e.g. polyurethane foams, hydrocolloids, iodine-containing gels). In the mid-90's, a range of synthetic wound dressings developed, including hydrogen cups, hydrocolloids, Alginates, synthetic foam dressing, silicone meshes, adhesives, silver/collage dressing sheets (Leonida and Kumar, 2016).

Traditional clothing for wound:

Traditional materials of dressing such as gauze, lint, plasters, bandages of natural or synthetic materials and cotton wool have been dried and used as primary and secondary dressings of the wound to guard against contamination. Gaze dressings consisting of woven and unwoven cotton fibres, rays and polyesters are a kind of protection for germs. Some sterile gas pads are used for the absorption of exudates and fluids in the Open Wound using fibres (Aized *et al.*, 2017) in these dressings. These clothing should regularly be altered to prevent maceration of healthy tissue. Cheaper are gaze treatments. The dressings are moisturised by the abundant drainage of the wound and tend to adhere to the premises, making the wound uncomfortable. Cellulose or synthetic bandages, made of natural cotton and polyamide materials, serve diverse uses. Cotton bandages for the preservation of light-dressing, for high compression bands and for short-stretch compressible packs may be used to venous ulcers (Allizond *et al.*, 2017). The Bismuth Tribromophenate petrolatum gauze Xeroform™ (non-occlusive dressing), is designed to avoid the exudation into the lean exuding wound. Tulle clothing like Bactigras, Paratulle and Jeloneti are examples of the marketable tulle wear that is helpful for superficial clean wounds as paraffin-impregnated clothes. Conventional dressings are typically suited for clean and dry wounds with low exudate levels or for use as secondary dressings. Since traditional clothing does not provide the damp surroundings for the wound, they have been

replaced with current clothing and better formulations (Gogoi *et al.*, 2017).

Modern dressing of wound:

In addition to covering the wound, modern wound clothing was intended to enable it to operate. These clothing systems are intended for preventing and facilitating dehydration of the wound. On the market, different solutions based on the cause and kind of damage are available and make the choice a very difficult procedure. Mainly synthetic polymers are modern wound dressings, classified as passive, interactive and bioactive (Pinho and Soares, 2018). The non-occlusive use of passive products such as gazettes and tulle dressings covers the wound in order to regain its function. Semi-occlusive or occlusive are interactive dressings in films, foam, hydrogel and hydrocolloids. This is a barrier to microorganisms that penetrate the harmful environment (Saravanakumar *et al.*, 2018).

Dressings for bioactive wound:

The fourth kind of modern wound dressing treatment is biomaterials which play a vital role in the healing process. It is known to be biocompatible, biodegradable and non-toxic, and produced from natural tissues and artificial sources such as collagen, hyaluronic acid, chitosan, alginate and elastin (Shaghaleh *et al.*, 2018). Polymers of such materials shall depend on or in combination on the sort and nature of the wound. Organic dressing with growth factors and non-bacterial items is sometimes incorporated in the process of wound healing (Abdelraof *et al.*, 2019). Numerous investigators have examined collagen, an important structural protein that plays an active role in the natural healing process. The collagen stimulating the production of fibroblasts and accelerating endothel migration when it comes to touch with injured tissue. A glycoaminoglycaine component of the Extra-Cell Matrix (CEM) (Donnarumma *et al.*, 2019) is hyaluronic acid (HA) with its unique biological and physicochemical properties. HA is biocompatible, biodegradable and naturally lacks collagen-like

immunogenicity. Chitosan promotes the formation of granulated tissue during the proliferation stage of wound healing. In contrast with other dressings (Fasiku *et al.*, 2019), biological dressings are regarded superior to other dressings.

Dressings medicated:

A major component of the therapeutic procedure is in medical dressings which incorporate directly or indirectly medications by removing necrotic tissues. Necrotic tissue may be used, antibacterial products that prevent infection and encourage tissue regeneration, cleansing or decomposition. Typically, some compounds include antibacterial agents, growth hormones and enzymes (Fasiku *et al.*, 2019). Cutisorb™ for antimicrobial dressings is available commercially. The silver impregnated dressings are available as fibrous hydrocolloid, polyurethane films and silicone gels. Antiseptic dressing of iodine acts on the bacterial cells via oxidative cell disintegration, interfering with the extensively harmful protein activity. Long usage of iodine leads to skin inflammation. In specific diabetic foot ulcer (Gnanasekaran, 2019) antimicrobial medicines are primarily intended as a prevention or control for infections.

The normal tissue healing process in the body is governed by cellular activity as our organisms expand organically. Cell and growth factors inside the injury cells are retained in the event of chronic wounds which have an effect on the treatment process (Ghaem *et al.*, 2020). This enhances the wound-healing process via exogenous injection of growth factors. The PDGF is the most often used growth factor in chemical recruitment, cell proliferation and angiogenesis, amongst the other growth factors (Sadrmanesh, 2020). The usage of PDGF, Fibroblast Growth Factor (FGF), Epidermal Growth Factor and Autologous Thrombin Platelet is also widely investigated for the healing process (Naskar and Kim, 2020). The FDA authorised PDGF and EGF for human use. Necrotic tissue breakdown is also necessary for the natural healing process without harming healthy tissue. Necrotic tissue is now processed using papain and ointment based on collagenase. Collagenase acts

against collagen by targeted native collagen and by the slow disintegration of tissue into a viable collagen (Lin *et al.*, 2020).

Transparency:

The ulcer surface can be checked visually using a transparent dressing. An ulcer with an untransparent dressing might be progressively infected without being noticed. It is important to remove and alter clothing regularly when wearing non-transparent clothing (Ahari and Lahijani, 2021).

Composite dressing:

Composite dressings are flexible and practical for both partial and whole thickness wounds. There are numerous complex layers and each layer is physiologically distinct. Most compound dressings include three layers. In addition, composite dressings may have a nonwoven fabric tape adhesive or clear film edge (Elnagar *et al.*, 2021). It can function as a main or secondary garment against a range of wounds and can be used with topical drugs. The wound is typically composed of absorbent material, moisture-preserving and autolytic debridging; the middle layer outside the most strata protects the wound from infection, and the bottom layer, which is not adhesive to material which prevents attachment to young grained tissues. Composite dressings are cheaper and less flexible (Moreno *et al.*, 2021).

Although over seven million Americans have been chronically injured every year, modern research have produced more than a handful of therapy products that the FDA deems to be effective. In view of the intricacy of the chronic disease and rehabilitation of the tissue, it is not unexpected that there has not been adequate proof of existing therapeutic techniques to get FDA approval, which increased the demand for viable and efficient alternative therapeutic treatment methods (Mahesh *et al.*, 2021). Many contributors prevent more rapid therapy for chronic lesion patients, including a limited understanding of the pathophysiology of the patients; variability and comorbidity of the population of the patient

(patient-to-patient) and of a group variability (diabetic foot, vein legs, pressure ulcers) (Ul-Islam *et al.*, 2021). The fundamental challenge is the design of appropriate combination therapies that can target different cell failure pathways and select the perfect delivery time for each therapeutic medicine. Currently, however, we are in the midst of BD2 endeavor K's to revolutionise computer biomedicine (Ojha and Das, 2021). In B D2K programmes and institutions, tools and 'ecosystems' are being developed rapidly to integrate huge information sets, including medical records, genomes, genotyping of drug customization or available to the public on drug response (Pérez-Moreno *et al.*, 2021). A look at how tailor-made therapies are based on the genotype's unique characteristics presents major possibilities for adapting traditional nanotechnology techniques and for simplifying and facilitating personalised treatment plans resulting from these platforms (Kramerov *et al.*, 2021).

The huge number of articles that have been presented in the past several years that imply high hopes of nanotherapy interventions is proven in wound cure in the development of various nanotechnologies, in particular multifunctional systems. However, the barrier is acquiring knowledge about physicochemical properties of human body nanoscale systems, and their anticipated behaviour and toxicity (Tayel *et al.*, 2021). The high clarity of the fabric and nanoparticles recommended by the FDA for human use is also very challenging since bulk polymers and nanoparticles often cannot be easily produced and purified. There is also a continuous demand for enhanced synthetic tools and techniques to analyse the clinic which enable the translation of nanotechnology technologies. Comprehensive efforts are also necessary for the arming and efficiency of site-specific chronic wound therapies, to minimise undesirable accidents and interferences which might damage the biological activities of the nanosystems in the body (Wang *et al.*, 2021). Further study is needed in the long term to provide insights on how

nanotechnology-based results of research in the therapeutic arena may be used. Nanotechnology platforms are expected to develop and additional research is needed on these technologies to create global biocompatibility standards and toxicity for nanotherapy. The overall objective of the system is to assist the design and development, along with knowledge of chronic injuries, molecular pathology and phenotypic genotype characteristics of the future generation of wound healing nanotechnologies in current capacities (Zhu *et al.*, 2021).

Conclusion

The bandages market presently includes more than 3000 types which the doctor may give in all fields of injury treatment. However, persistent wounds such as venous leg ulcers, diabetic injuries and pressure ulcers are not yet cured in the perfect way. However, they generally are not fully cured. The design of clothing address the primary interfering factors that can greatly assist patients and injurers. The focus of this review was on the advantages of wound-healing nanomaterial. Nanoparticles' unique physiochemical properties make them the ideal choice for the wound healing process. It was also better than standard treatment of wounds, focused mostly on clothing, and on wound healing from nanomaterials. Nanomaterials can modify one or more stages of the wound treatment since the wound healing process contains antibacterial, anti-inflammatory and anti-proliferative features. In clinical wound therapy, however, some major obstacles have to be solved with regard to the successful use of nanomaterials. The possible detrimental effects in human cells are one of the main concerns regarding the toxicity of nanomaterials. This has to be resolved at the beginning for more advanced wound therapy or clinical studies. Another problem is that the non-material wound healing applicability of the in vivo model is rarely understood. Many of the studies reported for wound healing applications based on nanomaterial mostly depends on one-target bacterium or is in vitro. For in vivo wound healing

uses, gramme positive and negative bacterial strains and skin microbial strains, more comprehensive study are thus necessary. In future attempts will be made to improve the specificity and effectiveness of the site by applying wound healing more effectively in line with the nanomaterial utilised. The researchers must thus produce biocompatible, biodegradable nanomaterials capable of curing all healing wound stages.

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