



Nanotechnology-based approaches for acne treatment: A comprehensive review

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Abstract

Acne vulgaris is a prevalent dermatological condition affecting millions worldwide, causing significant physical and emotional distress. Nanotechnology has emerged as a promising approach in the treatment and management of acne, offering improved drug delivery, enhanced efficacy, and reduced side effects. This review aims to provide a comprehensive overview of the application of nanotechnology in acne treatment, focusing on pathophysiology, clinical presentation, and management strategies. A detailed examination of various nanoparticle types utilized in anti-acne treatment, including liposomes, niosomes, solid lipid nanoparticles, and metal-based nanoparticles. The benefits and challenges of using nanotechnology in acne treatment are also explored, including concerns about toxicity and safety. Additionally, the regulatory framework surrounding nanotechnology in the European Union. This review highlights the potential of nanotechnology to revolutionize acne treatment and management, providing a promising avenue for future research and development.

Keywords: Acne vulgaris; Nanotechnology; Nanoparticles; Acne treatment

1. Introduction

Acne vulgaris is one of the most common dermatologic conditions globally [01]. The skin condition known as acne vulgaris affects the oil glands located at the base of hair follicles [02]. Development of acne may be individual or site-specific, with only a few follicles presenting the inflamed area yet at some stage in serious disease manifestations [03]. Acne is the most prevalent skin disease in the world, affecting 85% of adolescents and over 10% of adults [04]. Acne is associated with increased risks of stigmatization, bullying, depression, anxiety, poor self-esteem, and suicidal ideation [05].

Considered the most promising technology of the twenty-first century, nanotechnology is expected to have a significant positive impact on the cosmetics sector. The words "technology" and the Greek numeral "nano," which meaning dwarf, are combined to form the term "nanotechnology." Therefore, science and technology utilised to create or work with particles that range in size from 1 to 100 nm is known as nanotechnology [06]. Nanocarriers, including liposomes, niosomes, solid lipid nanoparticles, and micro-emulsions, provide protection for the encapsulated hydrophilic and lipophilic actives, target and control delivery while minimizing their toxicity toward other tissues, and allow higher skin penetration and time residence [07]. Acne is an inflammatory disease of pilosebaceous units in the skin of the face, neck, chest and upper back [08]. As nanomedicine technology continues to advance, nanotechnology has gradually emerged as a research hotspot in the field of drug delivery. Using nanotechnology, drugs may be synthesized into tiny particles to increase their water solubility [09]. Nanocarrier based transdermal drug delivery systems offer multiple advantages over conventional modes, such as enhanced physicochemical stability of drugs, increased skin permeation, improved biodistribution, effective targeted accumulation, and controlled drug delivery. Especially, nanocarriers can be applied

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in the hair follicle (HF)-targeted delivery to enhance the topical treatment outcomes of some appendage-related skin diseases, such as alopecia and acne [10].

In figure 1 highlights the difference between a normal skin and the skin with acne. In the skin with acne due to the excess production of male hormone androgen and oil producing glands in the face comedone occurs on the face. The closed comedone (whitehead) and ripen comedone (blackhead) are the primary two noninflammatory lesions in acne. These lesions may progress to inflammatory papules and pustules when the contents rupture. Larger, more painful lesions, such as cysts and nodules, may also develop. The application of novel delivery systems to the skin distributes the topical agent gradually, reduces the irritancy of some antiacne drugs, and shows good efficacy [11].

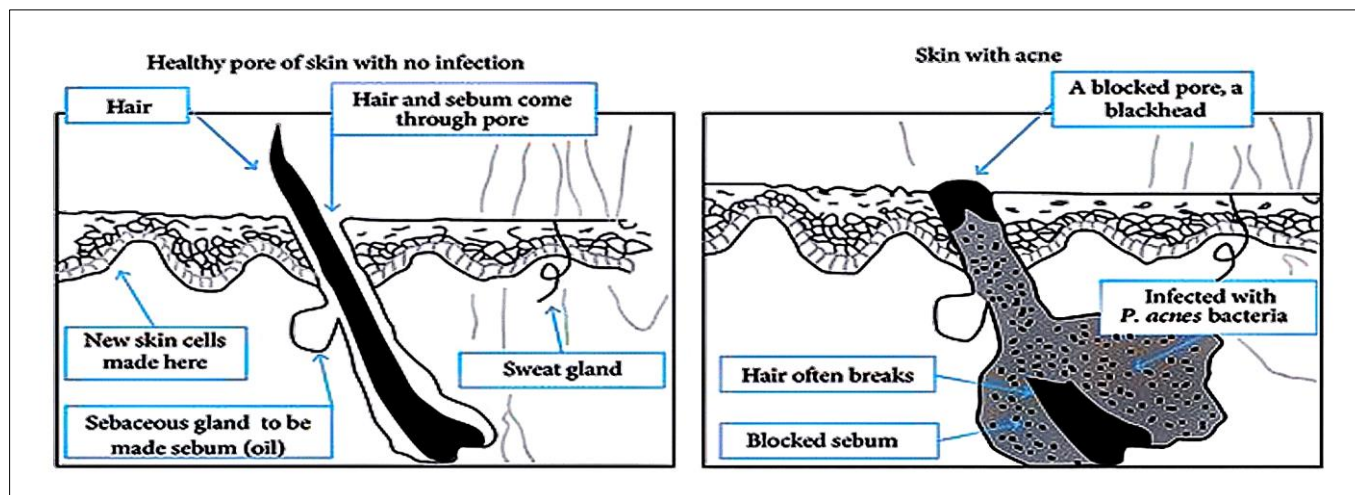


Figure 1 Difference between normal skin and skin with acne [11]

2. Causes of acne

Acne is mostly caused by a variety of reasons, including hormonal (Androgenic progestin), emotional, environmental (humidity, heat, etc.), and microorganisms.

One of the main reasons for acne forms is an increase in sebum production in the hair follicles [12].

Acne vulgaris is mainly caused by gram-positive, non-spore forming, micro-aerophilic and rod-like bacterium *P. acnes* [13].

Excessive colonization of these microbes promotes perifollicular inflammation. Therefore, increases in the development of Cutibacterium acne and inflammatory processes are additional causes of acne development [14].

One of many causes of acne vulgaris is the proliferation of Propionibacterium acnes (*P. acnes*), a normal anaerobic flora found in sebaceous glands [15].

3. Types of nanoparticles for Acne treatment

There have been several nanotechnology-based formulations that have been proved to be efficient for acne treatment and care. In this section, describe lipid nanoparticles polymeric-based nanosystems, metal-based nanoparticles and additional nanosystems (Figure 2) [16].

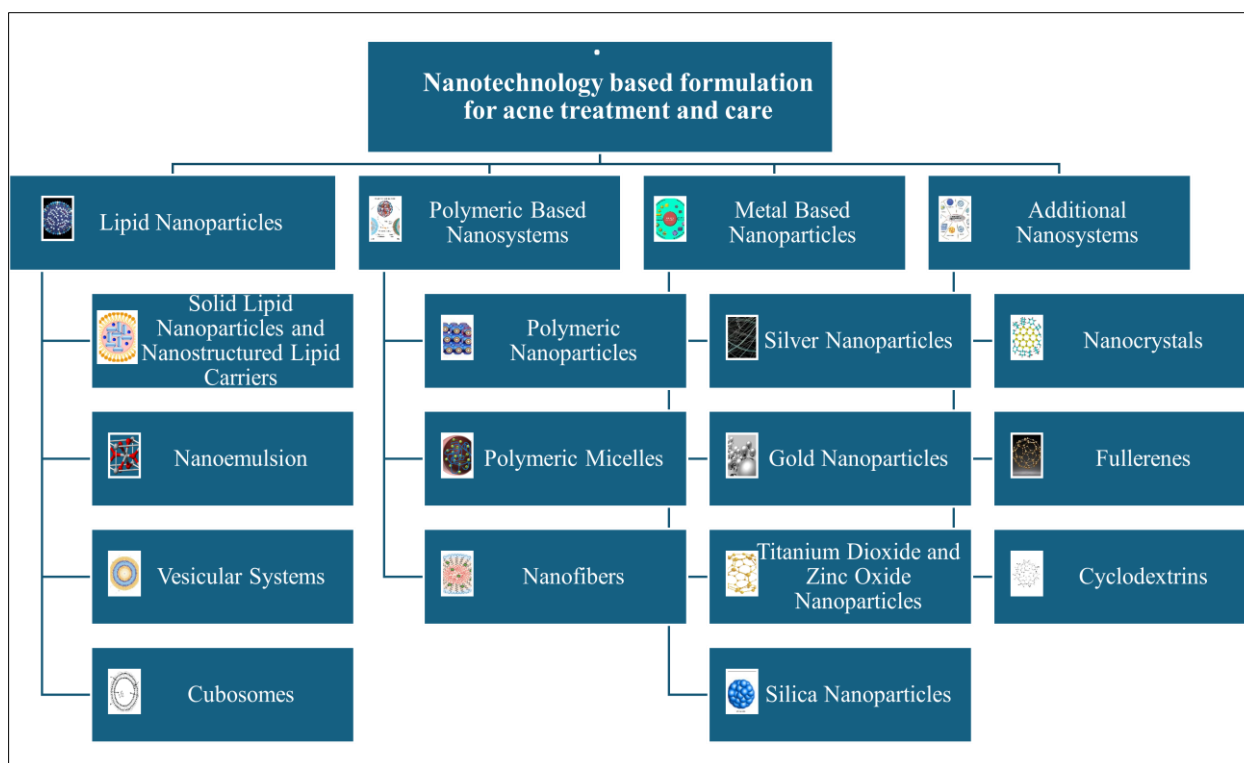


Figure 2 Schematic compilation of nanotechnology-based formulations for acne treatment and care [16]

3.1. Liposomes

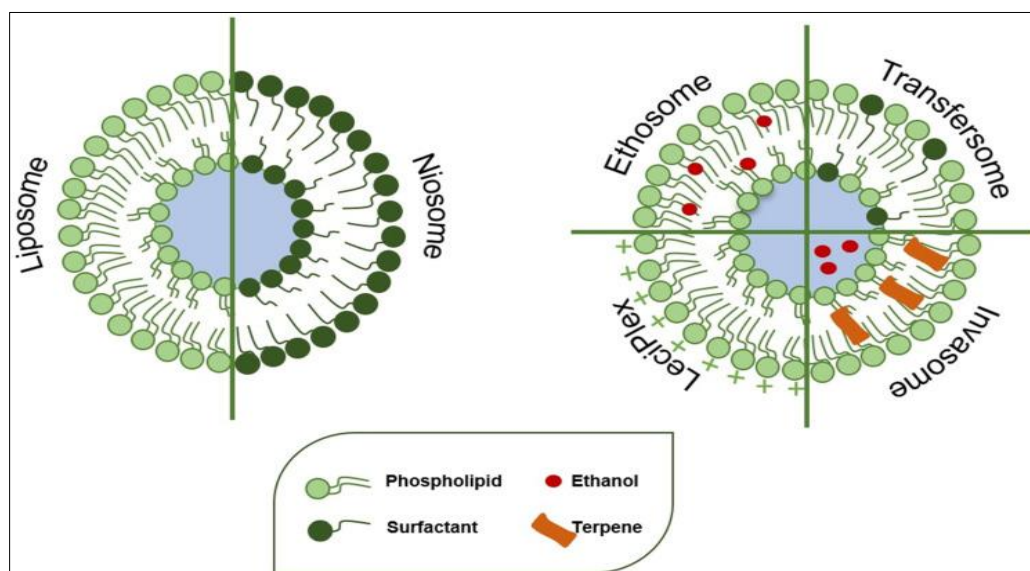


Figure 3 Schematic representation of liposomes and liposomal-like vesicles [18]

Liposomes can promote drug deposition in the skin at the site of action, decrease systemic absorption, and minimize adverse effects, all of which contribute to a localized effect of anti-acne medications. Additionally, they can boost the drug's systemic absorption by targeting it to skin appendages. They can lessen the negative effects and increase the therapeutic efficacy of medications [11]. Liposomes have found significant applications in dermal and transdermal drug delivery, including their targeted delivery to skin appendages. In the context of dermal drug delivery, liposomes are employed to enhance drug penetration into the skin ensuring localized therapeutic levels while minimizing percutaneous absorption [17]. Conventional liposomes are one of the most famous and extensively studied lipid vesicles, which are typically composed of phospholipids, cholesterol, and aqueous medium (water or buffer solution with varying pH). These vesicles are formed when naturally or synthetically occurring biodegradable lipids come into

contact with the aqueous medium, wherein the hydrophilic head group of the lipid surrounds the aqueous core while the hydrophobic tail group is exposed to the external medium. Due to this unique structural property, water-soluble drugs can be loaded in the aqueous core while the water-insoluble drugs can be loaded in the lipid bilayer [18].

3.2. Niosomes

Niosomes are vesicular drug delivery systems that have a long shelf life, a sustained drug release profile, and improved penetration of the stratum corneum or intradermal passageways [19]. One of the most promising drug delivery methods for the treatment of skin conditions is niosomes. When given topically, niosomes can decrease systemic absorption of the medicine while increasing its residence time in the stratum corneum and epidermis [11]. Microsponge delivery systems of (Benzoyl peroxide) BPO, niosomal gel formulations and solid lipid nanoparticles have been reported in the article [20] Niosomes are nanosized colloidal vesicular structures formed by the self-assembly of non-ionic surfactants in an aqueous environment resulting in a bilayer configuration [21].

3.3. Solid Lipid Nanoparticles (SLNs)

In the 1990s, SLNs were introduced as a viable alternative to traditional colloidal carriers, including liposomes and polymeric nanoparticles, to account for the potential for industrial-scale production [22]. In 2012, a study showed the improvement of the properties of SLNs, as a carrier, by adding chitosan to SLNs loaded with (Trusted Research Environment) TRE. It was found that low TRE concentrations in TRE loaded SLNs with chitosan inhibited acne-causing bacteria, such as *C. acnes* and *S. aureus*, with a MIC of 40 µg/ml and 300 µg/ml, respectively [16]. According to a published study, when compared to free medications for the treatment of psoriasis, SLNs could considerably improve the quantity and depth of cyclosporine A and calcipotriol skin penetration [23]. The liposome micelle structure is released by Solid Lipid Nanoparticles (SLN) and Nanostructured Lipid Carriers (NLC), which are safe for delivering and releasing active ingredients to the skin and stable in both lipophilic and hydrophilic environments [24].

4. Example of nanotechnology based on anti-acne product

4.1. Benzoyl peroxide (BPO)

A topical non-antibiotic disinfectant, is used to treat mild to moderate acne because of its antibacterial and disinfecting properties [12]. Benzoyl peroxide is a crucial treatment for mild to moderate acne. While it can be used alone for 6–8 weeks, topical antibiotics are frequently used in conjunction with it to lessen *P. acnes* species resistance and boost treatment effectiveness [25].

4.2. Salicylic acid

A beta hydroxyl molecule with bacteriostatic, fungistatic, and anti-inflammatory qualities is salicylic acid. When applied topically, it keeps pores clear and exfoliates the skin to lessen acne. Although salicylic acid is safe to use, it can occasionally irritate, itchy, and dry out the skin [12]. Salicylic acid is an over-the-counter topical comedolytic medication that comes in concentrations between 0.5% and 2% [05].

5. Therapeutic Agent for Acne Treatment: Antibacterial, Anti-inflammatory, and Comedolytic Properties

5.1. Antibacterial Activity

The health and medical fields have conducted the most extensive research and development of technology that utilises the antibacterial properties of nanoparticles [26]. Zinc lowers inflammation and has antibacterial qualities against candida acnes [27]. Benzoyl peroxide is used to treat mild to moderate acne since it is an antibacterial and topical non-antibiotic disinfectant [12]. Furthermore, tretinoin-containing SLNs with and without chitosan were made to evaluate their antibacterial activity against microorganisms linked to acne and cytotoxicity in keratinocytes [17].

5.2. Anti-inflammatory activity

An active version of vitamin B3, nicotinamide is being studied more and more in the field of dermatology. This nutraceutical is a great option for treating acne because it has anti-inflammatory qualities and has been demonstrated to lower sebum production [19]. Willow bark extract contains salicin, an alcoholic β-glucoside with anti-inflammatory and high comedolytic properties that are used to treat skin conditions including acne [22].

5.3. Comedolytic activity

Benzoyl peroxide (BPO) is an anti-bacterial topical medication that kills *P. acnes* by the generation of free oxygen radicals and is also weakly keratolytic and comedolytic action [28]. Physical excision of microcysts, macrocomedones or closed comedones will boost the therapeutic efficacy of topically applied comedolytic drugs [25].

6. Challenges and future direction

6.1. Concerns about toxicity and safety

The short and long-term health impacts of nanoparticles on organisms and the environment are little understood, despite their immense benefits. Because of the nanomaterials' documented toxicity and potential risks, safety concerns have been raised [06]. Clarifying the long-term health advantages and possible negative consequences should be the main goal of future research. The physicochemical, anti-inflammatory, immunomodulatory, antioxidant, and DNA repair capabilities of polyphenols primarily determine their effects on the skin and can be used to prevent a range of skin conditions [29].

The future challenges of nanotechnology-based preparations for the treatment as well as care of acne are as follows:

- The absence of systemic absorption
- The prevention of nano contamination
- Ensure human health and safety
- Stricter regulatory framework [16]

7. Conclusion

Nanotechnology has shown significant promise in the treatment and management of acne vulgaris, offering improved drug delivery, enhanced efficacy, and reduced side effects. The various types of nanoparticles used in anti-acne treatment, including liposomes, niosomes, solid lipid nanoparticles, and metal-based nanoparticles, have demonstrated potential in targeting the underlying causes of acne. However, concerns about toxicity and safety must be addressed through further research and development. The regulatory framework surrounding nanotechnology in the European Union provides a foundation for the safe and effective use of nanotechnology in treatment with acne. As research continues to advance, nanotechnology is likely to play an increasingly important role in the management of acne vulgaris, providing new and innovative treatment options for millions of people worldwide. Ultimately, the integration of nanotechnology into acne treatment has the potential to revolutionize the field, improving patient outcomes and quality of life.

Compliance with ethical standards

Disclosure of conflict of interest

No conflict of interest to be disclosed.

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