A Brief Review on the Current Trends in Microencapsulation

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Publication history: Received on 11th April; Revised on 14th May; Accepted on 17th May 2024

Article DOI: 10.5281/zenodo.11527372

Abstract: Microencapsulation has emerged as a pivotal technology in the field of novel drug delivery systems (NDDS), offering significant advantages in enhancing the efficacy, safety, and patient compliance of therapeutic agents. This cutting-edge technique involves encapsulating active pharmaceutical ingredients (APIs) within microscopic protective coatings, enabling controlled release, targeted delivery, and improved bioavailability. Microencapsulation has proven to be a versatile tool in addressing various pharmaceutical challenges, such as masking unpleasant tastes, protecting sensitive drugs from degradation, and modifying the absorption site of drugs. This comprehensive review aims to provide an in-depth understanding of microencapsulation, its principles, techniques, and diverse applications in the pharmaceutical industry. The article delves into the reasons for employing microencapsulation, including extended or sustained drug release, taste and odor masking, conversion of liquids into free-flowing powders, and stabilization of light, moisture, or oxygen-sensitive drugs. Additionally, the classification of microparticles based on their core and shell materials is discussed. Furthermore, the review explores the advantages and disadvantages of microencapsulation, highlighting its ability to protect encapsulated active agents, transform gases and liquids into solid particles, modify surface and colloidal properties, and alter the release profiles of drugs. Various microencapsulation techniques are extensively covered, including air suspension, coacervation phase separation, pan coating, spray drying and spray congealing, and solvent evaporation. The applications of microencapsulation span across diverse industries, such as pharmaceuticals, cosmetics, agrochemicals, and food. This review provides insights into the utilization of microencapsulated products in these sectors, emphasizing their role in enhancing product functionality, stability, and controlled release.

Keywords: Microencapsulation; Novel drug delivery systems; Controlled release; Targeted delivery; Sustained release; Taste masking; Bioavailability.

1. Introduction

Novel drug delivery systems (NDDS) have emerged as game-changing strategies in the pharmaceutical industry, aimed at improving the effectiveness, safety, and convenience of medication administration. These innovative systems are designed to maximize drug absorption, target specific sites within the body, and control the release of therapeutic substances, ultimately reducing adverse effects and enhancing therapeutic outcomes. [1-4] One of the most promising NDDS technologies is microencapsulation, a multidisciplinary approach that combines cutting-edge formulations, novel technologies, and innovative development strategies to achieve desired pharmacological effects safely and effectively. Microencapsulation involves encapsulating active pharmaceutical ingredients (APIs) within microscopic protective coatings, offering numerous advantages over traditional dosage forms. Compared to conventional formulations, microencapsulated drug delivery systems offer increased convenience and patient compliance, reduced toxicity, and enhanced efficacy. These systems often utilize macromolecules as drug carriers, enabling the delivery of treatments that were once considered unattainable. The field of pharmaceutical technology has witnessed significant growth and diversification in recent years, with controlled release techniques and the development of novel polymers playing a crucial role. [5-7]

The process of microencapsulation involves encapsulating active ingredients or core materials within a protective shell or coating. This coating can be composed of various natural or synthetic polymers, such as proteins, polysaccharides, or biodegradable polymers, chosen based on the desired release characteristics and compatibility with the core material. [8-10] The microencapsulation process not only protects the encapsulated active agents from environmental factors like light, moisture, and oxygen but also allows for the transformation of gases and liquids into solid particles, facilitating handling and storage.
One of the key advantages of microencapsulation is the ability to modify the surface and colloidal properties of different active agents, enabling improved solubility, bioavailability, and targeted delivery. Additionally, microencapsulation provides a means to alter and control the release profiles of drugs from various pharmaceutical dosage forms, allowing for extended or sustained release formulations. This can lead to the development of prolonged controlled release dosage forms, ultimately improving patient compliance and therapeutic outcomes. Various techniques have been developed and employed for microencapsulation, each with its unique principles and advantages. These include air suspension, coacervation phase separation, pan coating, spray drying and spray congealing, and solvent evaporation. The choice of technique depends on factors such as the nature of the active ingredient, desired release profile, and production scale. The applications of microencapsulation span across diverse industries, with the pharmaceutical industry being a major beneficiary. In pharmaceuticals, microencapsulation has proven invaluable in addressing challenges such as taste masking, bioavailability enhancement, targeted delivery, and sustained release formulations. These advancements have led to improved patient compliance, reduced dosing frequency, and minimized adverse effects, ultimately enhancing therapeutic outcomes.

### 2. Microencapsulation

#### 2.1. Principle

Microencapsulation is a versatile technique that employs a multidisciplinary scientific approach to improve human health through controlled drug delivery. This section will delve into the fundamental principles and various techniques employed in microencapsulation. A microparticle is generally defined as any particle with a diameter ranging from 1 to 1000 μm, regardless of its interior or exterior structure. Within the broad category of microparticles, "microspheres" specifically refer to spherical microparticles, while "microcapsules" refer to microparticles with a core made of a material that differs significantly from the shell, which can be solid, liquid, or even gas. A microcapsule is a spherical particle with a core substance and a size ranging from 50 nm to 2 mm. It is important to note that the terms "microsphere" and "microcapsule" are sometimes used interchangeably, despite their distinct definitions. Other related terms, such as "microbeads" and "beads," are also used synonymously.

#### 2.2. Reasons for microencapsulation

There are several reasons for employing microencapsulation in pharmaceutical formulations:

**2.2.1. Extended or sustained drug release**

Microencapsulation is widely utilized to achieve extended or sustained release of drugs, improving patient compliance and therapeutic outcomes.

**2.2.2. Taste and odor masking**

Microencapsulation can enhance patient acceptability and adherence by encapsulating drugs with unpleasant tastes or odors.

**2.2.3. Conversion of liquids into free-flowing powders**

Microencapsulation enables the transformation of liquid drugs into free-flowing powders, facilitating handling and processing.

**2.2.4. Stabilization of light, moisture, or oxygen-sensitive drugs**

The protective coating provided by microencapsulation can stabilize drugs that are sensitive to environmental factors, prolonging their shelf-life and ensuring efficacy.

**2.2.5. Avoidance of drug incompatibility**

Microencapsulation can prevent interactions between incompatible drugs or excipients, enhancing formulation stability.

**2.2.6. Modification of absorption site**

Microencapsulation can improve the bioavailability and efficacy of drugs by altering the release profile and targeting specific absorption sites.

**2.2.7. Protection against potential sensitization**

Encapsulation of toxic substances, such as pesticides, can minimize the risk of sensitization or adverse reactions.
2.3. Classification

Microparticles can be classified based on their core and shell materials (Figure 1):

2.3.1. Core Material

The core material refers to the active ingredient or substance encapsulated within the microparticle. It can be a drug, a nutrient, a cosmetic agent, or any other desired compound.

2.3.2. Shell Material

The shell material, also known as the coating material or wall material, forms the protective layer surrounding the core. Various natural and synthetic polymers, such as proteins, polysaccharides, and biodegradable polymers, can be employed as shell materials, depending on the desired release characteristics and compatibility with the core material. [18-20]

![Core and Shell material of microparticles](image)

**Figure 1.** Core and Shell material of microparticles

2.4. Advantages

Microencapsulation offers numerous advantages [21-23], making it a valuable technology in various industries, including pharmaceuticals, cosmetics, and agriculture:

- Protection of encapsulated active agents or core components from environmental factors, such as light, moisture, and oxygen.
- Transformation of gases and liquids into solid particles, facilitating handling and storage.
- Modification of surface and colloidal properties of different active agents, enabling improved solubility, bioavailability, and targeted delivery.
- Alteration and control of drug release profiles from various pharmaceutical dosage forms, enabling extended or sustained release formulations.
- Development of prolonged controlled release dosage forms by altering or delaying the release of encapsulated active agents or core materials

2.5. Disadvantages

While microencapsulation offers significant benefits, it is important to consider the following potential drawbacks:

- Expensive production processes: Certain microencapsulation techniques can be costly, particularly at larger scales.
- Reduced shelf-life for hygroscopic agents: Hygroscopic agents encapsulated within the microparticles may have a shortened shelf-life due to potential moisture absorption.
- Uneven coating: Inconsistencies in the microencapsulation coating can affect the release of encapsulated materials, potentially leading to variability in performance

3. Microencapsulation Techniques

Various techniques (Figure 1) have been developed and employed for microencapsulation, each with its own unique principles and advantages. In this section, we will explore some of the commonly used methods [25-27].
3.1.1. Air Suspension Technique

The air suspension technique involves dispersing solid and particulate core materials in a supporting air stream and spray coating the air-suspended particles. The core materials are suspended in an upward-moving air stream within the coating chamber. The coating substance is sprayed onto the particles as they recirculate through the coating zone. The coating is applied to the core material with each pass, and the product is dried by the supporting air stream during the encapsulation process. (Figure 2a)

3.1.2. Coacervation Phase Separation

The coacervation phase separation process involves three main steps:

- Formation of three immiscible phases: the coating substance, the core material, and the liquid manufacturing phase.
- Deposition of the liquid polymer coating onto the core material.
- Rigidization of the coating, typically through desolvation, cross-linking, or heat treatment, to form microcapsules.

The coating polymer is deposited around the interface formed between the liquid vehicle phase and the core material. Phase separation of the polymers can be induced by physical or chemical changes in the coating polymer solutions, leading to the formation of a two-phase liquid-liquid system.

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![Figure 1. Types of microencapsulation techniques](image)

![Figure 2. a. Air suspension process b. Pan coating c. Spray drying](image)
3.1.3. Pan Coating

The pan coating technique is widely used in the pharmaceutical industry for preparing controlled release particulates. It involves coating relatively large spherical core materials, typically larger than 600 μm, with various polymers. The coating is sprayed over the chosen solid core material in the coating pan as an atomized spray or solution. Heated air is often circulated over the coated items during the coating process to facilitate solvent removal. In some cases, a drying oven is used to complete the solvent removal process (Figure 2b).

3.1.4. Spray Drying and Spray Congealing

Spray drying and spray congealing are similar microencapsulation techniques that involve dispersing the core material in a liquefied coating agent and introducing the core-coating mixture into an environment that induces rapid solidification of the coating (Figure 2c). In spray drying, the coating material's rapid evaporation of a solvent causes solidification. The core-coating mixture is sprayed into a stream of hot air or gas, leading to the evaporation of the solvent and the formation of dried microparticles. Spray congealing, on the other hand, relies on the thermal congealing of a molten coating material or the solidification of a dissolved coating by introducing the core-coating mixture into a non-solvent. The non-solvent or solvent is subsequently removed from the coated product by various methods, such as evaporation or sorption extraction.

3.1.5. Solvent Evaporation

The solvent evaporation method is suitable for preparing liquid manufacturing vehicles, typically oil-in-water (O/W) emulsions. In this technique, the coating polymer is dissolved in a volatile solvent that is immiscible with the liquid manufacturing vehicle phase. The core substance (drug) to be microencapsulated is dissolved or dispersed in the coating polymer solution. The core-coating material mixture is then agitated and distributed throughout the liquid manufacturing vehicle phase to produce microcapsules of the desired size. Continuous agitation is maintained until the solvent evaporates and separates into the aqueous phase, resulting in hardened microcapsules.

4. Applications of Microencapsulation

Microencapsulation finds numerous applications across various industries due to its versatility and ability to impart beneficial properties to encapsulated substances. Some notable applications include:

4.1. Pharmaceutical Industry

- Extended or sustained drug release formulations
- Taste and odor masking of bitter or unpleasant drugs
- Protection of light, moisture, or oxygen-sensitive drugs
- Targeted drug delivery to specific sites

4.2. Cosmetic Industry

- Encapsulation of fragrances, essential oils, and active ingredients
- Controlled release of cosmetic agents
- Protection of unstable or volatile compounds

4.3. Food Industry

- Encapsulation of flavors, colors, and nutrients
- Protection of sensitive ingredients from degradation
- Controlled release of flavors or aromas

4.4. Agrochemical Industry

- Controlled release of pesticides, herbicides, and fertilizers
- Protection of active ingredients from environmental factors
- Reduction of environmental impact through targeted delivery

4.5. Textile Industry

- Encapsulation of dyes, fragrances, and antimicrobial agents
- Controlled release of functional finishes
5. Conclusion

Microencapsulation has emerged as a pivotal technology in the field of novel drug delivery systems and beyond, offering significant advantages in enhancing the efficacy, stability, and controlled release of various active ingredients. By encapsulating sensitive compounds within protective coatings, microencapsulation techniques have revolutionized the way drugs, cosmetics, agrochemicals, and food additives are formulated and delivered. The versatility of microencapsulation lies in the availability of various techniques, each tailored to specific requirements and applications. From spray drying and coacervation to solvent evaporation and pan coating, these methods enable the precise control of particle size, release profiles, and encapsulation efficiency. In the pharmaceutical industry, microencapsulation has proven invaluable in addressing challenges such as taste masking, bioavailability enhancement, targeted delivery, and sustained release formulations. These advancements have led to improved patient compliance, reduced dosing frequency, and minimized adverse effects, ultimately enhancing therapeutic outcomes.

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