Microemulsion technology: Nanoparticle-based emulsion as a sustainable natural product formulation

Mikael Ham Sembiring, Arsyadi, Taufik Riadi
Faculty of Military Pharmacy, Indonesia Defense University, Sentul, Bogor, Indonesia

Keywords
Essential oil
Microemulsion
Nanoparticle
Natural product
Sustainable technology

Introduction
Developing environmentally friendly formulations has emerged as a critical topic of discussion in various industries, including the cosmetics, personal care, and home care sectors. Increased consumer demand for environmentally and socially responsible goods drives this sustainability movement forward. Sustainable formulation ideas are becoming an increasingly important focus for businesses to align themselves with the most recent green trends and address the environmental effects of their goods (Marinho et al., 2022). Conventional formulations often produce excessive waste or depend on non-renewable resources. Sustainable formulations emphasise the efficient use of resources, focusing on renewable ingredients, decreased energy usage, and limited utilisation of hazardous compounds (Martins et al., 2023).

Natural products are environmentally friendly substances that provide several benefits. Natural products are widely used in pharmaceuticals as a source for drug discovery and as lead compounds for drug development. Approximately 35% of the yearly worldwide pharmaceutical market comprises natural goods or their derivatives. This category mostly includes plant-derived compounds, accounting for 25% of the market, followed by microorganisms at 13% and animal sources at 3% (Najmi et al., 2022).

Natural products are often used in the form of essential oils. Essential oils have complex combinations of volatile molecules obtained from plants. Essential oils have many benefits in pharmaceuticals, cosmetics, botany, and food and beverages (Irshad, et al., 2020).

Abstract
Natural products refer to chemicals or compounds from living creatures occurring naturally in the environment. Natural commodities, such as spices, herbs, and medical remedies, are often used as additions. Natural products have many advantages, such as their environmentally sustainable nature, compatibility with the skin, affordability, and widespread availability. Nevertheless, developing natural products has several challenges, such as limited options for ingredients, functional limitations of natural substances, considerations about stability and preservation, consumer perceptions and expectations, and the variability in the composition of natural products. Microemulsions are prevalent emulsions that incorporate nanoparticles. A microemulsion is a homogeneous, transparent liquid consisting of oil, water, and a surfactant, often combined with a cosurfactant. The use of microemulsions has been extensively observed in many chemical and pharmacological applications owing to their unique attributes, such as a significant interfacial area, little interfacial tension, and the ability to solubilise and deliver hydrophobic substances. Therefore, the present review paper aims to comprehensively analyse microemulsion technology and its applications as an environmentally friendly formulation for natural products. Here, we propose strategies and research on microemulsions to overcome problems in natural product formulations.
Most of the chemicals found in essential oils are terpenes, esters, aldehydes, ketones, alcohols, and phenols. Different plant types, growth conditions, harvest times, and the part of the plant used for extraction can all change the chemicals that make up essential oils (Dhifi et al., 2016).

**Microemulsion technology**

According to Danielsson and Lindman (1981), a microemulsion is a system of water, oil, and an amphiphile, a single optically isotropic and thermodynamically stable liquid solution. The components of a microemulsion mixing system include air, oil, surfactant, and cosurfactant. Preconditioning using surfactants and cosurfactants lowers surface tension, resulting in thermodynamically stable microemulsions (Souto et al., 2022). Because the size of the particles in microemulsions ranges from 10 nm to 100 nm, the microemulsions look visibly translucent. Thermodynamic stability, visual clarity, enhanced lipophilic substance solubility, enhanced bioavailability of active compounds, and enhanced penetration ability are among the advantages of using microemulsions (Sharma et al., 2020).

The use of high-frequency ultrasound has created microemulsions containing essential oils. In addition, several methods can be used in making microemulsions (Figure 1) (Racovita et al., 2023):

- **Method of phase inversion:** This approach involves adding an extra dispersed phase to the surfactant system under temperature control, resulting in phase inversion. Drug and agrochemical particle size decreases during phase inversion, resulting in active release kinetics.

- **Method of phase titration:** This mechanism, known as spontaneous emulsification, may be shown using phase diagrams. The phase diagram helps analyse interactions between microemulsion components during mixing.

![Figure 1: Essential oils are obtained by extracting the designated herbs. Subsequently, a comprehensive examination is conducted to determine the configuration of the desired microemulsion and whether it adopts the water-in-oil (W/O) or oil-in-water (O/W) composition. The selection of surfactant and co-surfactant should be predicated upon their safety and efficacy. Subsequently, a combination of surfactants, co-surfactants, and essential oils is blended inside a singular receptacle. Two distinct procedures are used in producing microemulsions: the titration method and the inverse approach. The titration approach does not use an inverse phase but utilises a phase diagram to ascertain the optimal combination of oil and surfactant. Conversely, the inverse method involves a phase inversion phenomenon occurring in the microemulsion due to temperature variation.](image-url)
Application of microemulsion essential oil

Previous research has demonstrated the antimicrobial properties of peppermint, oregano, and rosemary essential oils when formulated as microemulsions. These oils are effective against Escherichia coli and Listeria monocytogenes, with minimum inhibitory concentration (MIC) and minimum bactericidal concentration (MBC) values 20-75% lower than the non-encapsulated essential oils (Dávila et al., 2020). Microemulsions containing essential oils have shown notable efficacy in repelling insects, giving a viable and ecologically sustainable substitute for mosquito larvicides and agricultural pest control. Furthermore, Pavela et al. report that using microemulsions containing Apiaceae essential oils has shown efficacy as mosquito larvicides. These microemulsions have LC50 values ranging from 1.45 to 4.01 ml L-1, indicating their potency in inducing larval death (Pavela et al., 2019).

Deeper research proves the penetration-enhancing effects of clove oil, peppermint oil, and tea tree oil microemulsion, which may be attributed to the terpenic components present in these oils (Thakur et al., 2021). Specific terpene components, such as limonene, menthol, carvone, and thymol, have minimal systemic toxicity and excellent skin penetration-boosting qualities. In addition to these, because of their ability to increase penetration, fatty acid esters, such as ethyl oleate and isopropyl myristate, and monounsaturated fatty acids, such as oleic acid, have also been widely used for the hydrophobic phase of Microemulsion production. This finding underscores their therapeutic potential and suggests their use as vehicles for drug delivery (Ait-Touchente et al., 2023).

The composition of the essential oil obtained from Lavandula angustifolia Mill. Consists of many compounds, such as linalool, linalyl acetate, geraniol, and β-caryophyllene. The microemulsion oil derived from Lavandula angustifolia exhibited notable antibacterial effects against Klebsiella pneumoniae, MRSA, and Staphylococcus aureus. The MIC measurements for these bacteria were determined to be 32, 16, and 16 µg/mL, respectively. However, the microemulsion oil showed little effectiveness against Proteus mirabilis. The recently produced formulation of nano-gold and nano-Lavandula angustifolia demonstrated complete eradication of pre-existing P. mirabilis biofilm. The MIC and minimum biofilm eradication concentration (MBEC) values for this formulation were determined to be 8 and 16 g/mL, respectively (Figure 2) (Fadel et al., 2023).

Figure 2: Microemulsions containing essential oils have shown efficacy in preventing bacterial resistance, reducing minimum inhibitory concentration (MIC) and minimum bactericidal concentration (MBC), enhancing the anti-larval activity against mosquitoes, and improving the skin penetration of chemicals.
Conclusion
Microemulsion technology allows for particle size reduction from 100 nm to 400 nm. Thermodynamically stable microemulsions may be produced, and this approach can decrease surface voltage. Furthermore, using microemulsion technology for vegetable essential oils may improve the antibacterial effect, boost the killing impact against mosquito larvicides, and increase active chemical absorption. It can also be used as an agricultural pest management method.

Acknowledgement
Figure 1 & Figure 2 were made with BioRender.com.

Source of funding
This research received no external funding.

Conflicts of Interest
The authors declare no conflict of interest.

References


Racovita, R. C., Ciuca, M. D., Catana, D., Comanescu, C., & Ciociarla, O. (2023). Microemulsions of nonionic surfactant with water and various homologous esters: Preparation, phase transitions, physical property measurements, and application for extraction of tricyclic antidepressant drugs from aqueous media. Nanomaterials, 13(16), 2311. https://doi.org/10.3390/nano13162311

