



A comprehensive Review on Formulation, Characterization, Composition, Stability & Preparation, of Nanoemulsions

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Abstract— *The use of Essential Oils (Eos) alone is limited due to their disadvantages such as high volatility, low stability, bioavailability and solubility. Therefore, Essential Oils are generally use in the form of nanoemulsions. The food and pharmaceutical sectors have shown a great deal of interest in essential oil-based nanoemulsions (EO-NEs) because of their improved stability, bioavailability, and functional properties. Nano-based emulsions havereceived considerable attention recently due to their physico-chemical properties and kinetic stability , attractive appearance. Furthermore, their rheological properties make them superior over counter parts. The ability of nanoemulsion to improve the solubility, bioavailability and stability of weakly water-soluble drugs has emerged their roles as promising drug delivery systems, drug administration and vaccine formulation, cosmetics, and food, it also has numerous applications in the field of agriculture. Nevertheless, their benefits, some concerns may represent obstacles that hinder their applications such as formulation complexity, stability issues and industrial production scale-up. To get beyond these hurdles and completely realize the potential of nanoemulsion in commercial and therapeutic applications, more research and innovation are need. This review intends to offer an overview of the formulation, Characterization, Composition, Stability & Preparation, of nanoemul sions specifically, recent literature has been examined in order to define the most common practices adopted (materials and fabrication methods), highlighting their suitability and effectiveness. Finally, relevant to some points such as advantages disadvantages, and applications of nanoemulsions.*

Keywords— *Characterization, Essential Oils, Formulation, Nanoemulsions, Preparation.*

I. INTRODUCTION

Essential oils (EOs) are secondary metabolites from aromatic plants. Moreover, they are plant-derived oily bioactive compounds from various parts like roots, leaves, flowers, and seeds. Through methods such as steam distillation, cold pressing, steam distillation, microwave or ultrasound assisted extraction etc [1, 2]. Essential oils are a complex mixture of various bioactive chemical molecules, they are typically use in small quantities due to their potent aroma and flavor, they are recognised as safe by both the US Food and Drug Administration (FDA), applications and can used as alternatives to chemical preservatives [3 - 6]. Due to these diverse, complex bio-structures, essential oils have been widely used in the traditional medicine as essential oil posse various biological activities like

antimicrobial, antiviral, fungicidal, insecticidal and herbicidal, antioxidant, anti-inflammatory, anti-allergic, anticancer agents, expectorant, digestive, diuretic agents [7, 8]. The biggest challenges of modern botany is to move towards the replacement of synthetic compounds with natural ones, including EOs, in order to exploit their promote sustainable development based on a circular economy [8]. Interest in these kinds of plant based secondary metabolites is increasing day by day [9, 10]. Essential oils have poor physico-chemical properties, such as water insolubility [11]. However, these problems can be preventing through the encapsulation of EO within appropriate delivery systems, thereby amplifying their biological efficacy, [12].Therefore; nanoemulsification technology offers a good alternative for encapsulating these bioactive compounds, making them hydrophilic [13].

Recently, medical studies showed a possible anticancer activity of some EOs, attributable to an antimutagenic (i.e., *Salvia officinalis* L., *Origanum onites* L. and *compactum*, *Lavandula angustifolia* EOs [14, 15] and prooxidant activity (i.e., *Origanum compactum*, *Artemisia herba-alba* and *Cinnamomum camphora*, *Glandora rosmarinifolia* EOs [16, 17]. This latter is a fundamental property to reduce local tumor volume and cell proliferation [18]. The studies was confirmed that the main components of Cinnamon essential oils (CEO), such as limonene and Cinnamaldehyde, have strong antimicrobial effects on foodborne pathogens like *Escherichia coli* and *Listeria monocytogenes* [19]. In addition, encapsulating extracts from graviola leaves [20], garlic [21], pep-permint [22], basil oil [23], and CEO in nanocapsules enhances their antimicrobial and antioxidant properties, thereby optimizing growth, gastrointestinal health [21]. The interest in nanoemulsions has experienced a continuous increase in the last years, emulsification is the process of dispersing two or more immiscible liquids together to form a semi stable mixture ([24- 26]. Nanoemulsions are three main components used in the preparation of nanoemulsions are oil, surfactant and cosurfactants have unique physicochemical properties [27-32]. Emulsions (liquid droplets and/or liquid crystals dispersed in a liquid) are prepared using nanotechnology (1-100 nm) by reducing the surface tension between the dispersed and dispersed phases. This is usually achieved through vacuum and electrostatic repulsion between the emulsions, is preferred, the stability of emulsions, which ranges from a few hours to several years, is influenced by important characteristics related to the preparation procedures and the ingredients used. Mollet and Groppenmann emphasize that emulsification requires both chemical and physical energy [33- 38]. Nanoemulsion is a nonequilibrium mechanism that is create with the aid of an external or internal energy source. This is accomplish by a variety of methods that are grade as either high-energy or low-energy by using mechanical devices [39]. Emulsions, which are prepared using nanotechnology, characterized by; exhibit excellent skin penetration and long-lasting effects. In contemporary times, nano-drug delivery systems have emerged as a highly effective approach for delivering drugs within the human body [40-44]. Nanoencapsulation based nanoemulsions, play a pioneering role in encapsulating herbal, Tween-20 and 80 are nonionic, biocompatible, and non-toxic surfactants and widely implicated in medicine, cosmetic, food industries products as used as emulsion surfactants [45-50]. The nanoencapsulation approach could improve the essential oils' physico-chemical properties and stability, by enabling their water dispersability, and by protect them from the interaction with the environment [51].

This work will endow with new information about nanoemulsions based on essential oils in different areas such as food, cosmetic, and pharmaceutical drug delivery systems. In addition, offer a technical overview on the formulation, preparation and stability parameters of Molecular extracts and Nano-extracts when used as vehicle and delivery systems for essential oils, based on the most relevant works reported in the literature in the last years. Moreover, it aims to provide a formulation guide for researchers that need to encapsulate essential oils in such Nano systems.

II. NANOEMULSIONS FORMULATION

An understanding of the physics of nanoemulsion formation is critical for the control of nanoemulsion droplet size. Nanoemulsions are typically prepared in a two-step process where a macroemulsion is first prepared, and is then converted to a nanoemulsion in a second step [52]. There are a number of techniques for preparing nanoemulsions, such as high-pressure homogenization, microfluidization, phase inversion, spontaneous emulsification, solvent evaporation and hydrogel formation. Multiple emulsions are usually prepared using the double emulsion-solvent evaporation technique. [53].

Theories of Nanoemulsions Formulation

1. Solubilization Theories

it is believed that the nano emulsion is getting closer to thermodynamically stable monophasic solutions of spherical micelles that are swelled in water (w/o) or oil (o/w) [54].

2. Interfacial theory

It is also called as mixed film, the surfactant and co-surfactant form a complex layer at the oil-water interface, which is what cause the spontaneous creation of nanoemulsion droplets. According to the general provisions of the theory, an increase in the degree of dispersion of micro droplets can be achieved by lowering the interfacial tension at the water-oil interface to reaching extremely low values (from almost zero to negative). As Shulman's work [55]. This is represent by the equation below:

$$y_i = y_0/w - \pi_i \quad (\text{equation 1})$$

Where, y_0/w : Oil-water interfacial tension, π_i : Spreading pressure, y_i : interfacial tension [56].

3. Thermodynamic theory

When interfacial tension between two immiscible phases reduces to zero, causes spontaneous formation of micro emulsions and formed negative free energy helps to make emulsion thermodynamically stable [57]. T. P. Hoar and J. H. the term microemulsion when they used this term to

describe multiphase system . Discovery of microemulsions confirms well before use in the form liquid waxes., there are four types of micro emulsion phases exists in equilibrium, these phases are referred as Winsor phases [58, 59].

III. PHYSICOCHEMICAL CHARACTERIZATION ON NANOEMULSIONS

3.1. Droplet size & distribution

The normal range of droplet sizes for nanoemulsion is 20-500 nanometers. area-to-volume ratio produced by the small droplet size improves the interaction between the nanoemulsion and biological membranes, enhancing the absorption and bioavailability of the materials that are entrapped [60, 61]. As the droplet size is significantly smaller than the wavelength of visible light, nanoemulsions are often transparent in appearance [62]. Nano-emulsions size distribution is the most important parameters of nano-emulsions since it is relate to applicative properties [63, 64]. Methods used for size characterization include photon correlation spectroscopy (PCS) (i.e., dynamic light scattering, DLS), fluorescence correlation spectroscopy (FCS), electron microscop [65, 66].

3.2. Zeta potential analysis

The liquid layer surrounding the particle exists as two parts; an inner region (Stern layer) where the ions are strongly bound and an outer (diffuse) region where they are less firmly associated. Within the diffuse layer, the ions and particles form a stable entity inside a notional boundary. When a particle moves (e.g. due to gravity), ions within the boundary move it. Those ions beyond the boundary stay with the bulk dispersant. The potential at this boundary is the zeta potential [67, 68]. Sakhalin et al. designed the first zeta potential changing nanoemulsion exhibiting a zeta potential change from just -1 mV to +1 mV [69]. Up to date, all zeta potential changing nanoemulsions contain a cationic surfactant being responsible for a positive zeta potential [70]. Concept of zeta potential is rooted in the examination of electro kinetic phenomena, which describe the relative motion between a charged surface and a liquid under the effect of an electric field. The most frequently encountered electro kinetic phenomena used to measure zeta potential are Electrophoresis, Electro osmosis [71, 72].

3.3. Viscosity

The viscosity of freshly prepared nanoemulsion samples usually measured using a Brookfield DV-ETM viscometer. [73]. The proportions of water, oil, and surfactant in a nanoemulsion determine its viscosity, Lowering the content of cosurfactants and surfactants raises viscosity by raising the interfacial tension between the water and oil, whereas raising the water content lowers viscosity [74, 75].

3.4. Encapsulation efficiency (EE)

Encapsulation of essential oils in nanoemulsions systems represents a viable and efficient approach to increase the physical stability of bioactive compounds. Thus protecting them from loss and undesirable changes [76]. Indeed, the use of this technology gains special relevance when it comes to the formulation of biopharmaceuticals [77]. Encapsulation Efficiency (EE) was regarded as a bulk property and was measured as the average of all DDS particles in a solution. Recent technical advances however, allowed the monitoring of single particles [78, 79], to unique information on heterogeneous properties that were lost due to averaging [80]. Encapsulation Efficiency is calculated by following equation:

$$(EE) \% = \frac{\text{Total oil} - \text{Free oil}}{\text{Total oil}} \times 100 \text{ (equation 2) [81 - 83]}$$

The encapsulation efficiency of the micro particle or microcapsule or microsphere will be affect by different parameters. Figure 1 illustrates the factors influencing encapsulation efficiency [84].

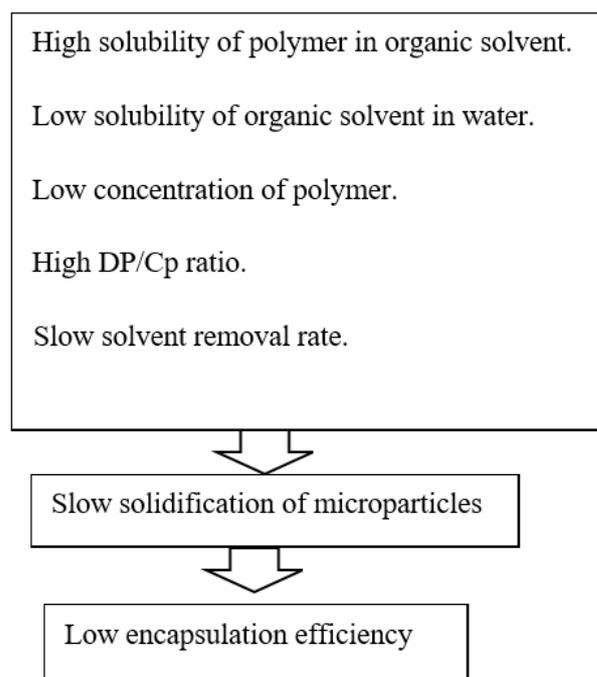


Fig.1: factors influencing encapsulation efficiency [85].

3.5. Surface tension

Surface tension usually measured using a thermostatically controlled processor tensiometer K10 PST at 22 °C [86].

3.6. Rheological measurements

The rheological properties of Nano-emulsions can be turn from flowing fluid to a solely relaxing fluid or to a gel-like system by controlling the dispersed phase volume fraction and droplet size, or by adding additives in the bulk [87].the

study of interfacial properties, surface tension and dilatational rheology is getting more and more attention in recent years [88]. Rheological measurements usually done perform at $25 \pm 0.1^\circ\text{C}$ using a Bohlen remoter equipped with a cone/plate apparatus 40 mm per 4° . For each sample, continuous variation of shear rate $\dot{\gamma}$ will applied and the resulting shear stress σ will measured. Viscosity of dispersions with Newtonian flow properties calculated according to the relation: $\eta = \sigma / \dot{\gamma}$ (equation 3) [89].

IV. COMPOSITION OF NANO EMULSIONS

4.1. Emulsifier

An emulsifier has the ability that mixes two immiscible liquids (oil and water) to form a homogeneous dispersion. It plays a significant role in the food industry and it could prevent phase separation. [90]. Food colloids emulsions and foams have their origins in nature, Milk, for example, has a naturally occurring membrane, which allows solid fat to be disperse into an aqueous phase [91].

Table 1: Properties of emulsions [92, 93].

Emulsion	Droplet Size	Thermodynamic Stability	Appearance
Nanoemulsion	5-200 nm	Unstable	Transparent
Microemulsion	5-100 nm	Stable	Transparent
Macroemulsion	0.1-100 μm	Unstable	Turbid

4.2. Oil phase

The oil phase could consist in a carrier oil in which the lipophilic bioactive compound dissolved [97]. Oil-in-water miniemulsions are prepared at slow agitation by an emulsification process that involves the use of mixed emulsifier solutions of an ionic surfactant [98]. Penetrating the tail group region of the surfactant monolayer and causing it to swell, the oil components also affect curvature [99]. Because short chain oils, penetrate the lipophilic group more [100]. Nano-emulsions are thermodynamically unfavorable systems [101], the chemical stability is also important for quality control of the drug-loaded Nano-emulsions [102, 103]. Some plant oils (corn, olive, sunflower, sesame, oils) defined as “ripening inhibitors” is needed [104-107]. Plant oils less water-soluble, they are able on positively influence their partitioning between the lipid droplets and the aqueous phase, for these reasons, several works in the literature reported the mixing of essential Oils, acting as active ingredient, and ripening

inhibitor oils, in order to guarantee the long-term stability of the system [108, 109].

4.3. Surfactant

A surfactant molecule consists of two parts. One part is hydrophobic and the other one is hydrophilic. These molecules are highly active in the interfaces between air and water or oil and water. There are four main classifications of surfactants: anionic, nonionic, cationic and amphoteric surfactants, therefore, proper selection of surfactants becomes a crucial criterion. [110, 111]. Surfactants are essential for stabilizing nanoemulsions [112]. These amphiphilic molecules form a protective layer around droplets, offering steric, electrostatic, or dual electro-steric stabilization [113]. The selection of surfactants depends on the desired droplet size, stability, and application. Cosurfactants are often added to strengthen the interfacial and further stabilize the emulsion. Examples include polyethylene glycol [114, 115].

Table 2: Types of surfactant [116 - 119]

Sr. No	Types of surfactant	Examples
1	Non-ionic	Sorbitan monooleate (Span 80), Tween 80
2	Cationic	Ecithin , soybean egg,. Quaternary ammonium alkyl salts
3	Anionic	Sodium bis-2-ethylhexylsulphosuccinate (AOT)
4	Ampholytic	Phospholipids (Lecithin, Ndodicylalanine)

4.4. Co-surfactant

Drugs that are not water-soluble usually made into emulsion preparations. Emulsion preparations have poor stability; surfactants are ingredients that affect nanoemulsions to reduce surface tension [120]. Surfactants have unique

molecular characteristics; a short to medium hydrophobic chain, weak amphiphilic nature, and terminal small hydrophilic group (such as hydroxyl group characteristic of low molecular weight alcohols and diols). These properties increase the tendency of cosurfactants to interact with

surfactant monolayers at interfaces, perturbing the packing and long-range order of surfactant molecules and promoting better interface fluidity and curvature in favor of NE formation [121]. Furthermore, the balanced amphiphile nature of these molecules promotes their selective partitioning between phases, changing their composition and thereby their relative hydro/lipophilicity of the immiscible phases to improve better miscibility [122].

Table 3: List of co-surfactant used in nanoemulsions [123]

Sr. NO	Co-surfactant
1	TranscutolP
2	Glycerin, Ethylene glycol
3	Propylene glycol
4	Ethanol
5	Propanol

V. STABILITY

5.1. Centrifugal stability

The stability of nanoemulsions under centrifugation reflects the emulsion resistance and the integrity of the interfacial film [123,124]. use varying speeds (3000, 5000, and 8000 r/min) for 30 min, the absorbance of the non-layered emulsion is then measured at 210 nm before and after centrifugation, using distilled water as a reference sample. The permeability calculated using the following equation:

Transmittance formula: $T (\%) = A1/A2 \times 100$ (equation 4)

Where: T: transmittance; A1: absorbance of nanoemulsion before centrifugation; A2: absorbance of nanoemulsion after centrifugation [99].

5.2. Thermal stability

Increased temperatures can cause droplets to absorb energy in the Nano-emulsion, resulting in the emulsification, coalescence and flocculation of the Nano-emulsion [125]. Over time, they evolve into two distinct liquid phases, an irreversible process driven by the lower free energy level of the phase-separated state compared to the emulsion state [126]. The kinetics of Nano emulsion breakdown follow a first-order rate law, where the rate is exponentially proportional to $\exp(-\Delta E/kBT)$, with ΔE denoting the energy barrier, representing the Boltzmann constant, and T indicating the temperature [127]. The irreversible breakdown poses significant constraints on the utility of Nano emulsions, especially in scenarios where exposure to high temperatures during production, storage, or application is inevitable, such as in food processing, formulations, and pharmaceutical [128].

5.3. PH stability

The pH value of the Nano emulsion systems adjust by adding Hydrochloric acid or Potassium hydroxide to the system, which monitor by using a Sartorius basic pH [129]. To pH testing conduct use the methodology described by Shoviantari et al. [130]. Before evaluating the preparation's pH, the pH meter will be calibrate using buffer solutions (pH 4.00, 7.00, and 10.00). To guarantee its correct functioning. [131].

5.4. Storage stability

There are few studies have investigated the storage stability of Nano emulsion droplets; Environmental conditions that effect on the stability of emulsions include temperature, pH, and ionic strength [132- 134]. The emulsion stored at higher temperatures increases the thermal energy of the droplets and therefore increases the frequency of droplet collisions. It also reduces interface viscosity, which results in a faster film-drainage rate and faster droplet coalescence [135]. Nano-emulsions have a long kinetic stability that can be maintaining for months because of their small size and low dispersion index. When the Nano-emulsion droplet size is less than 200 nm, the Nano-emulsion is highly resistant to gravitational separation, which is ideal for the development of products with a long shelf life [136]. The shelf of Nano emulsions is tested by storing a sample of fresh emulsion (5 ml) in a sealed container at 25°C for 2, 4, 6, 12, and 24 days respectively. The droplet size, dispersion index (PDI), and zeta potential are then measured at each storage period [125].

VI. NANO EMULSIONS PREPARATION

Nano emulsions can be prepared by using high and low energy methods. In high-energy methods, mechanical devices deliver required large disruptive forces. On the other hand, in low energy methods, there is no need for an external force [137]. Production of Nano emulsions achieved by using the intrinsic physiological properties of the system. [138]. The most extensively used methods for the preparation of Nano emulsions: are Spontaneous Emulsification Method (SE), Phase inversion Temperature method (PIT), Emulsion phase inversion (EPI) method/ catastrophic phase, Emulsion inversion point (EIP) method, High-pressure homogenization (HPH) method, Micro fluidizers, High-Intensity Ultrasound (HIU), and Rotor stator mixer [139].

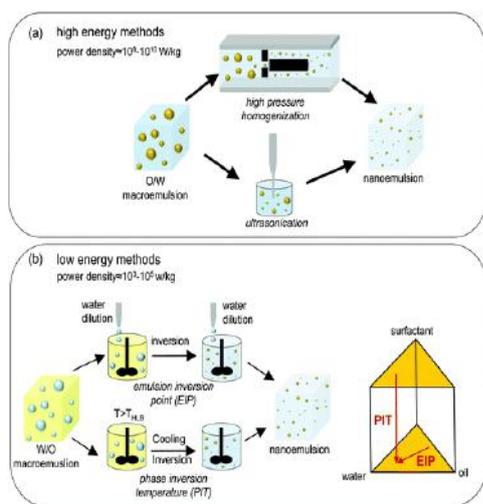


Fig.2: Nano emulsions Preparation

6.1. Low-energy methods

Low-energy approaches rely on the spontaneous formation of tiny oil droplets within oil-water-emulsifier mixtures when either their composition or the environment conditions are altered [140]. A number of different Nano emulsion preparation methods based on this principle, including: [141].

6.1.1. Spontaneous emulsification method (SE)

The spontaneous formation of an O/W Nano emulsion facilitated by specific temperatures relies on the chemical compounds present in both phases, as well as the utilized emulsifier [142]. Solvents can facilitate this spontaneous process [143]. It benefits from the chemical energy replacement based upon dilution process with the continuous phase, which occurs usually at constant temperature [144]. This method can produce Nano emulsions at room temperatures and no special devices are required. It subjected to interfacial tension, viscosity of interfacial and bulk, phase transition region, surfactant structure, and surfactant concentration. [145].

6.1.2. Phase inversion temperature method (PIT)

In the PIT method, Nano emulsions prepared by the formation of micro emulsions at its PIT followed by immediate cooling to room temperature, which can be divided into 3 main steps:

1. Non-ionic surfactant, oil and water stirred at room temperature to form a coarse emulsion [146].
2. The mixture gradually heated up to around PIT [147].
3. The solution rapidly cooled to the room temperature with continuous stirring, resulting in the formation of o/w Nano emulsions, mixture is rapid cooled either by immersing into an ice bath or by diluting with cold water [148]. Inversion Temperature method (PIT) has the advantage over SE

method in that its composition is without organic solvent which is an integral component of the SE method. The PIT method also has superiority over the PIC method in that the nanoemulsion droplets have lower diameter and polydispersity index (PDI). The emulsification efficiency of PIT method has also been found higher (1) than that of PIC method (0.35)40 [149, 150].

6.1.3. Emulsion phase inversion (EPI) method/ catastrophic phase

Nano emulsions can be fabricate using either low-energy emulsification (e.g., transitional phase inversion) [151,152]. Although low energy emulsification methods use higher amounts of surfactants, it requires only a simple stirring to produce an emulsion with desirable characteristics [153,154]. Which makes them more energy efficient opposed to high-energy methods, which require the use of mechanical devices [154]. The Emulsion Phase Inversion (EPI) method, is a low-energy technique for creating fine emulsions, by agitated water-in-oil (O/W), emulsion inverts into an oil-in-water (W/O) emulsion and vice versa. There are two types of phase inversion; transitional phase inversion and catastrophic phase inversion (CPI) [155]. Emulsification via phase inversion is widely used in fabrication of cosmetic products, pharmaceutical products (e.g., vesicles for drug delivery), foodstuff and detergents. Emulsification process strongly affected by preparation method; very different droplet size distribution could be achieved, which is strictly linked to the product stability [156]. The phase inversion of a W/O emulsion can be achieved not only through changes in thermodynamic state variables but also by the imposition of ultra-high deformation rates under constant thermodynamic conditions. Phase inversion carried out using a model static mixer, which is essentially a series of short capillaries with flow dividers. His degree of phase inversion is dependent on the rate of deformation and number of mixer units. The properties of the emulsions produce by FIPI-emulsification are evaluated through particle size and viscosity measurements and optical and electron microscopy studies [157].

6.1.4. Emulsion inversion point (EIP) method

In the emulsion inversion point (EIP) methods the change from one type of an emulsion to another (e.g., W/O to O/W or vice versa) is through a catastrophic phase inversion (CPI), rather than a transitional phase inversion (TPI) as with the PIC or PIT methods [158]. In this case, a W/O emulsion with a high oil-to-water ratio is formed using a particular surfactant, and then increasing amounts of water are added to the system with continuous stirring. Above a critical water content, the water droplet concentration is so high that they are packed very tightly together, and the emulsion reaches a phase inversion point where it changes

from a W/O to an O/W system. The size of the droplets formed depends on the process variables, such as the stirring speed and the rate of water addition [159]. Recently, it has been shown that the emulsion inversion point (CPI) method can be used to produce Nano emulsions ($r < 100$ nm) from food-grade ingredients [160, 161].

6.2. High energy methods

6.2.1. High-pressure homogenization (HPH) method

High pressure homogenization (HPH) was first introduced in the early 20th century by Auguste Gaulin, the basic operational principle of HPH remains unchanged since and it involves using a high-pressure pump to force the fluid through a small orifice. [162]. In the emulsion fabrication, a mixture of essential oil and Tween 80 (as surfactant) is disperse in distilled water, using a magnetic stirrer at 1500 rpm for 15 min, to form a primary emulsion. Subsequently, the primary emulsions subjected to a high-pressure homogenization [163].

Table 4: Preparation of nanoemulsions using Spontaneous Emulsification Method (SE) with different oils

Method	Bioactive Compound Encapsulated	Optimal Processing Conditions	Droplet Diameter (nm)	Ref
Spontaneous Emulsification Method (SE)	Peppermint oil	(1) titration of organic phase into aqueous phase, (2) constant stirring, 600 rpm, (3) room temperature	50	[164, 165]
	Eugenol oil	(1) deprotonated Eugenol in hot alkaline added to surfactant mixtures, (2) the mixtures acidic to pH 7.0, stirred, 600 rpm	109–139	[166]
	Black Cumin oil	Mix cumin essential oil with Tween 80 and distilled water using a magnetic blender for five minutes.	122.7	[167]
	mustard oil	1-Blends of Tween 20 and SDBS in a range of 1:1–1:6 ratios. 2-Mustard oil (5% w/w) add drop wise into different blends of emulsifier with slow mixing by magnetic stirrer at 50–60°C for final preparation of oily phase. 3- Aqueous phase form by adding two co-surfactants, in de-ionized water.	23	[168]
	pine oil	The method: Mix pine oil +Tween 80 + ethanol in + water with total cumulative value of all components not more than 100% v/v	633	[169]

Table 5: Preparation of nanoemulsions using Phase inversion & Emulsion inversion point Methods with different oils

Phase inversion Temperature method (PIT)	Curcuminoids	1. Non-ionic surfactant, oil and water are stirred at room temperature to form a coarse emulsion	20-100	[170]
	Lemon oil		100	[171]
	coconut oil	2. The mixture is then gradually heated up to around or above the PIT. 3. The solution is rapidly cooled to the room temperature with continuous stirring..	20-100	[146]
	ginger oil		100	[147] [148]
	Harmal oil	combining the essential oil with surfactants (like Tween 80/Span 20) and water, then mixing and homogenizing to create stable, nanometer-sized droplets	There is no study	
Emulsion phase inversion (EPI) method/ catastrophic phase	coconut oil	80 or Tween 20 & oil are prepare and add to it different amounts of water and the mixture was mixed using a vortex, for two minutes.	20-100	[172]
	Oregano Essential Oil		170-180	[173]
	Fenugreek oil		< 100	[174]
	canola oil		156.13 ± 2.3	[175]

	orange oil		146-200	[176]
	sesame oil		100	[177]
	Mexican Tea essential oil		Suggested:	
Emulsion inversion point (EIP) method	Castor oil	slowly adding water to an oil-surfactant(tween 80) mix, which suddenly transforms the system into tiny droplets at low interfacial tension.	35 ± 45	[178]
	coconut oil (VCO)		<300	[179]
	paraffin wax		<200	[180]

Table 6: Preparation of nanoemulsions using High-pressure homogenization (HPH) method with different oils

Method	Essential oil	Surfactant	Aqueous Phase	Size (nm)	Ref
High-pressure homogenization (HPH) method	fish oil	Tween80	sodium Caseinate	369.4 ± 3.8	[181]
	clove essential oil	Tween 80	Distilled water	90	[163, 182]
	lemon essential oil	Tween 80	Distilled water	100-140	[163]
	Saccocalyx satureioides Coss. essential oil	Tween 20	Distilled water	106,000	[183]
	Cinnamon essential oil	Tween 80	Lauric arginate	100	[184]
	cardamom oil	Tween 80	grape seed oil		[185]
	bergamot essential oil	xanthan gum	Distilled water	63.6 - 85.3	[186]
	groundnut oil	Tween 80	Distilled water	20-200	[187]
	borage oil	Tween® 80	Distilled water	334.4 ± 5.2	[188]
	Cellulose derivatives	Tween 20, 60, or 80	Distilled water	100-126	[189]

Table 7: Preparation of nanoemulsions using microfluidizers and high-intensity ultrasonic (HIU) devices with different oils

Method	Essential oil	Surfactant	Aqueous Phase	Size (nm)	Ref
Microfluidizers	Corn oil	quillaja saponin and soy lecithin	Whey protein isolate (WPI)	20-200	[190]
	linseed oil	lecithin	Distilled water	70.8	[191]
	Mineral oil	lecithin	Distilled water	<200	[192]
	Chia Oil	NaCas	Distilled water	< 200	[193]
	orange oil	gum Arabic and modified starch	Distilled water	77	[194]
High-Intensity Ultrasound (HIU)	olive oil	Tween 80, Span 60, or sucrose esters	Distilled water	< 200	[195]
	Castor Oil	Tween 80	deionized water	174.6	[196]
	Cady oil	Tween 80	deionized water	< 200	[196]
	Neem oil	Tween 20	Distilled water	< 100	[197, 198]
	Almond Oil	Tween 20	Distilled water	< 100	[199]
	juniper berry essential oil	Tween 20	deionized water	182-250	[200]

	Curcumin	soybean lecithin	glycerol in the aqueous phase	108	[201]
	Thyme Essential Oil	Tween 80	deionized water	54.6	[202]
	ginger oil	Tween 80	buffer or distilled water, ethanol	< 100	[203, 204]
	grapeseed oil	Tween 80	Distilled water	144.2 - 162.6	[205]
	Moringa Oil	whey proteins	Distilled water		[206]
	Garlic essential oil	Tween 80	Distilled water	215-820	[207]
	Aloe vera oil	Span® 85	Distilled water	< 200	[208]
	safflower oil	polysorbate 80	Distilled water	88.14 ± 1.52	[209]
	Basil oil	Saponins extracted from Quillaja lancifolia	Distilled water	174.1	[188]
	celery seed oil	Tween 80	Distilled water	23.4 ± 1.80	[210]
	lavender oil	Tween-80, Span-80, ethanol	Distilled water	-----	[211]
	Qassum oil	Tween 80/Span 80	Distilled water	Suggested: There is no study	

Table 8: Preparation of nanoemulsions using Phase inversion composition (PIC) & Spontaneous emulsification methods (SE) with different oils

Method	Essential oil	Surfactant	Aqueous Phase	Size (nm)	Ref
Phase inversion composition (PIC)	Neem oil	Tween 80	Distilled water	194	[198]
	Ginger oil	Tween 80	Distilled water	20–75	[212]
	coconut oil	Tween 80/Span 80 or lecithin	Distilled water	48.63	[179]
	Origanum Essential Oil	linseed mucilage	Distilled water	170-180	[173]
Spontaneous emulsification method (SE)	Lime Essential oil	Tween 80	Distilled water	20–60	[163]
	Cinnamon oil	Tween 80	Distilled water	100	[184]
	Peppermint Essential oil	Tween 20, 40, 60	Distilled water	70–170	[164]
	Clove oil				[182]
Phase Inversion Temperature (PIT)	oregano essential oil	Cremophor RH 40, Span 80	Millipore water	30–60	[213]
dynamic light scattering	Elongate oil	deionized water	non-heated water	115	[214]

6.2.2. Micro fluidizers

Two solutions (aqueous phase and oily phase) are united together and processed in an in-line homogenizer to create

a coarse emulsion, which allowed into a Microfluidizer where it is further developed to obtain a steady Nano emulsion. [215]. It is most widely employed in the

pharmaceutical industry [216]. In nanoparticle production, this process facilitates the size reduction and uniform dispersion of materials. For example, in the production of liposomes and lipid nanoparticles (LNPs), [217, 218].

6.2.3. High-intensity ultrasound (HIU)

Industrial scale emulsion formation typically takes place in high-pressure homogenizers (HPHs) [219]. HPHs used widely in many branches of chemical engineering, ranging from traditional dairy applications to Nano emulsification in pharmaceutical production and more [220]. Advantages of these devices high-energy efficiency, and Good emulsion stability - HIU devices are easy to operate and clean [221]. Ultrasound devices can produce acoustic cavitation, releasing of reactive radicals promoting chemical reactions [222]. The physical effects of ultrasound induced by acoustic cavitation can boost the disruption of oil droplets, facilitating the formation of stable O/W [223]. High power ultrasound has mechanical, chemical and/or biochemical effects, which are used to modify the and enhance the quality of various food systems during processing [224].

VII. ADVANTAGES OF NANO EMULSIONS [225 - 230]

Nanoemulsions have a number of advantages due to the small size of their droplets (100 nanometers), most notably low cost, transparent appearance, no clumping or settling, and a large surface area. Non-toxic and non-irritating, therefore it can be taken via the digestive system. It has a superior ability to improve the solubility of hydrophobic substances, it can also be given topically, intravenously, by inhalation, nasally, intramuscularly, or orally, the metallic and bitter flavors of medicines that may cause unwanted side effects such as nausea and vomiting can be successfully masked using nanoemulsifiers. Nanoemulsifier formulations protect the active ingredients from degradation, oxidation, or reactions with external factors, such as light or heat, and the possibility of altering their rheological properties.

VIII. DISADVANTAGES OF NANO EMULSIONS [231- 237]

The stability of nanoemulsions is affected by environmental factors such as temperature and pH, and these factors change when nanoemulsions are given to patients, the homogenization apparatus (the tool needed to prepare nanoemulsions) is also an expensive process, and Ostwald maturation is the main problem associated with the non-acceptance of nanoemulsion formulations. Additionally, stabilizing nanodroplets requires the use of a high concentration of surfactants and auxiliary materials.

IX. APPLICATIONS OF NANO EMULSIONS [238-244]

Nanoemulsions appear highly attractive from an industrial perspective, given their ease of large-scale production and the absence of complex manufacturing techniques. Nanoemulsions have potential applications in the food industry for delivering pharmaceutical nutrients, coloring and flavoring agents, and antibiotics. Delivery systems made from smart materials with adjustable physical and biological properties also enhance current treatment methods. The development of nanosystems has shown a significant impact in many agricultural fields, including pesticide formulations, food coatings, fungicides, and antiparasitics. Recently, nanoemulsions have gained increasing importance as potential means of delivering cosmetics in a controlled manner, and of optimally distributing active ingredients in different layers of the skin.

X. RECOMMENDATIONS

The most important recommendations for developing nanoemulsions include improving their physical stability by using surfactants (non-toxic, safe, environmentally friendly), adopting high-energy production methods (ultrasonic homogenization), integrating artificial intelligence to improve formulations, and applying packaging techniques (drug loading and release), using Metal-organic frameworks in the preparation of crystalline nanocomposites because they have a large surface area, and have high porosity.

XI. CONCLUSION

To date Nano emulsions have been shown to be able to protect labile drug, control drug release, increase drug solubility,. The main thing is that it contain small particle size reduced to the nanometric scale shows various physical properties. Nano emulsion can be used in various disciplines including cosmetics, biotechnology, food, agriculture and, and pharmaceuticals.

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