Arti S. Rathod et al



Available online at: <u>http://www.iajps.com</u>

Review Article

A REVIEW ON TECHNIQUES FOR FORMULATION OF NANOEMULSION DRUG DELIVERY SYSTEM

Arti S. Rathod¹, Gayatri V. Apothikar², Sakshi Y. Pathrikar³, Vinayak A. Katekar⁴, Dr. Swati P. Deshmukh⁵

^{1,2,3}Department of Pharmacy, Shraddha Institute of Pharmacy, Washim (444505)
⁴Department of Quality Asurance, Shraddha Institute of Pharmacy, Washim
⁵Department of Pharmacology, Shraddha Institute of Pharmacy, Washim

Abstract: -

Nanoemulsions, characterized by their nanoscale droplet size and enhanced stability, have emerged as a promising technology with diverse applications. This abstract provides a concise overview of Nanoemulsions, emphasizing their unique properties and applications across pharmaceuticals, food and beverages, cosmetics, agriculture, and more. The small droplet size of Nanoemulsions contributes to improved bioavailability and targeted drug delivery, particularly beneficial for poorly water-soluble drugs. Their versatility extends to food formulations, where Nanoemulsions enhance flavor, color, and nutrient delivery. In the cosmetic industry, these emulsions improve the absorption of active ingredients in skincare products. Additionally, Nanoemulsions find utility in agriculture for the formulation of pesticides, offering enhanced stability and efficacy. While nanoemulsions present numerous advantages, considerations such as toxicity, biocompatibility, and environmental impact warrant attention. Ongoing research aims to optimize formulation science, providing solutions to challenges in drug delivery, food technology, and beyond.

Keywords- Microfluidization, Emulsifier, Ultrasonic emulsification method etc.

Corresponding author:

Arti S. Rathod,

Department of Pharmacy, Shraddha Institute of Pharmacy, Washim (444505)



Please cite this article in press Arti S. Rathod et al., A Review On Techniques For Formulation Of Nanoemulsion Drug Delivery System, Indo Am. J. P. Sci, 2024; 11 (01).

IAJPS 2024, 11 (01), 145-154

INTRODUCTION:

Nanoemulsions are emulsions with droplet size on the order of100 nm. A typical nanoemulsion contains oil, water and anemulsifier. The addition of an emulsifier is critical for thecreation of small sized droplets as it decreases the interfacialtension i.e., the surface energy per unit area, between the oiland water phases of the emulsion. The emulsifier also plays arole in stabilizing nanoemulsions through repulsive electro-static interactions and steric hindrance.



Fig. O/W Nanoemulsion

The emulsifier used isgenerally a surfactant, but proteins and lipids have also beeneffective in the preparation of nanoemulsions.2–12Over the pastdecade or more, the research focus has been on preparing nano-emulsions through various methods, broadly classified into twoprimary categories: high-energy and low-energy methods.13–15High energy methods such as high pressure homogenization (HPH) and ultrasonication15consume significant energy(B108–1010Wkg to make small droplet.

Advantages Of Nanoemulsion

- 1) **Transparent Appearance:** Nanoemulsions often have a transparent or translucent appearance, making them aesthetically appealing for use in products like clear beverages, skincare formulations, and other transparent applications.
- 2) Efficient Delivery Systems: Nanoemulsions can be tailored to encapsulate and deliver specific active ingredients, such as drugs, flavors, or nutrients, with precision. This targeted delivery can improve the overall efficacy of the incorporated substances
- 3) **Versatility:** Nanoemulsions find applications in various industries, including pharmaceuticals, food and beverages, cosmetics, and agriculture. Their versatility stems from the ability to customize formulations for specific needs.
- 4) **Improved Penetration and Absorption:** In skincare products and pharmaceuticals, Nanoemulsions facilitate better penetration of

active ingredients into the skin or biological tissues, enhancing their therapeutic effects.

- 5) **Reduced Dosage Requirements:** The increased bioavailability and efficient delivery of substances in Nanoemulsions may lead to reduced dosage requirements for drugs or other active compounds, minimizing potential side effects.
- 6) **Ease of Manufacturing:** Nanoemulsions can be produced using relatively simple manufacturing processes, contributing to their scalability and feasibility for large-scale production.
- 7) **Potential Cost Savings:** The improved efficacy and reduced dosage requirements associated with Nanoemulsions may result in cost savings, especially in pharmaceutical and agricultural applications.

Disadvantage Of Nanoemulsion

While Nanoemulsions offer numerous advantages, they also come with certain challenges and potential disadvantages:

- 1) **Cost of Production:** The manufacturing processes involved in creating Nanoemulsions can sometimes be more intricate and may require specialized equipment, which could contribute to higher production costs compared to conventional emulsions.
- 2) Potential Instability: Despite enhanced stability, Nanoemulsions may still be susceptible to destabilization over time, leading to issues such as phase separation, coalescence, or Ostwald ripening. Careful formulation and storage conditions are necessary to mitigate these risks
- 3) **Surfactant Dependency:** Nanoemulsions often rely on surfactants or other stabilizing agents. High surfactant concentrations may be required, and in some cases, these surfactants can affect the taste, appearance, or overall acceptability of the final product.
- 4) Limited Loading Capacity: The small droplet size in Nanoemulsions may limit the amount of active ingredient that can be loaded, potentially impacting the delivery of higher doses in certain applications.
- 5) **Biocompatibility Concerns:** In pharmaceutical and biomedical applications, concerns may arise regarding the biocompatibility of the materials used in the formulation Nanoemulsions. The impact on biological tissues or long-term effects should be thoroughly studied.
- 6) **Environmental Impact:** The production and disposal of Nanoemulsions-containing products could raise environmental concerns, especially if the Nano-sized droplets or associated components pose risks to ecosystem.

Arti S. Rathod et al

- 7) **Regulatory Challenges:** As Nanoemulsions are a relatively advanced technology, regulatory frameworks may not be fully established or standardized across all industries. Meeting regulatory requirements and ensuring product safety can present challenges.
- 8) **Scale-Up Issues:** Transitioning from laboratoryscale to large-scale production may pose challenges in maintaining the desired properties of Nanoemulsions. Achieving uniformity and stability on a larger scale can be complex.
- 9) Limited Understanding: Despite significant progress, the complete understanding of the longterm effects and interactions of Nanoemulsions within the human body or the environment is an ongoing area of research.
- 10) **Potential for Unintended Consequences:** The unique properties of Nanoemulsions may lead to unforeseen consequences, such as altered drug release profiles or unexpected changes in product characteristics, requiring careful consideration during formulations.

> FACTOR AFFECTING THE NANOEMULSION –

Several factors influence the formation and stability of Nanoemulsions. These factors need careful consideration during the formulation process to achieve the desired properties. Some key factors affecting Nanoemulsions include:

- Surfactant Type and Concentration: The choice of surfactant or emulsifying agent, as well as its concentration, plays a crucial role in stabilizing Nanoemulsions. The type of surfactant affects the interfacial tension and determines the stability of the droplets.
- 2) Oil and Water Phases: The nature of the oil and water phases used in the formulation impacts the physical and chemical properties of the Nanoemulsions. The selection of suitable oils and water phases is essential for achieving the desired characteristics.
- 3) Droplet Size and Distribution: The target droplet size and size distribution are critical parameters influencing the stability and performance of Nanoemulsions. Controlling these factors is essential for applications such as drug delivery or improving the sensory attributes of food and cosmetic products.
- Temperature: The temperature during the formulation process can affect the viscosity, solubility, and stability of Nanoemulsions. Some formulations may require specific

temperature conditions for optimal production

- 5) Energy Input: Techniques used for emulsification, such as high-pressure homogenization, ultra sonication, or micro fluidization, determine the energy input during the formulation process. The choice of method can influence the droplet size and overall stability.
- 6) pH Levels: The pH of the Nanoemulsions formulation can impact the stability of the emulsion system. It is crucial to consider the compatibility of the chosen components with the desired pH range.
- 7) Surfactant-to-Oil Ratio: Achieving an appropriate balance between the surfactant and oil phases is essential for stabilizing Nanoemulsions. An excess or deficiency of surfactant can affect the emulsion's stability and performance.
- 8) Ionic Strength: The presence of ions in the formulation, often from salts or other additives, can influence the stability of Nanoemulsions. Ionic strength can affect the electrical double layer around droplets, impacting their stability.
- 9) Presence of Cosolvents: The inclusion of Cosolvents or co-surfactants can influence the solubility of components and the overall stability of the Nanoemulsions. Careful selection and control of these additives are necessary
- 10) Processing Conditions: The specific conditions during the emulsification process, such as shear rate, mixing time, and homogenization pressure, can significantly affect the Nano emulsion's characteristics. Optimal processing conditions are crucial for reproducibility

METHOD OF PREPRATION

Nanoemulsions 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. Production of Nanoemulsions is achieved by using the intrinsic physiological Properties of the system. In this Nanoemulsions preparation method, stored energy of the system is utilized by Alteration of parameters such as temperature, composition of the system (Satya et al., 2014). At the initial Studies of Nanoemulsions, the high energy methods were only choice for researches and thus high-energy Stirring and ultrasonic emulsification were the most widely used methods. Nowadays, low-energy methods Have drawn considerable attention since they are 'soft'. nondestructive and cause no damage to encapsulated Molecules Several methods have been suggested for the preparation of Nanoemulsions. The basic objectives of the Nano emulsions preparation to achieve the droplet size range of 100-600 nm and another is to provide the Stability condition. Formation of Nanoemulsions system required a high amount of energy. This energy can be Provided either by mechanical equipment or the chemical potential inherent within the component. Here some Methods are discussed which is freely used for the Nanoemulsions preparation

High-Energy Emulsification Methods

Nanoemulsions are non-equilibrium systems which cannot be formed spontaneously. For this reason, Mechanical or chemical energy input is necessary to form them. Nanoemulsions are generally prepared by Using high energy methods in which mechanical energy input is applied by high pressure homogenizers, High shear stirring, and ultrasound generators (Sole et al., 2012). These mechanical devices provide strong Forces that disrupt oil and water phases to form Nanoemulsions. In high energy methods, input energy density is about 108 -1010 W kg-1 (Gupta et al., 2016).

Required energy is supplied in a shortest time to the system in order to obtain homogeneous small sized Particles. High-pressure homogenizers are capable of doing this and therefore they are the most widely used Devices for preparing nanoemulsions (Solons et al, 2005). Moreover, producing emulsions using ultrasound is a Cost-effective process which needs less surfactant use (Katas et al., 2013). Therefore, considering conventional Mechanical processes more homogeneous batches are achieved.

High Pressure Homogenization

It is the most popular method used for the production of nanoemulsions. This method benefits from the High-pressure homogenizer or the piston homogenizer to manufacture nanoemulsions that particle sizes are up To 1 nm. During the method, the macro emulsion is forced to pass through in a small orifice at an operating Pressure between 500 to 5000 psi. Extremely small droplet sized nanoemulsions are achieved because during The process several forces like hydraulic shear, intense turbulence and cavitation act together. This process can Be repeated until the final product reaches the desired droplet size and polydispersity index (PDI). The Uniformity of droplet size in nanoemulsions is specified by PDI. Higher PDI means lower uniformity of droplet Size in nanoemulsions. Nondispersive samples have PDI lower than 0.08, PDI between 0.08 and 0.3 states a Narrow size distribution, whereas PDI greater than 0.3 indicates broad size distribution. However, obtaining of Small droplets that are in submicron levels requires large amount of energy (Lovelyn&Attama, 2010). This Amount of energy and increasing temperatures during high pressure homogenization process might cause Deterioration of the components. Terminable compounds such as proteins, enzymes and nucleic acids may be damage.

High-Shear Stirring

In this method, high-energy mixers and rotor-stator systems are used for the preparation of Nanoemulsions. Droplet sizes of the internal phase can be significantly decreased by increasing the mixing Intensity of these devices. However, obtaining emulsions with the average droplet size less than 200-300 nm is Rather difficult.

Ultrasonic Emulsification

There are two mechanisms which take part in ultrasonic emulsification. Firstly, acoustic field creates Interfacial waves that makes oil phase to disperse in the continuous phase as droplets. Secondly, ultrasoundProvokes acoustic cavitation which provides formation and collapse of microbubbles respectively due to Pressure fluctuations of a single sound wave. In this way, enormous levels of highly localized turbulence is Generated and this causes micro implosions which disrupt large droplets into sub-micron size (Zhang, 2011). In

This method, premixed macro emulsion is agitated by vibrating solid surface at 29 kHz or larger frequencies. High-power ultrasonic devices such as focusing horns and pointed tips cause extreme shear and cavitation that Result in breaking up of droplets. It has been observed that in most of the ultrasonic systems emitted sound field Is inhomogeneous. For this reason, in order to have all droplets to experience highest shear rate, recirculation Of the emulsion through the region of high power must be provided. Moreover, by doing this type of Recirculation many times it is possible to obtain emulsions with uniform droplet size at dilute concentrations. Emulsifier type, the amount emulsifier, and viscosity of phases are the most critical parameters that affect Homogenization efficiency. Thus, optimization of these parameters is necessary to prepare nanoemulsions Having fine droplets. However, there are some concerns about sonication Methods due to fact that they have possibility to induce protein denaturation, polysaccharide depolymerization and liquid oxidation.

> Micro fluidization

It is most widely employed in the pharmaceutical industry in order to acquire fine emulsions. In this Method, a device called micro fluidizer is used which provides high pressures. During the process, high Pressure forces the macro emulsion to go through to the interaction chamber and thus nanoemulsions with Submicron ranged particles can be produced. Uniform nanoemulsions production can be achieved by repeating

The process many times and varying the operating pressure in order to get desired particle size. There is a Collision between crude emulsion jets from two opposite channels in the nozzle of microfilariae which is also Called as the interaction chamber. The mobility of crude emulsion is provided by a pneumatically powered Pump that has capability of compressing air up to pressures between 150 to 650 MPa. This high pressure forces

The crude emulsion stream to go through microchannel and after the Collison of two opposite channels Enormous level of shearing force is obtained. Therefore, by the help of this force fine emulsions are produced.

Low-Energy Emulsification Methods

Nanomulsification can also be achieved with lowenergymethods which provides small size and more Uniform droplets. These methods such as phase inversion temperature and phase inversion component provide Smaller and more uniform droplets by using physicochemical properties of the system (Caldero et al., 2011).

Although low energy procedures are generally more effective to produce small droplet sizes than high energy Procedures, there are someLimitations for them about the using of some types of oils and emulsifiers like proteins and polysaccharides. In Order to overcome this problem high level of synthetic surfactant concentrations are used to produce Nanoemulsions in low energy techniques but this narrows down their application area, especially for many food process.

Spontaneous Nanomulsification

It benefits from the chemical energy replacement based upon dilution process with the continuous phase Which occurs usually at constant temperature without

any phase transitions in the system during the Emulsification process. This method can produce nanoemulsions at room temperatures and no special devices Are required. It basically subjected to interfacial tension, viscosity of interfacial and bulk, phase transition Region, surfactant structure, and surfactant concentration. In the pharmaceutical industry, systems prepared by Using this method are usually called as self emulsifying drug-delivery systems (SEDDS) or self nanoemulsifying drugdelivery systems (SNEDDS). When an oil phase with a water soluble substance is mixed With water, oil droplets spontaneously forms. The mechanism depends on the movement of water dispersible Substance from the oil phase to the water phase, indicated as red arrows in Figure 3. This leads to interfacial Turbulence and thus formation of spontaneous oil droplets.

> Phase Inversion Methods

These methods utilize the chemical energy that is released because of the phase transitions during Emulsification process. Required amount of phase transitions are achieved by changing the composition at Constant temperature or by changing the temperature at constant composition.

Phase Inversion Temperature (PIC)

In this method, temperature is changed at constant composition. Non-ionic surfactants which have temperature dependent solubility like polyethoxylated surfactants play important role. Emulsification is Achieved by modifying affinities of surfactants for water and oil as a function of temperature (Lovelyn&Attama, 2010; Chime et al., 2014). During heating of polyethoxylated surfactants they become Lipophilic due to dehydration of polyoxyethylene groups. Therefore, this circumstance establishes the principle Of producing nanoemulsions by PIT method. In order to prepare nanoemulsions by using PIT method, it is Necessary to bring sample temperature to its PIT level or hydrophilic-lipophilic balance (HLB) level (Anandharamakrishnan, 2014). In the PIT method, the droplet sizes and the interfacial tensions reach their Minimum value. This method promotes emulsification by benefiting from the extremely low interfacial Tensions at the HLB temperature. Nevertheless, it has been observed that although emulsification is Spontaneous at the HLB temperature, coalescence rate is greatly fast and emulsions are highly unstable. It has Been reported that stable and fine emulsion droplets can be produced by rapid cooling of the emulsion near the Temperature of PIT.

Phase Inversion Composition (PIC)

In obtained by Consistently adding water or oil to the mixture of oil surfactant or water-surfactant. The PIC method is more Suitable for a large scale production than the PIT method since adding one component to an emulsion is easier Than to generate abrupt change in temperature (Sloan's& Sole, 2012). By adding water to the system, volume of Water increases and this result to reach a transition composition. In other words. the level of hydration of the Thepolyoxyethylene chains of the surfactant increases and thus spontaneous curvature of the surfactant goes to A change from negative to zero. As in the HLB temperature, in the transition composition a balance is obtained for the surfactant hydrophilic-lipophilic properties. When this transition composition is exceeded, small sized Metastable oil in water droplet are composed due to the separation of the structures that have zero curvature.

Solvent Displacement Method

This method of nanoemulsions has been adopted from the Nano precipitation method used for Polymeric nanoparticles. In this the oily phase is dissolved in water-miscible organic solvents like ethanol And acetone. Then this organic solvent is poured into aqueous phase containing surfactant which leads to the Formation of nanoemulsions by the rapid diffusion of organic solvent.this method, composition is changed at constant temperature.

Classification Of Nanoemulsion –

Nanoemulsions can be classified based on various characteristics, including their composition, structure, and application. Here are common classifications:

1) Based on Composition:

Oil-in-Water (O/W) Nano emulsion: The dispersed phase consists of oil droplets within a continuous water phase. This type is common in pharmaceutical and cosmetic applications.

Water-in-Oil (W/O) Nano emulsions: The dispersed phase consists of water droplets within a continuous oil phase. This type is used in certain food and pharmaceutical formulations

2) Based on Structure:

a) **Biphasic Nanoemulsions:** Consisting of two immiscible phases, typically oil and water, stabilized by surfactants. These can be O/W or W/O, depending on the external phase.

- b) Multiple Nanoemulsions (Multiple Emulsions): Nanoemulsions containing more than one dispersed phase, forming complex structures like water-in-oil-in-water (W/O/W) or oil-in-water-in-oil (O/W/O)
- 3) Based on Droplet Size:
- a) **Micro emulsions:** Nanoemulsions with even smaller droplet sizes, often below 20 nanometers. These exhibit thermodynamic stability and unique properties, suitable for specific applications.

4) Based on Application:

- a) **Pharmaceutical Nanoemulsions:** Designed for drug delivery to improve solubility, bioavailability, and controlled release of pharmaceutical compounds.
- b) **Food Nanoemulsions:** Used in the food and beverage industry for applications such as flavor encapsulation, nutrient delivery, and improving the stability of certain products.
- c) Cosmetic Nanoemulsions: Applied in skincare and personal care products for better delivery of active ingredients, improved texture, and enhanced sensory properties.

5) Based on Manufacturing Method:

- a) **High-Energy Methods:** Produced using techniques such as high-pressure homogenization, ultra sonication, or Microfluidization, which provide the energy needed for small droplet formation.
- b) **Low-Energy Methods:** Created using lowenergy methods such as phase inversion temperature or phase inversion composition, which involve phase transitions for emulsion formation.

6) Based on Charge:

Cationic Nanoemulsions: Containing positively charged droplets, often used in applications where interaction with negatively charged surfaces or tissues is desired. Anionic Nanoemulsions: Containing negatively charged droplets, suitable for specific applications where electrostatic interactions play a role.

Understanding these classifications helps researchers and formulators choose the appropriate nanoemulsions type based on the intended use and desired properties. Each classification has its advantages and is tailored to meet specific application requirements.

Component Of Nanoemulsion – Nanoemulsions, twicelly, consist, of the

Nanoemulsions typically consist of three main components:

- 1) Dispersed Phase (Oil Phase): This is the internal phase of the nanoemulsions and comprises the oil or lipophilic component. The choice of oil depends on the application and desired characteristics. Common oils medium-chain include triglycerides, triglycerides (MCTs), essential oils, or other lipids.
- Continuous Phase (Water Phase): This is 2) the external phase surrounding the dispersed oil droplets and consists of water or another hydrophilic liquid. The selection of the continuous phase depends on the application, and it may include water, aqueous solutions, or other polar solvents.
- Surfactants/Emulsifiers: Surfactants play a 3) crucial role in stabilizing nanoemulsions by reducing the interfacial tension between the oil and water phases. They help prevent droplet coalescence and maintain the small droplet size. Emulsifiers are a subset of surfactants specifically used in emulsion formulations.
- 4) Cosolvents/Co-Surfactants: In some formulations, Cosolvents or co-surfactants may be added to enhance solubility, stability, the properties of modify or the nanoemulsions. These components can include alcohols, glycols, or other amphiphilic molecules.
- 5) Antioxidants/Preservatives: Depending on the intended application and stability requirements, antioxidants or preservatives may be added to prevent oxidation or microbial growth and extend the shelf life of the nanoemulsions.
- 6) Active Ingredients: In pharmaceutical, food applications, cosmetic, or nanoemulsions may encapsulate active ingredients such as drugs, flavors, fragrances, or nutrients. The small droplet size enhances the bioavailability and efficacy of these active components.
- 7) pH Adjusting Agents: In certain applications, pH adjusting agents may be used to maintain the desired pH level for stability and compatibility with specific formulations.
- 8) Thickeners/Rheology Modifiers: These components may be added to adjust the viscosity and improve the texture of the nanoemulsions. Thickeners can enhance the stability and sensory attributes in cosmetic or food formulations.

Markated Nanoemulsion Formulation - \triangleright

1. Pharmaceuticals:

Annexes: A nanoemulsions formulation of paclitaxel, a chemotherapy drug. Nanoemulsions enhances the solubility of paclitaxel, improving its bioavailability and potential effectiveness in cancer treatment.

2. Food and Beverages:

Nano emulsified CBD Products: Nanoemulsions technology is used to enhance the bioavailability of conidial (CBD) in various food and beverage products, such as CBD-infused drinks and edibles 3. Cosmetics and Skincare:

Nanoemulsions-Based Sunscreens: Some sunscreen formulations use nanoemulsions technology to create transparent and non-greasy sunscreens with improved UV protection.

4. Agriculture:

Nanoemulsions Pesticide Formulations: Nanoemulsions are employed in the agricultural sector for the development of pesticide formulations, improving the delivery and effectiveness of active ingredients.

5. Nutraceuticals:

Nano emulsified Nutrient Supplements: Nanoemulsions are used in the formulation of nutrient supplements, enhancing the absorption of fat-soluble vitamins and other bioactive compounds

It's crucial to note that the availability of nanoemulsions formulations can vary by region, and new products may have been introduced since my last update. Additionally, the regulatory landscape and consumer preferences can influence the types of nanoemulsions-based products that are commercially available. Always refer to current sources or consult with industry professionals for the latest information on market

Pharmacological Action Of Nanoemulsion - \geq

Nanoemulsions, owing to their unique properties, exhibit various pharmacological actions that make them attractive for pharmaceutical applications. Some key pharmacological actions include

Improved Bioavailability: Nanoemulsions enhance the bioavailability of poorly water-soluble drugs by reducing particle size, increasing the surface area available for absorption. This leads to improved and more efficient drug uptake in the body.

Enhanced Drug Solubility: The small droplet size in nanoemulsions provides a larger interfacial area, improving the solubility of lipophilic (fat-soluble) drugs. This is particularly beneficial for drugs with low aqueous solubility.

- 1) Targeted Drug Delivery: Nanoemulsions can be designed for targeted drug delivery, allowing the controlled release of drugs at specific sites in the body. This can enhance therapeutic efficacy while minimizing side effects.
- 2) Prolonged Drug Release: Nanoemulsions can be engineered to achieve sustained or prolonged drug release, leading to a more controlled and consistent therapeutic effect over an extended period.
- Intravenous Administration: Nanoemulsions are well-suited for intravenous administration, providing a stable and homogeneous drug delivery system. This is advantageous in the treatment of various medical conditions, including cancer
- 4) CNS Drug Delivery: Nanoemulsions can overcome the blood-brain barrier, facilitating the delivery of drugs to the central nervous system (CNS). This is significant for treating neurological disorders where conventional drug delivery methods may be less effective.
- 5) Ocular Drug Delivery: Nanoemulsions can be formulated for ophthalmic applications, allowing improved drug penetration into ocular tissues. This is relevant for the treatment of eye disease
- 6) Anti-Inflammatory Effects: Nanoemulsions can be designed to carry anti-inflammatory drugs, providing targeted delivery to inflamed tissues. This can enhance the therapeutic effects while minimizing systemic side effects.
- Antimicrobial Actions: Nanoemulsions have demonstrated antimicrobial properties and can be utilized for delivering antimicrobial agents. This is relevant in the treatment of infections.
- Vaccine Delivery: Nanoemulsions can serve as carriers for vaccine antigens, enhancing the immune response. They may improve vaccine stability and provide controlled release for sustained immunity.

It's Important to note that the pharmacological actions of nanoemulsions can vary based on the specific formulation, drug characteristics, and targeted application. Researchers continue to explore and optimize nanoemulsions formulations to improve drug delivery efficiency and therapeutic outcomes in various medical fields.

Uses Of Nanoemulsion –

Nanoemulsions find diverse applications across several industries due to their unique properties. Here are some notable uses of nanoemulsions:

1) Pharmaceuticals:

Drug Delivery: Nanoemulsions enhance the bioavailability of poorly water-soluble drugs, enabling more efficient drug delivery. They are used for oral, intravenous, transdermal, and ocular drug administration

2) Food and Beverages:

Improved Flavor and Nutrient Delivery: Nanoemulsions are used to encapsulate flavors, colors, and nutrients, improving their stability and delivery in food and beverage products. Fat Replacement: Nanoemulsions can serve as fat substitutes, providing a creamy texture without the need for traditional fats. This is useful in low-fat or fat-free food products.

- Cosmetics and Personal Care: Skincare Products:Nanoemulsions in cosmetics enhance the delivery of active ingredients, such as vitamins, antioxidants, and moisturizers, improving their absorption into the skin. Sunscreen Formulations: Nanoemulsions contribute to the development of transparent and non-greasy sunscreens with improved UV protection.
- 4) Agriculture:

Pesticide Delivery: Nanoemulsions are used in agriculture for the formulation of pesticides, enabling better coverage, increased stability, and improved efficacy.

5) Textile Industry:

Fabric Finishing: Nanoemulsions can be applied for fabric finishing, providing water and stain repellency to textiles.

6) Petroleum Industry:

Enhanced Oil Recovery:Nanoemulsions are explored for applications in enhanced oil recovery, helping to improve the efficiency of oil extraction.

Cleaning Products:

Cleaning Solutions:*Nanoemulsions are used in cleaning products to enhance the solubilization of oils and improve the overall cleaning efficiency.

7) Inks and Coatings:

Printing Inks: Nanoemulsions are employed in the formulation of inks, providing improved stability and color dispersion.

Coatings: Nanoemulsions contribute to coatings with enhanced properties, such as increased adhesion and durability.

8) Biomedical Applications:

Diagnostic Imaging:** Nanoemulsions can be used as contrast agents in diagnostic imaging techniques, providing better resolution and targeting capabilities.

9) Vaccines:

Vaccine Adjuvants: Nanoemulsions are explored as adjuvants in vaccine formulations, improving the stability and efficacy of vaccines.

The versatility of nanoemulsions arises from their ability to encapsulate, solubilize, and deliver various substances, making them valuable in multiple industries. Ongoing research continues to uncover new and innovative applications for nanoemulsions technology.

Adverse Effects Of Nanoemulsion –

While nanoemulsions offer many advantages, it's important to consider potential adverse effects, especially in the context of certain applications. Some concerns associated with nanoemulsions include:

- Toxicity and Biocompatibility: The materials used in the formulation of nanoemulsions, including surfactants and other components, may raise concerns about potential toxicity. Biocompatibility and long-term effects on biological tissues need careful consideration.
- Immunogenicity: Certain nanoemulsions formulations may trigger immune responses, and the impact of repeated exposure or chronic use should be thoroughly investigated, especially in pharmaceutical and medical applications.
- Environmental Impact: The disposal of nanoemulsions-containing products may raise environmental concerns. The persistence and potential Eco toxicity of Nano-sized particles require attention, particularly in formulations used in agriculture or other environmental settings.
- a. Allergic Reactions: Some individuals may be sensitive or allergic to specific components used in nanoemulsions, such as surfactants. It's essential to assess the potential for allergic reactions, especially in cosmetic and skincare products.
- b. Interaction with Biological Barriers: Nanoemulsions designed for drug delivery, particularly those targeting the central nervous system or crossing biological barriers, may raise concerns about unintended interactions or effects on these barriers

Incompatibility with Certain Drugs: The encapsulation of drugs within nanoemulsions may alter drug stability, release profiles, or interactions with the body, potentially leading to unintended consequences or side effects.

Accumulation in Organs: Concerns have been raised about the potential for the accumulation of Nano-sized particles in organs, particularly if used in pharmaceuticals over extended periods. The long-term effects of such accumulation need thorough investigation.

- c. Skin Irritation: In cosmetic and skincare formulations, nanoemulsions may pose a risk of skin irritation, especially if the formulation contains components that are known to cause skin sensitivities.
- d. Potential for Agglomeration: Despite efforts to maintain stability, nanoemulsions may undergo agglomeration or coalescence over time, leading to changes in particle size distribution and potential loss of desired properties.

It's important to note that the adverse effects mentioned above are dependent on factors such as the specific formulation, components used, application, and the intended use of the nanoemulsions. Rigorous testing, safety assessments, and regulatory compliance are crucial in mitigating these concerns and ensuring the safe use of nanoemulsions in various industries. Researchers and manufacturers continue to explore ways to address these challenges and optimize nanoemulsions formulations for safety and efficac

CONCLUSION:

In conclusion, nanoemulsions represent a versatile and promising technology with wide-ranging applications across pharmaceuticals, food and beverages, cosmetics, agriculture, and more. Their unique properties, including small droplet size and enhanced stability, contribute to improved bioavailability, targeted drug delivery, and efficient encapsulation of active ingredients. Nanoemulsions have demonstrated significant advancements in drug delivery systems, providing solutions for poorly water-soluble drugs and enabling controlled release profiles.

While nanoemulsions offer numerous benefits, it is essential to acknowledge and address potential challenges, such as concerns about toxicity, biocompatibility, and environmental impact. Ongoing research and development aim to optimize formulations, ensuring safety and efficacy in various applications.

As the field continues to evolve, nanoemulsions hold promise for addressing complex challenges in different industries. Their ability to enhance solubility, stability, and controlled release makes them a valuable tool for formulators and researchers seeking innovative solutions. However, careful consideration of regulatory frameworks, biocompatibility, and longterm effects remains crucial to unlocking the full potential of nanoemulsions technology. Overall, nanoemulsions stand at the forefront of advancements in formulation science, offering solutions to improve the delivery and performance of a wide array of substances in diverse applications.

REFERANCES:

- Bhosale RR, Omani RA, Ghodake PP, Shaikh SM, Chavan SR. 2014 Nanoemulsions: A Review on Novel profusion in advanced drug delivery. Indian J Pharm Biol Res;2:122-7.
- Borate V, Pathak S, Sharma S, Patravale V. 2012 Clotrimazolenanoemulsion for malaria Chemotherapy. Part I: Preformulation studies, formulation design and physicochemicalevaluation. Int J Pharm;431:138-48.
- Caldero, G., Maria, J.G.C. and Sloan's, C., 2011Formation of polymeric nano-emulsions by a Low energy method and their use for nanoparticle preparation, Journal of Colloid and Interface Science, 353, 406–411.
- 4) Chen H, Hu X, Chen E, Wu S, McClements DJ, Liu S, et al. 2016 Preparation, characterization, and Properties of chitosan films with cinnamaldehydenanoemulsions. Food Hydrocoll 61:662-71.
- Chime, S.A., Kenechukwu, F.C., and Attama, A.A.2014, Nanoemulsions-Advances in Formulation, Characterization and Applications in Drug Delivery, Ali DS, Application of Nanotechnology in Drug Delivery, Crotia: InTech, 77-111,
- 6) Chouksey R, Jain AK, Pandey H, Maithil, 2011 A. In vivo assessment of atorvastatin nanoemulsions .Formulation. Bull Pharm Res;1:10-4.
- 7) Đorđević SM, Cekić ND, Savić MM, Isailović TM, Ranđelović DV, Marković BD, et al.2015 Parenteral nanoemulsions as promising carriers for brain delivery of risperidone: Design, Characterization and in vivo pharmacokinetic evaluation. Int J Pharm;493:40-54.
- Fernandez, P., Andre, V., Rigger, J. and Kühnle A., 2004 Nano-emulsion formation byemulsion phase Inversion, Colloids and Surfaces A: Physicochem. Eng. Aspects, 251, 53–58.
- 9) Floury, J., Desrumaux, A., and Larders', J., 2000 Effect of high-pressure homogenization on droplet Size distributions and rheological properties of model oil-in-water emulsions, Innovative Food Science & Emerging Technologies, 1(2), 127-134.

- Ghosh V, Mukherjee A, Chandrasekaran N. 2013 Ultrasonic emulsification of food-grade Nanoemulsions formulation and evaluation of its bactericidal activity. UltrasonSonochem;20:338-44.
- Gupta, P.K., Pandit, J.K., Kumar, A., Swaroop, P., and Gupta, S., 2010 Pharmaceutical Nanotechnology Novel Nanoemulsions–High Energy EPreparation, Evaluation and Application, ThePharma Research, 3, 117-138.
- 12) Harika K, Debnath S, Babu MN. 2015 Formulation and evaluation of nanoemulsions of amphotericin B. IJNTPS;5:114-22
- 13) J Ahmad Farhan, Ali Mushir, Shekel Faiyaz, Talegaonkar Cushman, K KharRoop, Shafiq Sheikh. 2008 Investigation of Nanoemulsions System for Transdermal Delivery of Domperidone: Ex-vivo and in vivo Studies. Current Nanoscience, 382-639.