

NANOTECHNOLOGY: AN AEON IN THE FORMULATION DEVELOPMENT

MR. KSHATRIYA PRAVIN JAMNADAS

Student of M Pharmacy Pharmaceutical Department, PDEA's S.G.R.S. College of Pharmacy, Saswad.

MR. SHINDE JITENDRA V.

Asso. Prof. Department of Pharmaceutics, PDEA's S.G.R.S. College of Pharmacy, Saswad.

MR. MUNDE VIJAY PREMCHAND

Student of M Pharmacy Pharmaceutical Department, PDEA's S.G.R.S. College of Pharmacy, Saswad.

ABSTRACT

Solid lipid Nanoparticles (SLN) are unit at the rising stage of the quickest developing field of engineering with several helpful application in drug delivery and analysis. Owing to their distinctive nano size properties, solid lipid Nanoparticles offer the chance to develop new medicine. The talent to include medication into nanocarriers offers a brand new through in drug delivery that would be used for drug targeting. So solid lipid Nanoparticles hold nice promise for achieving the goal of web site specific drug delivery. This review presents a broad introduction of solid lipid Nanoparticles discussing their arms, procedures for preparation advantages, limitations, application, patents, associated with the sector and marketed merchandise for skin development as solid lipid Nanoparticles.

KYE WORDS:

Engineering , solid lipid Nanoparticles, nano, size, patents.

INTRODUCTION:

According to the NNI (National Nanotechnology Initiative), definition Nanoparticles are the solid mixture having the dimensions in nanometres that ranges from 10-1000nm a minimum of in one dimension (generally 50-500nm). During which the drug is dissolved or entrapped within the chemical compound or lipoidal matrix. 60 Material accustomed formulate Nanoparticles embrace lipid (solid at area temperature), natural chemical compound (chitosan, guar gum, gelatin, bovine albumen. Human albumen, sod. Alginate) artificial chemical compound (poly D lactide, poly-e-caprolactane, polymethylmethacrylate and semisynthetic chemical compound and supermolecule bovine albumen, gelatin). These all material is chosen in line with their encapsulation capability, drug stability, and drug unharness pattern and by their targeting capability.

Nanoparticles is also used as drug carrier; the most benefits of Nanoparticles area unit to extend stability of drug and to the management drug unharness with smart targeting capability. The little sizes of Nanoparticles, offer them distinctive chemistry and biological properties (eg., extremely active expansive and also to cross cell and tissue barriers) that makes it an acceptable candidate for medicine applications. The smaller size of Nanoparticles has the power to avoid phagocyte uptake thanks to that its circulatory lifetime of the drug is increased and also the drug free in sustained manner from a slow geological process chemical compound it, an acceptable quantity of drug will be delivered to the actual cell or target web site by active or passive drug absorption mechanisms. Within the active

targeting technique, ligands will attach with drug nano carrier to extend the affinity of Nanoparticles towards the target website, eg., Folic acid, peptide, etc. this may even be achieved by physical stimuli (eg., Temperature, pH, magnesium). On the opposite hand passive targeting captivated with size eg., Neoplasm tissue area unit leaky in nature and increased tube-shapped structure porousness and retention the little size of Nanoparticles favours the passive targeting. In another example the Nanoparticles will penetrate blood brain barrier by mimicking the transport mechanism of lipoproteins thanks to their little size. On planning to its binding web site the drug free from the formulation. Nanoparticles have shown nice potential for the higher drug delivery in each clinical and pharmaceutical analysis trade. As a drug carrier, Nanoparticles have vital benefits opt bioavailability, general stability, high drug loading, long blood circulation time and selective distribution within the organs tissues with longer 0.5 life.

ADVANTAGES OF NANOPARTICLES:

- 1.Stability enhancement of drugs.
- 2.Solubility enhancement
- 3.Drug targeting
- 4.Enhancement the solubility of a poorly soluble drugs.[1]

POLYMERIC NANOPARTICLES:

Polymeric Nanoparticles area unit mixture particles, but 1000nm, created up either by natural or artificial compound. The drug molecules is encircled by compound or distributed or dissolved in a very matrix. Numerous studied area unit current to develop compound Nanoparticles these days for targeting and controlled drug delivery, significantly focusing neoplasm and brain targeting. Polymers area unit the building block of Nanoparticles polymers conjointly having biomedical application. Various polymers are being used for preparation of Nanoparticles. Compound Nanoparticles have many benefits like smart biocompatibility, straightforward style, straightforward preparation and bio-mimetic character straight forward to regulate size, surface properties. It targets the drug on to its meant space at controlled rate. Compound Nanoparticles supply stability to drug molecule like lipid molecule, peptide or polymer from environmental conditions. Nanoparticles conjointly supply sequence delivery, solubility sweetening of water insoluble medicine. The compound cold also be classified into natural, artificial and semi- synthetic compound.

NATURAL POLYMER:

Natural polymers area under obtained from natural sources, they embrace proteins (gelatin, legume and albumin), and polysaccharides (starch, alginate), natural polymer area unit hydrophilic in nature. They have some disadvantages like batch to batch variation, conditional biodegradability and antigenicity. The protection feature assessment of compound is still anticipated. Gelatin, dextran, albumin area unit acceptable compound for canal use, but show immunogenic response with cross linking agents.

SYNTHETIC POLYMER:

Various artificial polymers area unit accustomed prepare Nanoparticles that embrace poly(e caprolactone) poly carboxylic acid poly (isobutyl cyanoacrylate), poly (butyl cyanoacrylate), poly (lactidecoglycolide), poly (methyl methacrylate) and numerous enteric coating compound of edragit compound etc. there area unit numerous ways of preparation of compound Nanoparticles like ionic gelation, emulsification, section coaservation methodology. [2]

METHOD OF PREPARATION

1. Cross linking amphiphilic macro molecules

The amphiphilic macromolecule like gelatin, {albumin, albumen simple lipid molecules} and protein Nanoparticles are often prepared by mistreatment this methodology. The amphiphilic compound is initially dissolved in binary compound solution containing an acceptable wetter to arrange o/w or w/o microemulsion; the microemulsion is then cross linked to make the compound Nanoparticles. The cross linking is completed by two ways. Heat denaturation or by employing an appropriate cross linking agent like glutaraldehyde, methanol.

1. Section separation or desolvation

In this methodology, an aqueous solution of lipid or polypeptide is ready, the drug is dissolved therein and so the section separation or desolvation is done by ever-changing pH scale, ever-changing temperature or addition of counter particle.

2. Phase separation or desolvation by changing Ph

Insulin containing gelatin Nanoparticles were ready by pH scale modification methodology. In which an aqueous solution of gelatin containing Tween 80 was ready initially. The pH scale was adjusted to optimum pH to induce a transparent solution. This solution was then heated at 400C.

3. Phase separation by addition of counter ion

In this methodology the compound is initially solubilized in an aqueous solution containing an acceptable wetter then an aqueous solution of o/w or w/o emulsion is ready. When this cross linking is completed by an acceptable counter particle e.g. Chitosan solution is ready in zero. 5% carboxylic acid aqueous solution. Drug is then distributed or dissolved during this solution and so the section separation is completed by adding TPP sodium counter ions and precipitation takes place because of section separation. The Nanoparticles are then controlled by natural process.

4. Spray drying technique

In this technique the chemical compound solution containing the active is dried on the nice and cozy air. The chemical compound solution is ready by dissolving the compound during a volatile organic liquid system. The solution of compound and drug is then sprayed during a stream of hot air. The dried particles are then collected e.g. chitosan was dissolved in 1% carboxylic acid aqueous solution, then more drop informed poly 9methyl vinyl ether-co-maleic anhydride). The solution is then spray dried exploitation water worker. 1100C outlet worker 490C at 600N1/h airflow rate.

SOLVENT DISPLACEMENT OR NANOPRECIPITATION

In nanoprecipitation technique the compound gets deposited to drug initially so solvent is displaced with solvent different solvent. Two miscible solvents are employed in this technique. The hydrophobic material used that are solubilized in organic solvent like ethylalcohol or ether but has low solubility in water. The compound and drug are dissolved in organic solvent then poured to non-solvent (water) on continuous stirring, that cause a speedy precipitation of compound into formation of Nanoparticles.[2,3]

SOLID LIPID NANOPARTICLES

In 1991 solid lipid Nanoparticles was discovered, solid lipid Nanoparticles is an alternative drug carrier system than chemical compound Nanoparticle. This carrier consists of solid lipid sphere

within the nano size, that are spread in water or in liquid surface-active agents solution. Solid lipid particles are fabricated from solid hydrophobic core having a monolayer of phospholipid coating. The solid core contains the drug either dissolved or spread in the high melting solid fat matrix. The hydrophobic chains of phospholipids are embedded within the fat matrix. They have potential to hold water insoluble medication or diagnostic agents. Oral bioavailability of poorly water soluble drug are often improved by exploitation this approach. Solid lipid nanoparticles have the advantages of over chemical compound Nanoparticles because lipid matrix is formed of physiologically tolerated lipid elements, which decreases the potential for acute and chronic toxicity. It is another drug delivery system to mixture drug delivery systems like lipid emulsions, liposomes and chemical compound nanopartivles. Solid lipid nanoparrticles have the several benefits over completely different mixture carriers and conjointly it avoids many disadvantages of chemical compound nanoparticle, like physical stability, protection of incorporated labile medication from degradation, controlled unharness, and wonderful tolerability. Solid lipid Nanoparticles formulations are often applied for numerous routes like parenteral, oral, dermal, ocular, pulmonary, and rectal. The in- vivo and in-vitro studies have shown have terribly positive results. Solid lipid Nanoparticles ready from lipid than compound. The lipid are solid at temperature. Solid lipid Nanoparticles combine the properties of liposomes like biocompatibility and chemical compound particles stability, higher production potency and the surface solid lipis Nanoparticles will modified for drug targeting by attaching matter or by PEGylation they are ready by blend or emulsion precipitation for delivering medication as a solid molecular dispersion oe as a drug encapsulating lipid shell.

OBJECTIVE OF SOLID LIPID NANOPARTICLES

- 1.For dominant the drug unharness
- 2.To reinforce stability of drug
- 3.To entrap high drug content
- 4.Incorporation of lipotropic and deliquescent medication.

ADVANTAGES

Their tiny size and slim size distribution, facilitate within the permeation through biological membrane, the tiny size conjointly facilitate to focus on the drug into brain lipid are physiological biodegradable that lowered possibilities of the acute chronic toxicity .

- 1.Typically organic solvent isn't employed in the preparation of solid lipid Nanoparticles; generally a really less quantity could also be used.
- 2.SLNs have higher stability compared to liposomes.
- 3.It will increase the bioavailability { of medication| of medicine| of medication } and protects drugs from chemical degradation'
- 4.It are often ready simply.
- 5.It protects, degrading the drug within the dirty dog or from different condition.
- 6.It are often sterilized by standard technique.

7. It are often used for web site specific drug targeting as a result of surface of solid lipid Nanoparticles are often changed simply.

DISADVANTAGES

1. Solid lipid Nanoparticles have restricted drug defense potency thanks to drug solubility within the lipid often.
2. Action of drug could also be determined throughout the storage as a result of excellent crystal is formed that facilitate in action.
3. The particles have gelation tendency and size could also be redoubled.
4. Unsure dynamics of organism transition.
5. The upper aq. Content of solid lipid Nanoparticles dispersion(70-99.9%).
6. Adjustment of drug unharness profile.

INGREDIENTS USED FOR SOLID LIPID NANOPARTICLES

Various ingredients square measure accustomed prepare solid Nanoparticles embody solid lipid surfactant, co-surfactant and water. The choice of ingredients depend upon the aim of formulation like web site of targeting, dominant the drug unharness, particles homegenecity, route of administration, stability on storage. Within the preparation of solid lipid Nanoparticles charge modifier, co surface active agents, cryoprotectent and preservative also are used as a additives.[3,4,5,6]

LIPID MATRIX

Lipid that square measure solid at temperature square measure use to arrange solid lipid Nanoparticles, are perishable, biocompatible with body and have low unhealthful. The crystalline lipid has ordered unit it enhance lipids stability thanks to their polymorphism and crystalline kinetics. The lipid reborn into choice is most significant issue. The lipid embody trimyristae, spinoff of calyx arene, chemical group momostearate, saturated fatty acid, glycer unhealthful metabolites square measure made. Palmitostearate, hydrodenated animal oil, paraffin, caprylicapric acylglycerol, bees wax and coca butter. The miscibility and solubility of drug in lipid matrix, crystallinity and physicochemical property of lipid affects drug defense, the lipid with high crystalline structure show low loading and natural action of drug. They will be used alone or in combination of two or lot of lipid to boost the drug defense, unharness behavior and stability.

SURFACTANT

Surfactant square measure used as stabilizer within the preparation of solid lipid Nanoparticles. Surfactants build the emulsion stable and stay at interface for while that protects formulation. The surfactants ought to be non unhealthful, non infliction, compare with another ingredients. The number of the surface active agents ought to be optimum as a result of low amount result the formation of enormous particles wherever as higher conc. Leads to tiny size to associate degree extent. At higher conc. of surface active agents the viscousness of the system is high that decrease diffusion speed and therefore the particles stay huge. Numerous surface active agent square measure used to prepare solid lipid Nanoparticles which has alkyl group organic compound, bile salt, cetyl pyridinium chloride, cetyl ammonium ion bromide, curdlan, egg lecithin, lauryl amino acid, poloxamer 237, poloxamer 238, poloxamer 239, polyethylene glycol 4500, stearate, sorbitantrioleate, soyabean plant phospholipid, sod. Lauryl sulphate, span60, span20, span80, tween80, tween 60 and tween 20 etc. The surface active agent will have an effect

on unharness of drug, particle size and in- vivo behavior. Several researchers evaluated numerous surfactant for brain targeting the tween 80 has shown the promising effects.

CO-SURFACTANTS

Co-surfactant additionally employed in the preparation of solid lipid Nanoparticles as a result of the surfactant has low quality to hide the surface throughout recrystallization thus the particles is also combination which may cause instability, to beat this drawback the nonionic or ionic surface active agents is also used. They kind micells which may function reservoir. Ordinarily benzalkonium chloride, butanol, glycerol, tristearin square measure used as co-surfactant. The little and stable particles square measure fashioned thanks to formation of microemulsion.[2,3]

METHOD OF PREPARATION OF SOLID LIPID NANOPARTICLES

1.MECHANICAL TECHNOLOGY

Mechanical technology wide accustomed scale back the particle into tiny particles by include mechanical edge, air mass homogenization victimization high energy input which end in high impact thanks to that cause particle size reduction takes place.

A.Air mass homogenization

In this technique a high shear pressure is employed to interrupt down the particles into tiny Nanoparticles, a air mass (100-200 bars) pushed the lipids to with stand a slender gap; this causes turbulence, cavitations and collision of particles to every alternative that results in breakdown of the particles into nano vary. The fluid circulated to a awfully small distance at high viscousness of regarding 1000-2000km/h. in distinction to a different preparation technique, this technique has no scaling drawback. Air mass homogenization is to two type hot- homogenization and cold – homogenization.

a.Hot Homogenization

In the hot homogenization first pre-emulsion of liquid lipid containing drug and aq. part having associate degree wetter is ready by victimization the equal worker with the assistance of a high shear combining instrumentation. For this lipid is molten higher than the temperature of the lipid. A hot o/w emulsion is ready on cooling of the emulsion crystallization of lipid takes place and solid lipid Nanoparticles square measure fashioned, thanks to lowered viscousness of the lipid at higher temperature part terribly tiny particle square measure made. At extreme temperature lipid and drug is also degrade. Thus 3-5 homogenization cycles square measure used. A nanoemulsion is created that on cooling at temperature, the lipid reborn into solid particles.

b.Cold Homogenization

Ultra-sonication / high speed homogenization

Ultrasonication or high speed homogenization techniques square measure accustomed prepare smaller solid lipid Nanoparticles, ultrasonication and high speed homogenization employed in combination. Solid lipid Nanoparticles square measure ready by melting the lipids ana new technique core material, associate degree emulsion is ready by adding binary compound medium into liquid lipid at higher temperature by combining with the assistance of mechanical stirrer, or sonication the method is done at low shear stress however metal contamination and particles growth upon storage, to big particles is determined. To emulsify the lipid in binary compound part, the lipid is dissolved in a water- immiscible organic solvent (eg. Cyclohexane). Then the organic solvent is evaporated from the emulsion by evaporation below reduced pressure (40-60mmHg) in rotary evaporator, precipitation of the lipid

dispersion within the binary compound medium takes place and 25nm mean size Nanoparticles square measure fashioned. If the coarse emulsion immediately homogenized the potency of fine emulsification will be improved.

1.CRITICAL FLUID METHODOLOGY

A new technique for the preparation of solid lipid Nanoparticles within which solvent isn't needed for process, is employed these days. A supercritical fluid particularly green house gas, is used during this technology as a result of greenhouse gas has low toxicity, low price high solubility power, and having a coffee essential 31C. 10C at pressure of seventy three bar. When the pressure and temperature of a fluid redoubled to the essential worth it's celebrated as critical fluid and solubilizing capability of the fluid is will increase. There two methods area unit unremaekably employed; one is speedy enlargement of critical resolution (RESS) and second is speedy enlargement from critical to aq. resolution of emulsions (RESAS).

Rapid Enlargement of Critical Resolution (RESS)

Solid lipid Nanoparticles area unit ready by the speedy enlargement of critical carbon dioxide solutions methodology. During this methodology critical fluid (CO₂) is employed as solvent, containing the substance dissolved in it that is then exhausted through nozzle to the collection chamber at gas pressure, the answer, on enlargement the atomization and evaporation of fluid cause nucleation and precipitation of substance into Nanoparticles. Rapid enlargement from critical to aq. resolution of emulsion (RESAS) particles get aggregating in RESS methodology this methodology is employed to urge tiny particles, in this methodology the surface active agent get diffuse to particles surface and forestall particles from agglomeration, this methodology is employed to make Nanoparticles of water insoluble medicine. In this methodology Nanoparticles area unit ready by part separation technique, the drug is dissolved into water immiscible organic solvent, on addition into the liquid part containing water soluble surface active agents, associate degree liquid dispersion is created, the temperature of the system is elevated which ends up in speedy evaporation of organic solvent and thence phase separation is occurred. The particles formation stated, the stabilizer the formulation. This system is employed to boost the solubility of medicine, because of formation of tiny particles and formation of the constancy crystalline or remittent crystallinity or increased wetting capability of drug.

2.Microemulsion Primary Based Methodology

Solid lipid Nanoparticles ready by microemulsification followed by dilution of microemulsions. The ,icroemulsion is ready by stirring the low melting carboxylic acid in aq. solution of associate degree wetting agents associate degreeed co – surface active agent at 65-70°C to make an clear mixture. The low melting carboxytic acid (stearic acid) is employed associate degreeed an wetting agent (poplyisorbate 20, polysorbate 60, soy phosphatidylcholine, and metallic element teurodeoxycholate), co-emulsifiers (sodium mono octylphosphate). The ready hot microemulsion is then dispersed in cold water (2-4°C) below stirring. The dilution method is critically determined by the composition of microemulsion.

3.Double Emulsion Methodology

To load water soluble medicine in solid lipid Nanoparticles solvent emulsification evaporation methodology is employed, a hot w/o/w double microemulsion is ready in two steps. In the first step the melted lipid supplementary to associate degree solution containing drug surfactant and co-surfactant the process worker is slightly higher than the freezing point of lipid to prepare w/o microemulsion. Within the second step, to arrange transparent w/o/w system the prepared w/o microemulsion is then poured into mix of liquid surfactants and co-surfactant to urge a

double emulsion. Then heat double small emulsion distributed in ice cold water, so laundry is finished by form by extremist filtration system.

4.Precipitation methodology

The precipitation methodology is accustomed prepare solid lipid Nanoparticles the organic medium is needed for this methodology. The lipid is dissolved on associate degree medium (eg. Chloroform) and also the emulsification of organic medium is finished is associate degree lipid medium. The organic medium is then gaseous at the reduced pressure in turning evaporator, the precipitation of lipid is takes place in lipid medium and Nanoparticles area unit developed.

5.Solvent Injection Technique

In the solvent injection technique the answer solid lipid Nanoparticles areaunit ready in water-miscible solvent system (eg. Ethanol, acetone, isopropanol). Or a mixture of two or lot water mixable solvent. To arrange the nanoparticle lipid resolution is then injected victimization associate degree injection needle into liquid part which can or might not containing surface-active-agent with a relentless stirring. A dispersion is then obtained that is then filtered with a paper so as to get rid of any excess lipid. The wetting agent is basically utilized in the liquid droplets at positioning of injection and stabilize solid lipid nanoparticles till solvent diffusion is completed by reducing the surface tension between the water and solvent.

6.Spray Drying Methodology

A cheaper various technique than the freeze drying methodology. The lipid having high melting worth area unit used. The carbohydrate and low lipid content area unit supplementary during this, which help to preserve the dimension of mixture particles in spray drying to stop the melting of the lipid ethanol- water mixture is utilized in place of victimization water as result of on cooling tiny and heterogeneous crystals formation takes place, at low recess temperature. This particles will aggregate as a result of the high worker, high shear force and partial melting of the particles. Secondary production steps.

a.Sterilization

If nanoparticles area unit given by epithelial duct route, the sterilization of formulation is needed for sterilization in the main autoclaving is finished that uses the damp heat. When the sterilization is to be done numerous issue to be contemplate concerning the drug and excipients of the formulation, if drug is temperature sensitive, at higher worker. It is degrade the particles size is modified because of the consequences of sterilization usually the it's found to cause a rise in particles size. The essential parameter embody sterilization temperature and also the composition solid lipid Nanoparticles. Additionally the choice of an honest emulsifier incorporate vital and necessary role on the physical stability of the sample at high temperature. If the worker is redoubled, it will modification the quality and also the water solubility of all emulsifiers to a distinct extent. Stream sterilization cause the formation of associate degree o/w-emulsion because of the melting of the lipid particles. Solid particles area unit fashioned when recrystallization. γ - irradiation can be another methodology to stream sterilization for temperature sensitive samples.

b.Evaporation

To enhance the evaporation or freeze drying is that the most promising means. In case of nanosuspension for orally administered drug which may be precipitated because of transformation product into solid state because of Ostwald ripening of drug, which can results in hydrolytic degradation of drug, to stop such conditions associate degree adequate amount of cryptotectant is supplementary to stop solid lipid Nanoparticles aggregation at the time evaporation method.[3,4,5]

List of Patents on SLN

Title of Patent	Patent No/ Publication No./Application No.
Lipid nanopellets for oral administration	EP0167825 [8]
Medication vehicles made of solid lipid particles	WO9305768 [9]
Method for producing solid lipid microspheres having a narrow size distribution	US5250236 [10]
Pharmaceutical cyclosporine formulation with improved biopharmaceutical properties, improved physical quality and greater stability, and method for producing said formulation	US6551619 [11]
Lipid nanoparticles as vehicles for nucleic acids, process for their preparation and use	WO05120469 [12]
Lipid emulsion and solid lipid nanoparticle as a gene or drug carrier	WO0006120 [13]
Compositions for the targeted release of fragrances and aromas	CA2524589 [14]
Novel method for preparing solid lipid particles using a membrane reactor	WO07000531 [15]
Solid lipid nanoparticle as a gene or drug carrier, formulation and method for preparing the same	KR2033665A [16]
Colloidal solid lipid vehicle for pharmaceutical use	US2006222716 [17]
Use of solid lipid nanoparticles comprising cholesteryl propionate and/ or cholesteryl butyrate	WO06128888 [18]
Method for producing solid-lipid composite drug particles	US2006008531 [19]
New topical formulation which includes active agent as liquid lipid nanoparticles in an oil-in-water emulsion	DE19825856 [20]
Topical preparation containing a suspension of solid lipid particles	US5667800 [21]
Pharmaceutical compositions suitable for the treatment of ophthalmic diseases	US2006024374 [22]
Solid lipid nanoparticles containing a UV filter material are useful in aqueous dispersion form as high filter content sunscreen compositions	DE19952410 [23]
Formulation of UV absorbers by incorporation in solid lipid Nanoparticles	US7147841 [24]

Microemulsions as precursors to solid nanoparticles	US7153525 [25]
Lipid matrix-drug conjugates Particle for controlled release of active ingredient	US6770299 [26]
UV radiation reflecting or absorbing agents, protecting against harmful UV radiation and reinforcing the natural skin barrier	US6814959 [27]
Solid lipid Nanoparticles	430/DEL/2004 [28]
Pharmaceutical compositions suitable for treatment of ophthalmic diseases	975/KOLNP/2005 [29]

APPLICATIONS OF SLNs

SLNs For Parenteral application

Wising et al. (2004) reviewed that parenteral use of SLS is short of safe and well tolerated as they incorporate physiologically well-tolerated ingredients and that they have smart storage capabilities once evaporation or sterilization. Once injected intravenously, SLNs are terribly little to flow into within the microvascular system. Therefore, SLS are prompt for infectious agent and non-viral factor delivery. Cationic SLS have potential advantages in targeted factor medical care in treatment of cancer. Treatment of central system disease like brain tumors, AIDS, medicine and psychiatric disorders is feasible via SLSs as deliquescent coating of collides improve the transport of those through BBB and tissue distribution.

SLNs For Nasal Application

Nasal administration was a promising various route of drug administration because of quick absorption and fast onset of drug action and avoiding degradation of labile medication within the GI. SLNs were projected as various trans mucosal delivery systems by numerous analysis teams. The role of PEG coating of polylactic acid Nanoparticles in rising the trans mucosal transport of the encapsulated bioactive molecule rumored to achieve success by tobio et al, 1998.

SLNs For Metastasis Application

The lungs supply a high area for drug absorption by avoiding first-pass effect. SLSs will be projected as carrier of anti-cancer medication in carcinoma treatment or amide medication to enhance their bioavailability. In a very recent study, anti tubular medication (rifampicin, antibacterial and pyrazinamide) were incorporated into numerous formulations of solid lipid particles ranged from one. 1-2.1 μ m and formulations were nebulized to guinea pigs orally for direct pulmonary delivery. Nebulization of solid lipid particles carrying antitubular medication was determined to achieve success in rising drug bioavailability and reducing the dosing frequency for higher management of tuberculosis.

SLNs For Ocular Application

Biocompatibility and mucoadhesive properties of SLNs improve their interaction with ocular mucous membrane and prolong tissue layer duration of the drug, with the aim of ocular drug targeting. Cavalla et al., (2002) evaluated SLNs as carrier for ocular delivery of antibiotic in rabbit eyes. As a results SLNs considerably increased the drug bioavailability within the body fluid . cavilli et al., (1995) conjointly studies alkaloid delivery via SLNs, that is usually utilized in eye disease treatment, earlier. They rumored terribly similar leads to order to reinforce the ocular bioavailability of drug.

SLNs For Topical Application

SLNs and NLC are terribly enticing mixture carrier system for skin application because of their numerous fascinating effects on skin besides the characteristics of a mixture carrier system. They are like minded to be used on broken or inflamed skin as a result of they are supported non- irritant and non-toxic lipids. Researchers have rumored intensively on the topical application of SLS. Throughout the previous few years, SLN and NLC are studied with active compounds like vitamin E ,vitamin E acetate, retinol, ascorbyl palmitate, clotrimazole, triptolide and phodphyllotoxin for topical application. A very new, recently discovered space of application is that the use of SLN in sun-protective creams.

SLN in Cancer Chemotherapy

From the last twenty years many chemotherapeutical agents are encapsulated in SLN and their in-vitro and in-vivo effectivity are evaluated. Tamoxifen, associate in nursing antitumor drug has been i.v. administration in carcinoma (Kambale and Jagdale 2010).

Market Potential

The projected marketing research on skin care merchandise signals study growth perspective in future. The market growth { is expected|is predicted| is associate in nursing ticipated} to rise apace at an annual rate of growth of 7%.the projected market potential is anticipated to succeed in \$31.84billion by 2016. The Asian countries like Japan, China and India expected to supply a large impact on this world skin care market (Venkateshswarlu and Manjunath, 2004).

CONCLUSION

In the period of the 20th century, bacteriologist unreal his curative concept; the though that medication reaches the proper website within the body, at the proper time, at right concentration. It should not exert facet effects, neither on its thanks to the therapeutic target, nor at the target website, nor throughout the clearance method. The SLNs have the potential to realize, a minimum of partially, these broad objectives. A side from these, the regular objective of controlled drug delivery is with competence achieved with SLNs. They are comparatively young drug delivery systems, having received primary attention from 1990s and have hold nice promise for its systematic investigation and exploitation. We will expect several proprietary does form within the style of SLNs within the feature.

REFERENCES

1. Agnieszka Z, Wilczewska, Katarzyna N, Karolina H. Markiewicz, Halina C, Nanoparticles as drug delivery systems, *Pharmacological Reports*, 2012, 64, 1020.1037.
2. Vyas S P, Khar R K, Targeted And Controlled Drug Delivery: Novel Carrier System, Vallabh Prakashan, Delhi, 2010 reprint
3. Ravi MNV, Handbook of particulate drug delivery, American Sci publishers, California, vol 1, 96-161.
4. Kaur I P, Bhandari R, Bhandari S, Kakkar V. Potential of solid lipid nanoparticles in brain targeting *Journal of Controlled Release* 127 (2008) 97 –109.
5. Lakshmi DP, Nair R, Chakrapani M, Venkatkrishnakiran P, Solid lipid Nanoparticle systems for delivery of drugs to the brain, *International Journal of Biopharmaceutics* . 2012, 3(2), 70-77.
6. Müller, R.H., K. Mäder, and S. Gohla, A review on SLNs for controlled drug delivery- *European journal of pharmaceutics and biopharmaceutics*, 2000, 50(1): p. 161-177.
7. Wong H, Rauth A M, Bendayan R, Wu X Y, In vivo evaluation of a new polymer-lipid hybrid nanoparticles (PLN) formulation of doxorubicin in a murine solid tumor model. *European Journal of Pharmaceutics and Biopharmaceutics*, 2007, 65, 300–308.
8. Speiser, P.: EP0167825 (1990).
9. Muller, R.H., Lucks, J.S.: WO93/05768 (1996).
10. Gasco, M. R.: US5250236 (1993).
11. Penkler, L. J., Muller, R.H., Runge, S.A., Ravelli, V.: US20036551619 (2003).
12. Gasco. M. R.: WO05120469 (2005).
13. Jeong, S.Y., Kwon, I. C., Chung, H.: WO00006120 (2000).
14. Dahms. G., Seidel, H., Jung, A.: CA2524589 (2004).
15. Catherine, C., Fessi, H.: WO07000531 (2007).
16. Jung, H.S., Jung S. Y., Kwon I.C.: KR2033665A (2002).
17. Schwarz, J., Weissapir. M.: US2006222716 (2006).
18. Gasco, M.R.: WO06128888 (2006).
19. Shekunov, B.Y., Chatopadhav, P., Huff, R.W.: US200 (2006).
20. Labtec, G.F.T.: DE825856 (1999).
21. Vringer, T.D.: US5667800 (1997).
22. Gasco, M.R., Saettone, M.F., Zara, G.: US20060243 (2006).
23. Heppner, A., Hansen, P., Schumann, C.: DE19952410 (2001).
24. Herzog, B.: US20067147841 (2006).
25. Mumper, R.J., Jay, M.: us20067153525 (2006).
26. Muller, R.H., Carsten, O.: US20046770299 (2004).
27. Muller, R. H., Wissing, S., Mader, K.: US20046814959 (2004).
28. Khuller, G.K., Pandey, R.: 430/DEL/2004 (2004).
29. Gasco, M.R., Zara, G. P., Saettone, M.F.: 975/KOLNP/2005 (2005).