

Development and Characterization of Lipid Based Nano Drug Delivery Systems for Some Poorly Bioavailable Drugs

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Abstract— Considering lipid-based nanocarriers have several advantages, such as biocompatibility, biodegradability, low toxicity, and non-immunogenicity, nanomaterials have undergone extensive research for the delivery of drugs. However, the efficacy of therapy or the effectiveness of administering the medication are limited by the disadvantages that accompany traditional lipid-based nanocarriers as particularly inappropriate aiming for, acquisition by the system of reticuloendothelial cells, and rapid clearance. To contribute to facilitate identify and address the causes of a variety of diseases, a number of multifunctional nanocarriers constructed from lipids have been created constructed that improve the development of pharmaceuticals in the area of the lesions. Everyone described the latest advances along with applications of nanocarriers made with lipids in this review, wrapping both traditional and intriguing operational a lipid formulation.

Index Terms- Bioavailable drugs, Self-micro emulsifying drug delivery system.

I. INTRODUCTION

Conventional "free drugs" in medicinal products frequently exhibit an assortment of common worries, including limited solubility, detrimental pharmaceutical kinetics, inferior biological distribution, and inadequate tissue sensitivity. Drug delivery systems (DDS) have enhanced pharmaceutical kinetics, efficacy in therapy, and solubility over drug molecules, which are besides a number of other advantages. Substantially nanodrug delivery system might hypothetically supply increased drug distribution efficiency by particular tissue targeting, which is improved cell internalizing it, size delivery to specific mitochondria because of its small size and structure. But the toxicity of nanocarriers

always brings up an important consideration, in particular with reference to medical uses. a variety of reports, ferrous and polymers the nanocarriers may be deleterious to the renal nervous system, brain, or pulmonary.

II. LITERATURE REVIEW

1. H Shrestha et al. (2014): - Improving the bioavailability of lipid-based medications is the main goal of their formulation. Although it is no longer a novel idea, the use of lipids in medication administration is nevertheless a promising one. One of the newest methods to solve issues such bioavailability and solubility of poorly water-soluble medicines is lipid-based drug delivery systems (LBDDS).
2. R Kumar et al. (2019): - The creation of workable new pharmaceuticals for the management of sickness minus any side effects is limited by the multiple phases involved in the discovery of novel therapeutic molecules. Nevertheless, despite demonstrating substantial biological activity in in vitro and in animal investigations, the majority of medications fail in the clinical stage. This could be because both in vitro and in vivo research were conducted without a physiological context.

III. MATERIALS AND METHODS

Inorganic Reactions

Alembics Pharmaceutical Ltd. (Vadodara, India) generously donated iloperidone (ILO) and vardenafil HCL Trihydrate (VDN). The additives that were utilized to develop the formulation are listed in Table 1 and were utilized exactly as supplied. Aqueous

buffers, analytical compounds, and purified HPLC grade water were created using reagents and chemicals that were acquired by passing water that has been double-distilled over nylon paper filters with a 0.45 μm pores (Pall Life sciences, Mumbai, India).

Excipients	Manufacturer/Supplier
Acconon C-80	Abitec Corporation, USA
Acconon CC-6	Abitec Corporation, USA
Brij 35	S.D. Fine Chemicals, Mumbai, India
Capmul MCM	Abitec Corporation, USA
Capmul MCM C8	Abitec Corporation, USA
Capmul MCM L8	Abitec Corporation, USA
Capmul PG 8 (Capryol 90)	Gattefosse, France
Capryol PGE 860	Abitec Corporation, USA
Capryol PGMC	Gattefosse, France
Captex 200 (Labrafac PG)	Abitec Corporation, USA
Captex 300 (Labrafac Lipophile WL 1349)	Abitec Corporation, USA
Captex 500	Abitec Corporation, USA
Castor Oil	S.D. Fine Chemicals, Mumbai, India
Cholesterol	Sigma-Aldrich, Mumbai, India
Coconut Oil	S.D. Fine Chemicals, Mumbai, India
Cremonophor EL	Sigma-Aldrich, Mumbai, India
Ethyl Oleate	S.D. Fine Chemicals, Mumbai, India
Glycerol	S.D. Fine Chemicals, Mumbai, India

(Table 1) Collection of excipients that for formulating

The overarching principles regarding cultivating cells The National Center of Cell Sciences in Pune, India, has a tissue archive from which the Caco2 (Homo sapien colonic carcinoma cells) line of cells was collected. The cell line (Jouan the IGO 150 an incubator, Thermo Fisher Scientific, Mumbai, India) was kept at 37°C in a moistened 5% carbon dioxide surroundings. The starter cultures were kept alive in MEM containing one percent antimicrobial solutions and ten percent heat-inactivated fetal bovine serum (FBS) [1].

Acquiring Information

To make the entire media outlets, MEM was mixed with 1% v/v antimicrobial solution and ten percent v/v heat-inactivated fetal bovine serum (FBS). This complete media set was kept cold (NMT 25° C) in an empty screw-capped container. After that, the jar was encased in aluminum foil and parafilm was used to seal it. The experiment was conducted beneath an inclined chamber with turbulent air flow.

Cell line subculturing

In T-25 culture cell containers, cells were kept as multilayer colonies and were subculture thrice per

week. Cultured lines of cells were thereafter maintained in full medium at 37°C in a humidified atmosphere with ninety-five percent air and five percent carbon dioxide (Jouan IGO150 carbon dioxide an incubator, Thermo Fisher Scientific, India). Every day for three days, an alternate full media was replaced [1,2].

Comprehensive Technique:

- The entire medium was a 37°C water bath that had been preheated.
- After removing the cells from the incubator, they were examined under a microscope to ensure that they had grown to about 80% confluence.
- Aspiration was used to extract the medium from the flask. The flask-attached cells underwent two rounds of washing in culture media devoid of serum.
- After introducing one milliliter of trypsin solution containing EDTA, the resulting mixture was let sit for five minutes at a time without periodic triggering in order to release cells.
- To halt trypsin action, 5 mL of the full medium was subsequently added.
- Following a gentle out-and-down pouring motion to separate up all cell clusters, the cells were collected after which they were reseeded at an appropriate seeding density into an empty flask.
- Following this, the test tubes were successfully nourished.

Standard operating protocol for characterisation methods

Dimensions

Arguably those most important components of a nano-sized composition is the dispersion and size of the fragments, since that influences its absorption and penetration into tissue. The application of dispersion techniques is more beneficial for size measurements, which happens in the nano range. Either the shape and dimension of nanoparticles can be identified via scattering methods. DLS, SANS, small-angle scattering of X-rays (SAXS), or static scattering of light (SLS) are just a few of the commonly used scattered methods. To determine the size, structure, and connections amongst the particles, one must

employ SLS, SAXS, and SANS. DLS hinges around the particle's physicochemical size-related propagation in liquids [3,4].

The kind of radiation used determines the kinds of specimens that can be researched using dispersion approaches, the sample habitat that can be used, the range of lengths that can be probed, and the details that can be obtained. For instance, SANS may be applied with ease to opaque samples that DLS is unable to study, thanks to its high penetration depth of monochromatic neutrons beam. While DLS analyses the particle's hemodynamic size, SANS measures a particle's actual size. The hemodynamic size is always bigger than the actual size since the LASER scatters through the hydration barrier around the particles themselves [5].

CONCLUSION

We talked about the method of synthesis and characterization of lipid-based drug delivery systems for vardenafil and iloperidone, two insoluble medications, in the current study. Here, we have created SMEDDS and Niosome using a methodical, QbD-enabled technique. It was the initial approach of using ANNs for optimizing the formulation of SMEDDS. When compared to drug suspension, the formulations demonstrated improved in vitro, ex vivo, and in vivo performance. By resolving PGp efflux, solubility problems, and the first pass metabolism effect, the lymphatic focused treatment strategy will lead to better bioavailability. Overall in vivo pharmacokinetics, as well as pharmacodynamics, investigation provided additional evidence of the generated methods' effectiveness in bolstering the results of ex vivo alongside in vitro line of cells experiments. The aim of this research was to generate LBDDS for the targeted medications in order to increase solubility and avoid the first-pass metabolism via capillary targeting, both of which will aid in enhancing the nano formulations' absorption. Formulated innovation provided by Quality by Design (QbD) creates a design space. Therefore, in order to strengthen the design space, we concentrated on developing the administration area. An additional improvement strategy which was taken into consideration was the Artificial Neural Network (ANN). The comparison was conducted comparing the

results of Design of Experiment (DoE) and ANN. To investigate the toxicity profile and formulation uptake, in vitro cell line experiments were conducted. After the formulation was evaluated in vivo, the absorption mechanism of LBDDS was identified. Among the various ways to boost oral their bioavailability better the dissolution in the GIT is a prominent way to improve absorption. LBDDS is one technique used in nanomedicine for the same purpose. The main component of LBDDS is the lipids portion, which may consist of a combination of various kinds of lipids or a single substance. liquid, solid, Vesicular in or particles delivery systems were a few instances of LBDDS. In this paper, medication manufacturing techniques for iloperidone (ILO) and vardenafil HCl trihydrate (VDN) are analysed using SMEDDS and Niosome.

REFERENCES

- [1] Torchilin, V. P. (2006). Introduction. Nanocarriers for Drug Delivery: Needs and Requirements. In V. P. Torchilin (Ed.), Nanoparticulates as drug carriers (pp. 1-8). London: Imperial college press.
- [2] Gao, P., Witt, M. J., Haskell, R. J., Zamora, K. M., & Shifflett, J. R. (2005). Application of a Mixture Experimental Design in the Optimization of a Self-Emulsifying Formulation with a High Drug Load. *Pharmaceutical Development and Technology*, 9(3), 301-309, doi:10.1081/PDT-200031441.
- [3] Date, A. A., & Nagarsenker, M. S. (2007). Design and evaluation of self-nanoemulsifying drug delivery systems (SNEDDS) for cefpodoxime proxetil. *International Journal of Pharmaceutics*, 329(1), 166-172, doi:https://doi.org/10.1016/j.ijpharm.2006.08.038.
- [4] Javed, A., Saima, A., Kanchan, K., & Showkat, M. (2013). Construction of Pseudoternary Phase Diagram and its Evaluation: Development of Self-dispersible Oral Formulation. *International Journal of Drug Development and Research*, 5(2), 84-90.
- [5] Patel, A. R., & Vavia, P. R. (2007). Preparation and in vivo evaluation of SMEDDS (selfmicro emulsifying drug delivery system) containing fenofibrate. *The AAPS Journal*, 9(3), E344-E352, doi:10.1208/aapsj0903041.