

A Critical Review on Applications of New Drug Delivery System

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ABSTRACT: *Great strides have been made in the development of novel drug delivery systems (NDDS) for plant actives and extracts over the past few years. Bioactive and plant extracts have been used to record the range of novel herbal formulations such as polymeric nanoparticles, nano capsules, liposomes, phytosomes, nano emulsions, microsphere, transferosomes, and ethosomes. The novel formulations are reported to have remarkable advantages over traditional formulations of plant actives and extracts, including solubility enhancement, bioavailability, toxicity safety, pharmacological activity enhancement, and stability enhancement, improved distribution of tissue macrophages, sustained delivery, and physical and chemical degradation protection. The present review highlights the current status of the development of novel herbal formulations and summarizes their method of preparation, type of active ingredients, size, and entrapment efficiency, route of administration, biological activity and applications of novel formulations.*

KEYWORDS: *Drug, Liposomes, Plant, Particles, Systems.*

INTRODUCTION

The liposomes are spherical particles in which a portion of the solvent is encapsulated and freely diffused (floated) into their interior. They can have one, two or multiple concentric membranes. Liposomes are composed of polar lipids characterised by the same molecules having a lipophilic and hydrophilic group. Polar lipids self-assemble and form self-organized colloidal particles upon contact with water[1]. Detergents, components form micelles, are simple examples, whereas polar lipids with bulkier hydrophobic components cannot interact with high curvature radii micelles but form bilayers that can self-close to liposomes or lipid vesicles. A liposome cross-section (Fig. 1) shows the amphiphile hydrophilic heads oriented towards the water compartment, while the lipophilic tails are oriented towards the middle of the vesicle away from the water, thereby forming a bilayer[2].

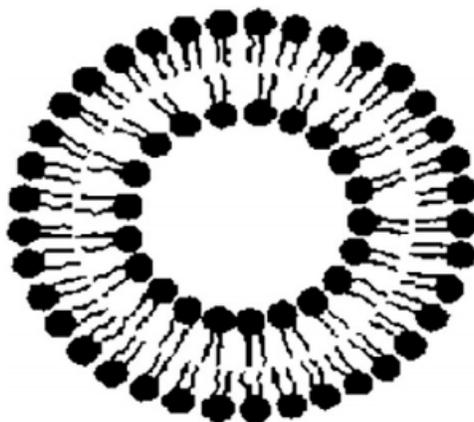


Figure 1: Illustrates the Cross-section of a liposome[3].

Table 1: Illustrates the Liposomal herbal formulation[4].

| Formulations | Active ingredients | Applications of liposome formulations | Biological activity | Method of preparation | % Entrapment efficiency | Route of administration |
|----------------------------------|-------------------------------------|---|---------------------------------|---|-------------------------|-------------------------|
| Quercetin liposomes | Quercetin | Reduced dose, enhance penetration in blood brain barrier | Antioxidant Anticancer | Reverse evaporation technique | 60% | Intranasal |
| Liposomes encapsulated silymarin | Silymarin | Improve bioavailability | Hepatoprotective | Reverse evaporation technique | 69.22 ± 0.6% | Buccal |
| Liposoma artemisia arborescens | Artemisia arborescens essential oil | Targeting of essential oils to cells, enhance penetration into, cytoplasmatic barrier | Antiviral | Film method and sonication | 60–74% | In vitro |
| Ampelopsin liposome | Ampelopsin | Increase efficiency | Anticancer | Film-ultrasound method | 62.30% | In vitro |
| Paclitaxel liposome | Paclitaxel | High entrapment efficiency and PH sensitive | Anticancer | Thin film hydration method | 94% | In vitro |
| Curcumin liposome | Curcumin | Long-circulating with high entrapment efficiency | Anticancer | Ethanol injection method | 88.27 ± 2.16% | In vitro |
| Garlicin liposome | Garlicin | Increase efficiency | Lungs | Reverse-phase evaporation method | 90.77 % | – |
| Flavonoids liposomes | Quercetin and rutin | Binding of flavonoids with Hb is enhanced | Hemoglobin | Solvent evaporation | – | In vitro |
| Usnea acid liposome with β-CD | Usnea acid | Increase solubility and localization with prolonged-release profile | Antimycobacterial | Hydration of a thin lipid film method with sonication | 99.5% | In vitro |
| Wogonin liposome | Wogonin | Sustained release effect | Anticancer | Film dispersion method | 81.20 ± 4.20% | In vivo |
| Colchicine Liposome | Colchicine | Enhance skin accumulation, prolong drug release and improve site specificity | Antigout | Rotary evaporation sonication method | 66.3 ± 2.2% | Topical |
| Catechins liposomes | Catechins | Increased permeation through skin | Antioxidant and chemopreventive | Rotary evaporation | 93.0 ± 0.1 | Transdermal |



Figure 2: Illustrates the (a) nano emulsion and (b) biopolymeric nanoparticle[5].

DISCUSSION

Table 2: Illustrates the Phytosomal herbal formulations.

| Formulations | Active ingredients | Applications of phytosomal formulations | Biological activity | Method of preparation | Dose | Route of administration |
|---------------------------------|--------------------|--|--|-------------------------------------|----------------------|-------------------------|
| <i>Ginkgo biloba</i> phytosomes | Flavonoids | Flavonoids of GBP stabilize the ROS | Cardio-protective, antioxidant activity | Phospholipids complexation | 100 mg and 200 mg/kg | Subcutaneous |
| Ginkgoselect phytosome | Flavonoids | Inhibits lipid peroxidation (LPO), stabilize the ROS | Hepatoprotective, antioxidant | Phospholipids complexation | 25 and 50 mg/kg | Oral |
| Silybin phytosome | Flavonoids | Absorption of silybin phytosome from silybin is approximately seven times greater | Hepatoprotective, antioxidant for liver and skin | Silybin-phospholipid complexation | 120 mg | Oral |
| Ginseng phytosome | Ginsenosides | Increase absorption | Nutraceutical, immunomodulator | Phospholipids complexation | 150 mg | Oral |
| Green tea phytosome | Epigallocatechin | Increase absorption | Nutraceutical, systemic antioxidant, anti-cancer | Phospholipids complexation | 50–100 mg | Oral |
| Grape seed phytosome | Procyanidins | The blood TRAP nTotal Radical-trapping Antioxidant Parameter) were significantly elevated over the control | Systemic antioxidant, cardio-protective | Phospholipids complexation | 50–100 mg | Oral |
| Hawthorn Phytosome | Flavonoids | Increase therapeutic efficacy and absorption | Cardio-protective and antihypertensive | Phospholipids Complexation | 100 mg | Oral |
| Quercetin phytosome | Quercetin | Exerted better therapeutic efficacy | Antioxidant, anticancer | Quercetin-phospholipid complexation | – | Oral |
| Curcumin phytosomes | Curcumin | Increase antioxidant activity and Increase bioavailability | Antioxidant, anticancer | Curcumin-phospholipid complexation | 360 mg/kg | Oral |
| Naringenin phytosomes | Naringenin | Prolonged duration of action | Antioxidant activity | Naringenin-phospholipid complex | 100 mg/kg | Oral |

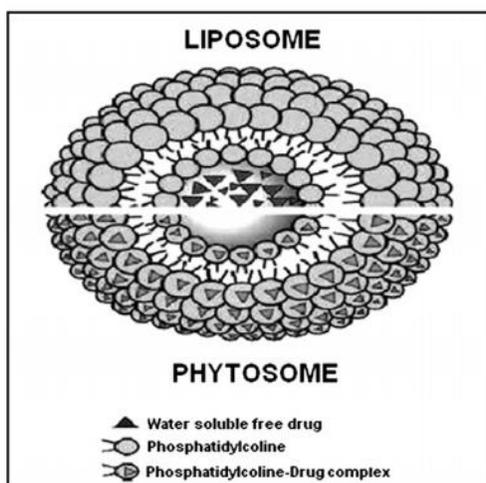


Figure 3: Depicts the difference between liposome and phytosome[6].

ROS has a role in the initiation, growth and maintenance of cancer cells' phenotypical characteristics in cancer. Compared to their normal counterparts, increased ROS generation in tumour cells is due to the involvement of oncogenes that are also involved in malignant transformation[7]. The relationship between the increase in levels of ROS and oncogene activation is still unclear. Inflammation plays an etiological role in cases of some malignancies, such as lung and oral carcinomas. However, the stress induces ROS generation that cause chronic inflammation. Another most commonly implicated mechanism is oxidative DNA damage. ROS can activate various signaling pathways related to persistent tumour survival, metastasis, vascularization and proliferation in cancer cells, in addition to oxidative DNA damage, which can promote cancer growth[8]. Table 1 illustrates the Liposomal herbal formulation. Figure 2 illustrates the (a) nano emulsion and (b) biopolymeric nanoparticle. Table 2 illustrates the Phytosomal herbal formulations. Figure 3 depicts the difference between liposome and phytosome.

CONCLUSION

In the field of novel drug delivery and targeting plant actives and extracts, comprehensive research is underway. Research is, however, still at the exploratory stage in this region. Many research, development and application problems need to be solved. Moreover, in order to create more suitable carriers that can reduce the toxicity of drugs, increase their activity and boost the overall efficiency of agents, more attention should be paid to research on carrier materials. Herbal products have tremendous therapeutic potential that should be explored by certain systems of delivery of drugs with added value. The significant limiting factors for drug molecules to pass through the biological membrane to be systematically absorbed after oral or topical administration are lipid solubility and molecular size. Despite having excellent in vitro bio-activity, many plant extracts and phytomolecules show less or no in vivo activities due to their low lipid solubility or improper molecular size or both, resulting in poor absorption and poor bioavailability.

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