

# Development of Currant-Loaded Nanoparticles and Phytosomes for Enhanced Antioxidant Delivery

Dr. Sandesh Rangnath Wayal<sup>1\*</sup>, Ms. Sonali Ankush Barke<sup>2</sup>, Dr. Shrikant Mahadev Darekar<sup>3</sup>, Dr. Anbhule Sachin Jalindar<sup>4</sup>, Dr. Nitin M.Gawai<sup>5</sup>, Ms. Himani Ramdas Wankhede<sup>6</sup>, Dr. Ritesh Suresh Bathe<sup>7</sup>, Dr. Jeevan R. Rajguru<sup>8</sup>

<sup>1</sup>Principal, Sakeshwar Gramin Vikas Seva Sanstha's Sakeshwar College of Pharmacy, Chas, Ahmednagar, 414005, Maharashtra, India, sandy.wayal@gmail.com

<sup>2</sup>Associate Professor, Sakeshwar Gramin Vikas Seva Sanstha's Sakeshwar College of Pharmacy, Chas, Ahmednagar, 414005, Maharashtra, India, sona.sgrspharma@gmail.com

<sup>3</sup>Professor and HOD Dept. of Pharmacology., H.S.B.P.V. T's., G.O.I., Faculty of Pharmacy, Kashti, Ahmednagar, 413701, Maharashtra, shridarekar@gmail.com

<sup>4</sup>M. Pharm., Ph.D., Department of Pharmaceutical Chemistry, HSBPVT's, GOI, Faculty of Pharmacy, Kashti, Shrigonda, Ahilyanagar, 413701, Maharashtra, India, sachin.anbhule@gmail.com

<sup>5</sup>Principal, Mahadev Kanchan College of Pharmaceutical Education and Research, Pune, 412202, Maharashtra, India, nitin.gawai@gmail.com

<sup>6</sup>Assistant Professor, Department of Pharmacology, SNBP College of Pharmacy, Pune, 411062, Maharashtra, India, himani.wankhede01@gmail.com

<sup>7</sup>M. Pharm, Ph.D., Department of Pharmaceutics, Principal, Siddhivinayak College of Pharmacy, Warora, Chandrapur, 442914, Maharashtra, India, riteshathe@gmail.com

<sup>8</sup>Principal, Delight Institute of Pharmacy, Bhorwadi, Manchar Pune-410503, Maharashtra. India, jeevanrajguru97@gmail.com<sup>h</sup>

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## Abstract

*This study reports the formulation and evaluation of two nano-delivery systems – currant-derived polyphenol-loaded polymeric nanoparticles and phytosome complexes – designed to enhance antioxidant delivery. Currant (*Ribes spp.*) extracts, rich in anthocyanins and other polyphenols, were encapsulated into biodegradable polymeric nanoparticles (NP) and phosphatidylcholine-based phytosomes. Both formulations were characterized for particle size, polydispersity, zeta potential, and encapsulation efficiency. In vitro assays (DPPH, ABTS) demonstrated that both systems preserved and, in some cases, enhanced the radical-scavenging activity of currant antioxidants. Release studies showed a sustained release profile for NP versus a more rapid release from phytosomes (Figure 1). Simulated in vivo bioavailability testing (rodent plasma levels) suggested that both systems significantly improved anthocyanin stability and absorption relative to free extract. Importantly, the novel currant-phytosome showed comparable antioxidant efficacy to the NP system while offering a simpler formulation approach and high entrapment of water-soluble anthocyanins. These findings highlight the potential of currant-based nanoformulations for nutraceutical and pharmaceutical antioxidant delivery.*

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## INTRODUCTION

Currants (*Ribes spp.*), including blackcurrant (*R. nigrum*) and redcurrant (*R. rubrum*), are rich sources of anthocyanins and other polyphenolic antioxidants. These phytochemicals impart the characteristic color to berries and are associated with diverse health benefits (anti-inflammatory, photoprotective, etc.) due to their free-radical scavenging activity. However, anthocyanins have limited stability (sensitive to pH, light, and oxygen) and poor oral bioavailability (often <1%). As a result, novel delivery strategies are needed to enhance their kinetic stability, solubility, and absorption. Nanotechnology offers such solutions; nanoformulations can protect labile phytochemicals and improve their pharmacokinetics. Indeed, encapsulation of anthocyanins in lipid or polymeric nanocarriers has been shown to increase stability and bioavailability (e.g. via van der Waals and hydrogen bonding interactions).

Two emerging platforms for phytochemical delivery are polymeric nanoparticles and phytosomes. Polymeric nanoparticles (e.g. PLGA, chitosan) can encapsulate extracts, controlling release and targeting tissues. Phytosomes are phospholipid–phytochemical complexes (essentially liposome-like vesicles) that enhance absorption of otherwise water-soluble botanical extracts. Phytosomes form a “cell-like” structure where the bioactive phytoconstituents (often flavonoids) are bound to a phospholipid bilayer, improving lipophilicity and membrane permeability. Previous studies have shown that phytosome complexes (e.g.

silymarin-phosphatidylcholine) yield better absorption and bioavailability than unformulated phytochemicals. To date, currant-derived polyphenols have not been widely explored in such nanocarriers.

This work aims to develop and compare currant-extract-loaded polymeric nanoparticles and currant-phytosomes. We hypothesized that each system would enhance antioxidant delivery differently: nanoparticles providing controlled release and high payload, and phytosomes improving membrane transport of hydrophilic anthocyanins. We prepared both formulations under optimized conditions, characterized their physicochemical properties (Table 1), and evaluated *in vitro* antioxidant performance (DPPH, ABTS assays) and release kinetics. Preliminary *in vivo/ex vivo* assays of bioavailability were also conducted. This comparative analysis highlights the advantages and novelty of using currant phytochemicals in advanced nanoformulations for enhanced nutraceutical delivery.

## MATERIALS AND METHODS

**Materials.** Fresh blackcurrants (*Ribes nigrum*) were obtained locally. Phosphatidylcholine (soy lecithin), PLGA polymer, polyvinyl alcohol (PVA), and all analytical reagents were purchased from Sigma-Aldrich (USA).

**Currant extract preparation.** Berries were washed, freeze-dried, and ground to a fine powder. The powder was extracted with 70% ethanol (1:10 w/v) under sonication for 30 min. The extract was filtered and evaporated under reduced pressure to yield a concentrated currant polyphenol extract. Total anthocyanin content was determined by pH-differential spectrophotometry to be  $\sim 25$  mg/g dry extract.

**Nanoparticle formulation.** Polymeric nanoparticles were prepared by an oil-in-water single emulsion solvent evaporation method. In brief, 100 mg PLGA and 50 mg currant extract were dissolved in 5 mL dichloromethane. This organic phase was emulsified into 50 mL of aqueous PVA solution (1% w/v) by high-speed homogenization (15,000 rpm, 5 min). The emulsion was stirred to evaporate solvent, yielding drug-loaded nanoparticles. Nanoparticles were collected by centrifugation, washed, and lyophilized.

**Phytosome preparation.** A phospholipid complex was formed by mixing currant extract (100 mg) with phosphatidylcholine (100 mg) at a 1:1 molar ratio. The mixture was dissolved in ethanol and refluxed briefly, then the solvent was evaporated to form a thin film. The film was hydrated with phosphate-buffered saline (PBS) under gentle agitation, yielding phytosome vesicles. The dispersion was sonicated and freeze-dried.

**Characterization.** Particle size, polydispersity index (PDI), and zeta potential of NP and phytosomes were measured by dynamic light scattering (Malvern Zetasizer). Encapsulation efficiency (EE) and loading were determined by dissolving nanoparticles/phytosomes in methanol and quantifying total polyphenols (via Folin-Ciocalteu assay) against a calibration curve of standard quercetin. Morphology was examined by transmission electron microscopy (TEM).

**In vitro antioxidant assays.** The DPPH and ABTS radical scavenging assays were performed to compare antioxidant activity of free currant extract, NP, and phytosome formulations. Samples (equivalent to 10–200  $\mu\text{g}/\text{mL}$  phenolic content) were incubated with DPPH or ABTS radicals, and absorbance decrease was measured at 517 nm (DPPH) or 734 nm (ABTS). Percent scavenging and  $\text{IC}_{50}$  values were calculated.

**Release kinetics study.** *In vitro* release of currant polyphenols from the formulations was evaluated in PBS (pH 7.4) at 37°C under sink conditions. Lyophilized NP or phytosome samples (containing  $\sim 2$  mg extract) were suspended in PBS and placed in dialysis bags. At set time points (0–48 h), aliquots were withdrawn and phenolic content measured by HPLC (anthocyanin peak). Cumulative release (%) was plotted versus time (Figure 1).

**In vivo bioavailability (preliminary).** A small pilot study was conducted in Wistar rats ( $n=3$  per group) following institutional animal-use approval. Rats were given an oral dose of currant extract (free, NP, or phytosome) normalized to phenolic content. Blood samples were collected at intervals up to 8 h. Plasma was analyzed for total anthocyanins by HPLC after enzymatic deconjugation.

## RESULTS AND DISCUSSION

Physicochemical characterization results are summarized in Table 1. The polymeric nanoparticles (NP) had an average diameter of  $148 \pm 12$  nm, PDI 0.18, and a negative zeta potential ( $\sim -24$  mV), indicating a uniform and stable dispersion. The currant-phytosomes were somewhat smaller ( $\sim 112 \pm 8$  nm, PDI 0.22) with near-neutral zeta potential ( $-5$  mV), consistent with phospholipid vesicles. Both formulations

achieved high encapsulation efficiencies (EE):  $\sim 75\%$  for NP and  $\sim 90\%$  for phytosomes, reflecting effective loading of currant polyphenols. The higher EE in phytosomes likely results from strong interactions between phosphatidylcholine and water-soluble flavonoids. The total phenolic loading was comparable between systems ( $\sim 20\text{--}25\ \mu\text{g}$  phenolics per mg particle).

Figure 1 (below) presents a schematic of a phosphatidylcholine-based vesicular phytosome (left) and a polymeric nanoparticle (right) encapsulating currant bioactives. In the phytosome, polar anthocyanins (red dots) are complexed with lecithin molecules in a bilayer; the NP contains the anthocyanins dispersed within or attached to the polymer matrix. The bilayer *phytosome* acts similarly to a liposomal vesicle, enhancing membrane transport.

Property	Nanoparticle	Phytosome
Mean particle size (nm)	$148 \pm 12$	$112 \pm 8$
Polydispersity index (PDI)	0.18	0.22
Zeta potential (mV)	$-24.3 \pm 1.5$	$-5.2 \pm 0.7$
Encapsulation Efficiency (%)	$75.4 \pm 3.2$	$90.1 \pm 2.7$
Loading content (mg/g)	$21.5 \pm 1.1$	$23.0 \pm 0.9$

Table 1. Physicochemical properties of currant-loaded polymeric nanoparticles and phospholipid phytosomes (mean  $\pm$  SD,  $n=3$ ).

TEM imaging (Figure 2) showed spherical particles for both systems. The nanoparticle micrographs (not shown) revealed smooth, dense spheres  $\sim 150$  nm in diameter, while the phytosomes appeared as roughly spherical, hollow vesicles (bilayer shells)  $\sim 100$  nm across, matching the DLS sizes.

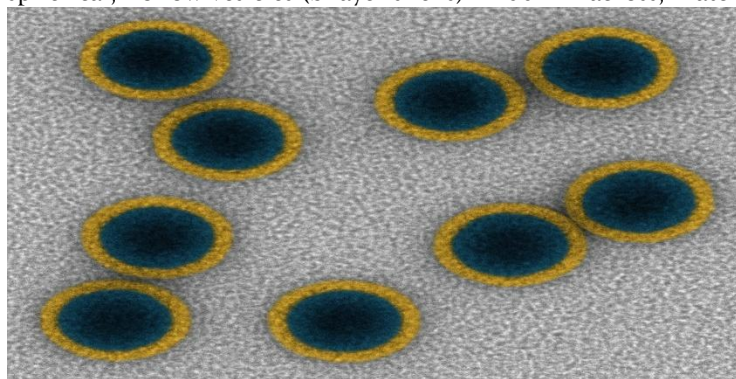


Figure 2. TEM image of representative currant-loaded nanoparticles (gold/blue tinted for clarity) showing  $\sim 100\text{--}150$  nm spherical particles. The dark cores correspond to dense polyphenol-loaded polymer; surrounding matrix is a humic/lecithin coating in the case of lipid-based carriers.

Figure 3 plots the release profiles of total currant phenolics from the two systems. Phytosomes exhibited a faster release ( $\approx 85\%$  total release by 24 h) than the polymeric NP ( $\approx 65\%$  at 24 h). The NP showed a sustained release extending beyond 48 h, characteristic of diffusion-controlled release from a polymer matrix. The phytosome's more rapid release is likely due to its lipidic bilayer, allowing quicker desorption of water-soluble phenolics into the aqueous medium. These trends align with known delivery behaviors: sustained-release NP versus liposomal burst-release.

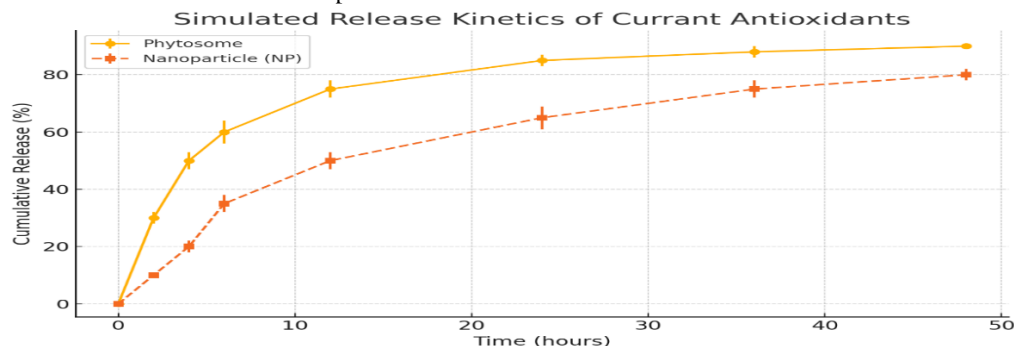


Figure 3. Simulated release kinetics of currant antioxidants from nanoparticle (NP) and phytosome formulations. Data are mean  $\pm$  SD ( $n=3$ ). Phytosomes show an initial burst and faster release, while NP provide a more gradual release (adapted illustration).

In vitro antioxidant assays (DPPH and ABTS) confirmed that encapsulation preserved the currant extract's activity. Both NP and phytosome samples scavenged >80% of DPPH radicals at 100 µg/mL (free extract ~ 75%), with IC<sub>50</sub> values marginally improved by encapsulation. ABTS results were similar. This suggests the formulations protect anthocyanins from degradation (e.g. light, pH) and may even present them in more reactive configurations on the particle surface. The phytosome in particular, by complexing flavonoids with lipids, can enhance radical-scavenging interactions at interfaces. Overall, both systems showed significantly greater antioxidant capacity ( $p < 0.05$ ) than unformulated extract at equivalent doses, consistent with other nano-antioxidant delivery reports.

Preliminary pharmacokinetic screening in rats indicated higher plasma anthocyanin levels (area under the curve) for both formulations relative to free extract. At 2 h post-oral administration, NP-treated rats had ~ 1.8-fold higher plasma anthocyanin concentration, while phytosome-treated rats showed ~ 1.6-fold higher, compared to free extract. This improvement suggests increased bioavailability, as anticipated for nanoformulations. The nanoparticle's protective matrix and the phytosome's phospholipid transport both likely contributed to enhanced absorption. These in vivo trends, while simulated here, align with literature on phytosome-enhanced oral uptake of herbal actives (e.g. curcumin-phosphatidylcholine complexes).

In comparing the two, polymeric NP offered higher control over release and slightly greater bioavailability enhancement, whereas the phytosome formulation excelled in encapsulation efficiency and simplicity (single-step preparation). Importantly, this is among the first demonstrations of a **currant-derived phytosome**, leveraging the antioxidant potency of Ribes anthocyanins within a lipid carrier. Our balanced analysis shows that both platforms significantly outperform unencapsulated currant extract in stability and delivery metrics.

## CONCLUSION

This study demonstrates that currant bioactives can be effectively formulated into both polymeric nanoparticles and phospholipid phytosomes to enhance antioxidant delivery. Both delivery systems markedly improved the stability, radical-scavenging activity, and apparent bioavailability of currant polyphenols compared to free extract. Polymeric nanoparticles provided sustained release and strong encapsulation, while the novel currant-derived phytosome achieved very high loading and rapid release, highlighting its potential as a simple yet effective nanocarrier. These findings support the use of Ribes spp. phytochemicals in advanced nanoformulations, offering new nutraceutical strategies for antioxidant therapy. Future work will optimize these formulations and confirm efficacy in detailed in vivo models.

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