Design, preparation and characterization of lactoferrinloaded sulfobutylether-\beta-cyclodextrin/chitosan nanoparticles as a therapeutic alternative for keratoconus treatment



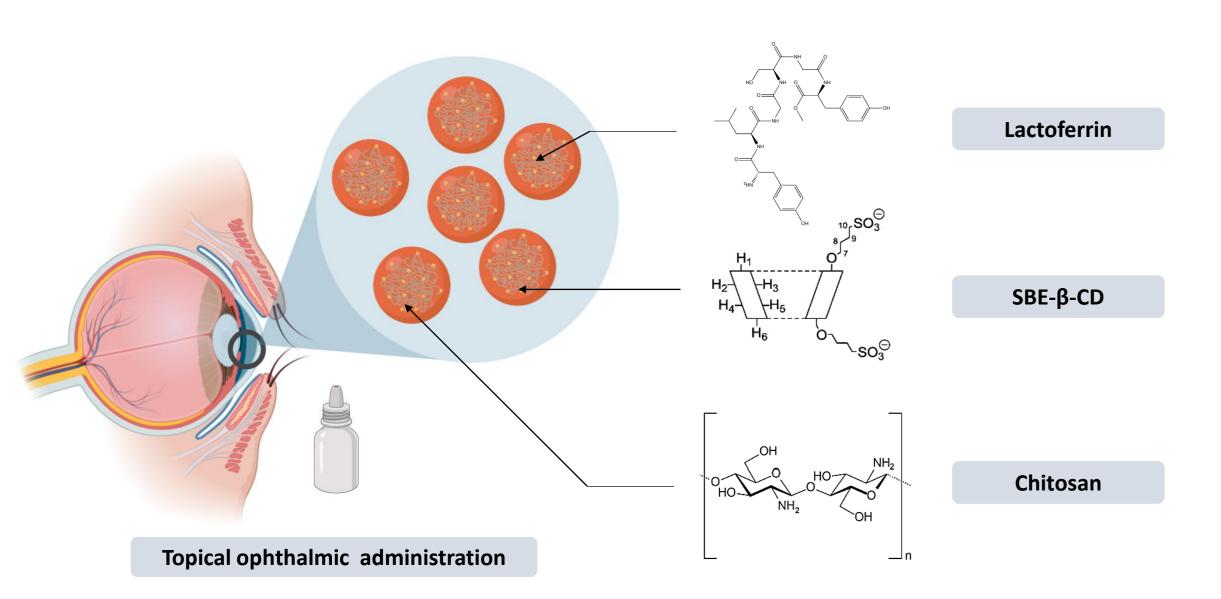
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Introduction and objectives

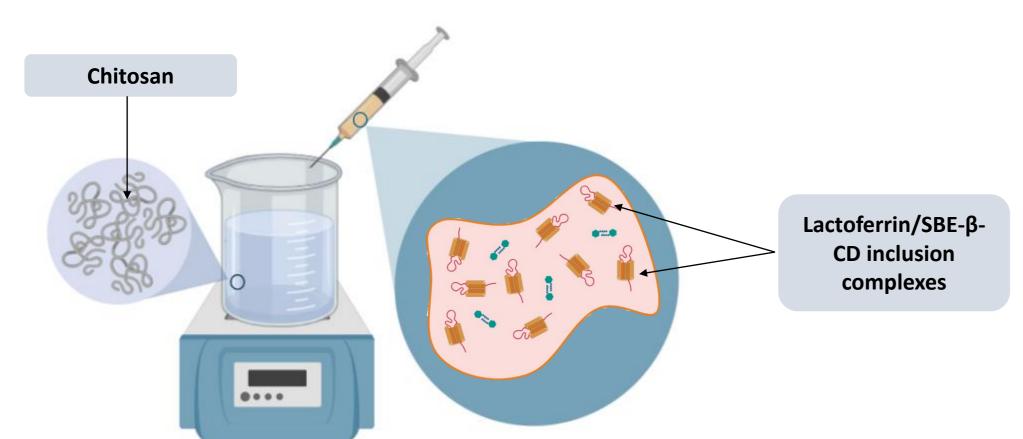
Lactoferrin has shown potential as a good therapeutic alternative in the treatment of Keratoconus [1]. Chitosan/Cyclodextrin nanoparticles as novel drug delivery systems (DDS) could successfully encapsulate hydrophobic drugs [2]. The aim of this work was based on the design, preparation, and characterization of lactoferrin-loaded CS/SBE-β-CD nanoparticles as topical ophthalmic DDS for the keratoconus treatment.



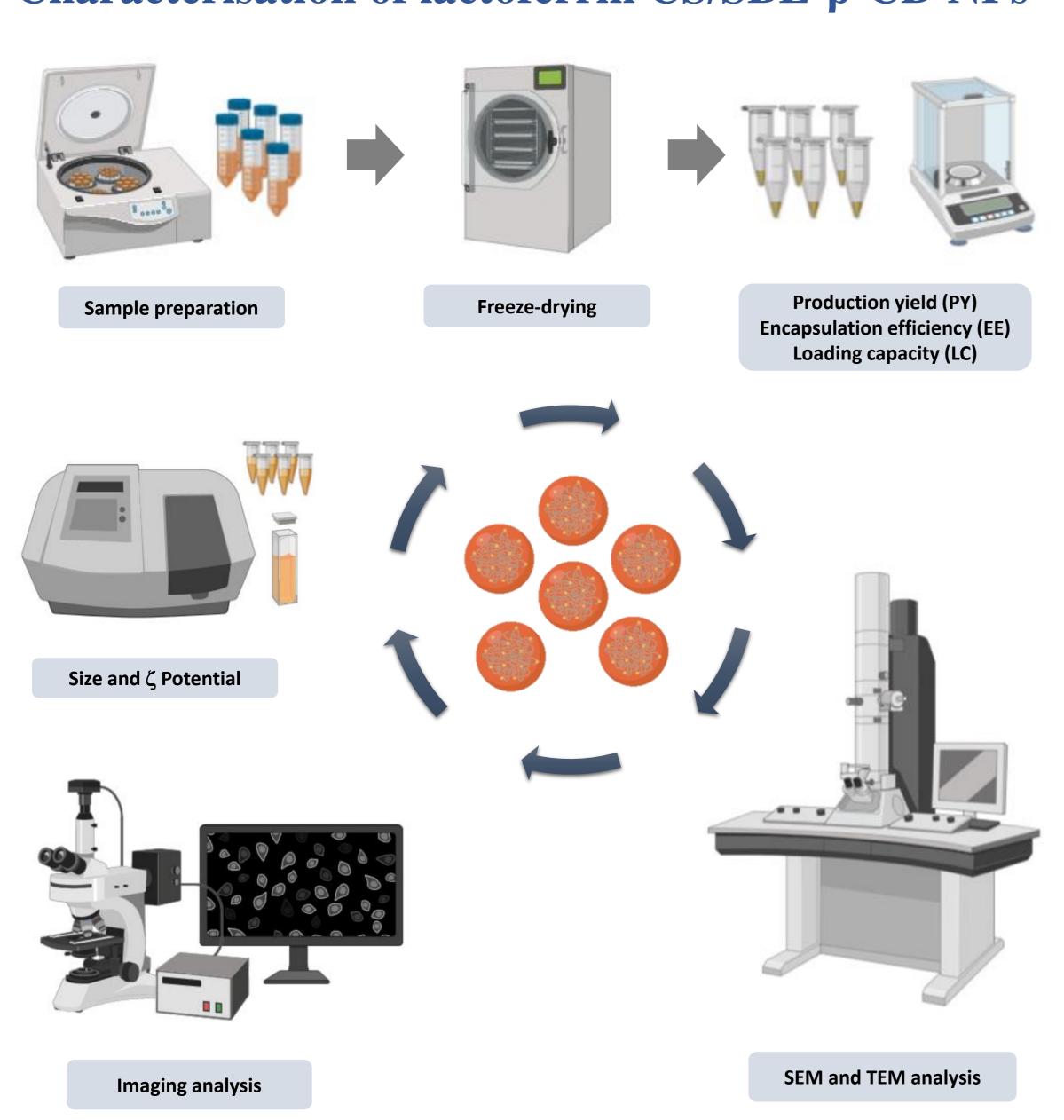
Methodology

Preparation of lactoferrin CS/SBE-β-CD NPs

Nanoparticles were spontaneously obtained via ionotropic gelation. 1 ml lactoferrin/SBE-β-CD aqueous solution was added to 3 ml CS acidic aqueous solution under magnetic stirring at room temperature.

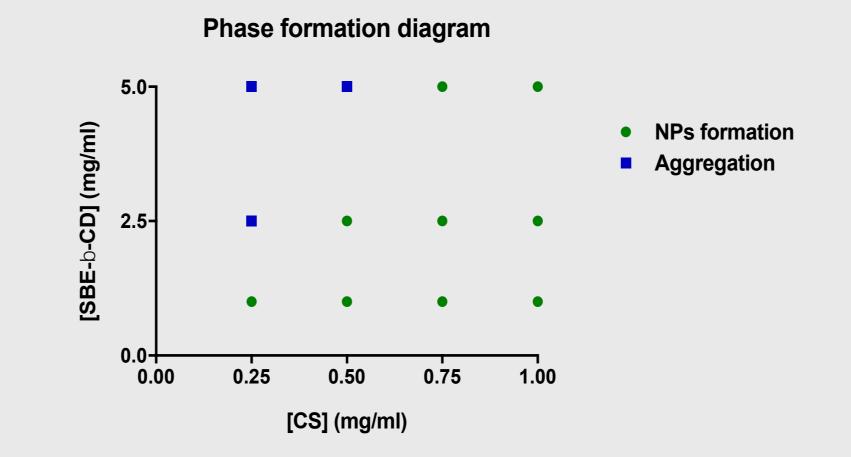


Characterisation of lactoferrin CS/SBE-\beta-CD NPs

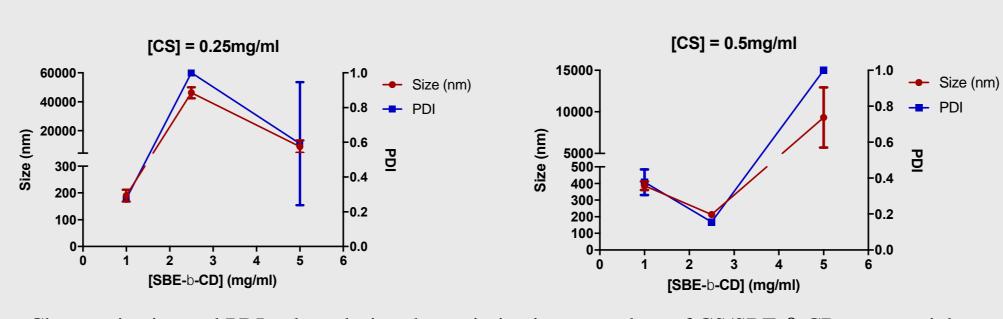


Results

Phase diagram of nanoparticle's formation and physicochemical characterization



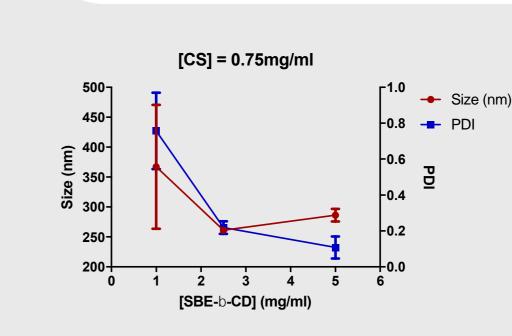
Phase diagram formation for lactoferrin-loaded CS/SBE-β-CD nanoparticle's formation.

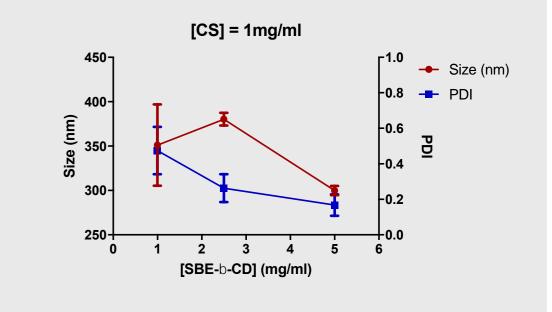


Changes in size and PDI values during the optimization procedure of CS/SBE-β-CD nanoparticles.

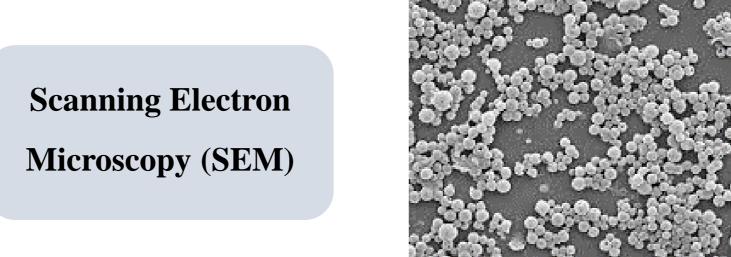
Phase diagram reveals that only CS/SBE-β-CD specific ratios lead to nanoparticle's formation. The appearance of opalescence was used as an indicator of nanoparticle formation, also confirmed by Dynamic Light Scattering (DLS).

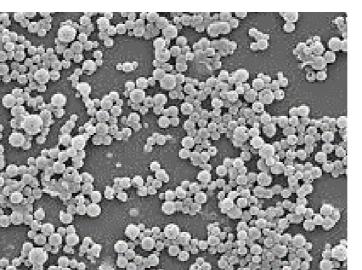
Low amounts of initial CS/SBE-β-CD give rise to no nanoparticle's formation, while precipitation occurred when high amounts of initial compounds were used.

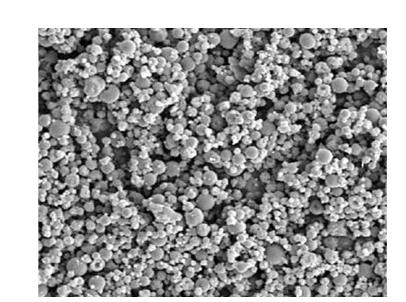


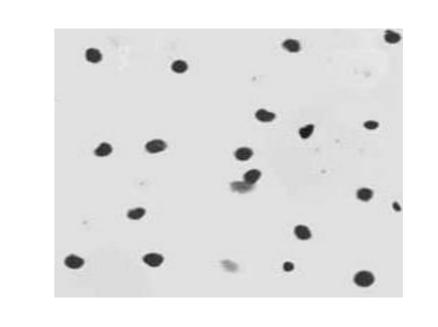


Morphological characterisation of nanoparticles





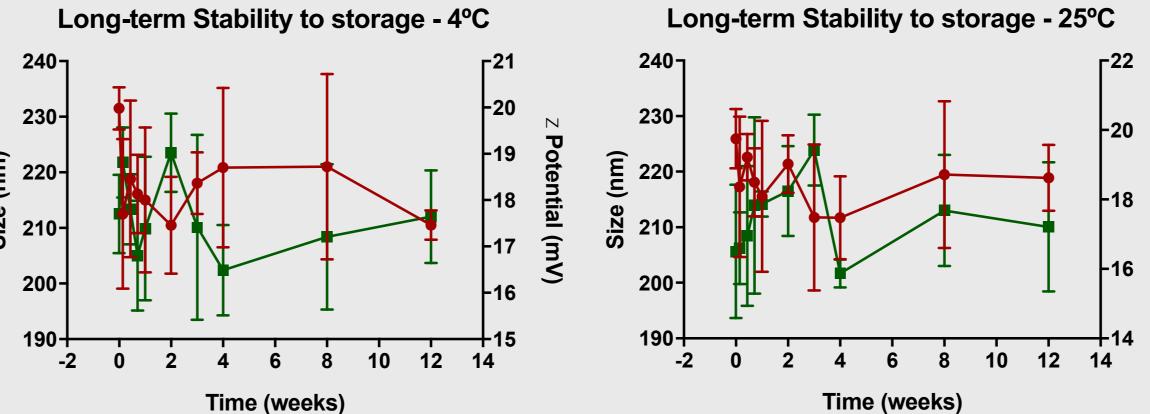


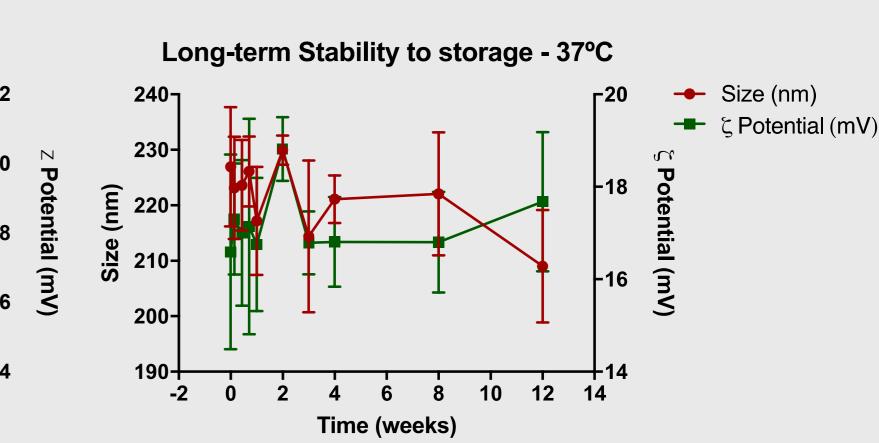


Scanning electron micrographs show that lactoferrin-loaded CS/SBE**β-CD** nanoparticles were predominantly spherical in shape.

Transmission electron microscopy images indicate that these nanosystems may have an irregular surface.

Long-term stability to storage study





Conclusion

Transmission

Electron Microscopy

(TEM)

Lactoferrin-loaded CS/SBE-β-CD nanoparticles were proposed as a new ocular drug delivery system with the virtue of easy administration, prolonged drug release time, improved ocular bioavailability and reduced dosing frequency. Lactoferrin CS/SBE-β-CD nanoparticles show considerable potential as hydrophilic drug carrier and have the capacity to deliver drugs to specific target sites, possibly revolutionizing the Keratoconus therapy.

Bibliography

[1] Mas Tur V, MacGregor C, Jayaswal R, O'Brart D, Maycock N. A review of keratoconus: Diagnosis, pathophysiology, and genetics. Surv Ophthalmol. 2017 Dec;62(6):770–83. [2] Jingou J, Shilei H, Weiqi L, Danjun W, Tengfei W, Yi X. Preparation, characterization of hydrophilic and hydrophobic drug in combine loaded chitosan/cyclodextrin nanoparticles and in vitro release study. Colloids Surf B Biointerfaces. 2011;83(1):103-7.