



## Supplementary Materials: In Vitro Evaluation of Curcuminand Quercetin-Loaded Nanoemulsions for Intranasal Administration: Effect of Surface Charge and Viscosity

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> On Figure S1, the EPR spectra of 16-DSA in the empty and loaded systems are presented.



**Figure S1.** EPR spectra of 16-DSA in the empty and loaded systems. Experimental conditions were center field: 0.349 T, scan range: 0.01 T, gain: 2.24 × 103, time constant: 5.12 ms, modulation amplitude: 0.4 mT and frequency: 9.78 GHz. The measurements were conducted at room temperature. monolayer.

On Figure S2, the EPR spectra of 5-DSA in the empty and loaded systems are presented.



**Figure S2.** EPR spectra of 5-DSA in the empty and loaded systems. Experimental spectra are plotted with solid lines and the corresponding simulated with red dotted line. Experimental conditions were center field: 0.349T, scan range: 0.01 T, gain: 5.64 × 103, time constant: 5.12 ms, modulation amplitude: 0.4 mT and frequency: 9.78 GHz. The measurements were conducted at room temperature.

Concerning photostability of quercetin and curcumin when loaded in negatively charged nanoemulsions under refrigerated conditions results are presented in Table S2.

Days	Quercetin		Curcumin	
	Average Content (%)	SD	Average Content (%)	SD
0	100.0	-	100.0	-
30	99.9	2.8	99.9	2.5
60	95.6	2.5	98.6	2.9
120	36.7	0.8	45.1	1.7

**Table S1.** Stability of Curcumin and Quercetin loaded nanoemulsion (CQ\_NE–) stored at 5 °C–8 °C.

SD: Standard Deviation, n = 3.

**Table S2.** Stability of Curcumin and Quercetin loaded nanoemulsion (CQ\_NE+) stored at 5 °C–8 °C.

	Quercetin		Curcumin	
Days	Average Content (%)	SD	Average Content (%)	SD
0	100.0	-	100.0	-
7	96.9	2.7	83.3	3.0
14	97.0	2.6	85.9	2.4
30	69.4	4.1	72.7	5.9

SD: Standard Deviation, n = 3.

In the present work, a preliminary study comparing the incorporation of CC or BC into the original negatively charged formulation by varying the amount of CC and BC was performed (Table S3), and the effect was evaluated on the size, polydispersity index (PDI), and zeta potential.

**Table S3.** Size, polydispersity index and zeta potential of the formulations prepared with cetalkonium chloride and benzalkonium chloride.

Nanoemulsion	Benzalkonium chloride (%)	Size (nm)	PDI <sup>1</sup>	ζ-potential (mV)	
Blank 1 – BC	0.0100	$126.9 \pm 1.02$	0.208	$-29.70 \pm 0.21$	
Blank 2 – BC	0.0175	$107.4\pm0.46$	0.189	$-2.89 \pm 0.18$	
Blank 3 – BC	0.0250	$117.2 \pm 1.12$	0.207	$+3.81 \pm 0.28$	
Blank 4 – BC	0.0500	$133.3\pm0.43$	0.217	$+17.40 \pm 0.15$	
Blank 5 – BC	0.1000	$125.1 \pm 1.11$	0.190	$+41.06 \pm 0.41$	
	Cetalkonium				
	chloride (%)				
Blank 1 – CC	0.0100	$160.3 \pm 1.34$	0.250	$-3.48 \pm 0.32$	
Blank 2 – CC	0.0175	$131.8\pm0.85$	0.246	$+6.45 \pm 0.21$	
DDL - a la dian anaita in day					

<sup>1</sup> PDI, polydispersity index.