



Article Oleuropein-Laded Ufasomes Improve the Nutraceutical Efficacy

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Abstract: Ufasomes are unsaturated fatty acid liposomes made up of oleic and linoleic acids, natural components required in various biological processes. This kind of nanocarrier is characterized by a simple and dynamic structure and is able to improve the bioavailability of unsaturated fatty acids. The aim of this investigation was to evaluate ufasomes as natural compound delivery systems to deliver oleuropein and improve its antioxidant activity. Oleuropein is a phenolic compound mainly present in olives and olive oil, with several biological properties, such as the antioxidant activity. However, to improve their biological activity, antioxidant compounds should be able to cross cell membranes and uniformly incorporate in cells. Because of the great similarity between their constituents and cell membranes, ufasomes could be advantageous carriers for oleuropein delivery. The physico-chemical characteristics of ufasomes were investigated. A regular shape was shown by transmission electron microscopy studies, while the mean sizes were dependent on the ufasomes composition. In vitro studies highlighted that empty ufasomes did not lead to cell mortality at the tested concentrations and a good carrier internalization in CaCo-2 cells, further studies in vitro studies demonstrated that oleuropein-loaded ufasomes were able to enhance the antioxidant activity of the free active substance making this carrier a suitable one for nutraceutical application.

Keywords: ufasomes; linoleic acid; oleic acid; oleuropein; antioxidant activity; unsaturated fatty acid liposomes

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Figure S1. CLSM micrographs of CaCo-2 human colorectal adenocarcinoma cells. Panel **A**: TRITC filter; panel **B**: Hoechst filter; panel **C**: overlay. No autofluorescence phenomenon was observed.



Figure S2. Antioxidant activity evaluation expressed as cell viability (**A**) and LDH released (**B**). The tests were carried out on CaCo-2 cells treated with empty Ufasomes (Formulation A), using the same concentrations necessary to deliver oleuropein at 10, 20 and 40 μ M, for 24 h and then with H₂O₂ (700 μ M) for 1 h. The results are the average of three independent experiments ± standard deviation. No significant antioxidant effect of blank formulation A was obtained. All results are not statistically significant compared to CTRL (+).