

The pleural cavity retains liposomal drugs providing extended topical drug bioavailability Antonia Marazioti ^{1#,}, Konstantina Papadia ², Georgios T Stathopoulos ³ and Sophia G Antimisiaris ^{1,2*}

¹ Institute of Chemical Engineering Sciences, FORTH/ICE-HT, Rio 26504, Greece

²Laboratory of Pharmaceutical Technology, Department of Pharmacy, University of Patras, Rio 26510, Greece

³Laboratory for Molecular Respiratory Carcinogenesis, Department of Physiology, Faculty of Medicine, University of Patras, Rio 26510, Greece # Presenting author: Antonia Marazioti, email: amarazioti@upatras.gr * Corresponding author: Sophia Antimisiaris, email: santimis@upatras.gr

ABSTRACT

Liposomes are well known for their potential applications as efficient carriers for drug delivery, with the most common administration route so far being intravenous administration. Little is known about the potential therapeutic advantages as well as the retention of topically administered liposomal drugs into confined body cavities. In the present study the retention of various types of liposomes in the pleural cavity following their intrapleural administration was monitored by live animal imaging. Different formulations of liposomes all incorporating DiR in the lipid membrane, were prepared, characterized and injected intrapleurally in healthy and pleural diseased mice. Bodily distribution of the DiR was followed by Biofluorescence imaging. The experimental results reveal that certain liposome preparative parameters significantly affect the local bioavailability of liposomes. However, the most important factor that prolonged liposome retention in the pleural cavity was the lipid dose injected. Moving a step further, we administered intrapleurally a liposomal formulation of the drug deltarasin, which is a novel KRAS inhibitor, to an experimental mouse model of malignant pleural effusion (MPE). A single injection was enough to halt MPE accumulation in mice indicating the high potential of this route of liposome administration as a method to dramatically increase the therapeutic potential of liposomal drugs for local diseases.

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