

Abstract Preview of 'Bile Acid Modified Lipopolymers' (63DWYR)

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Cholic Acid Modified Novel Lipopolymers: An Efficient Transfecting Agents Bramhanand Dube^{a,b} and Hasan Uludag^s, ^aChemical and Materials Engineering, U. of Alberta, Canada, T6G 2G6; ^bKrutika Sawant, ^tTIFAC-CORE in NDDS, Department of Pharmacy, The M S University of Baroda, India; ^sBasak Sahin and ^cCezary Kucharski

The bottleneck for treatment of genetic disorder is lack of an efficient and safe non-viral carrier for genes. The bile acids and their derivatives are known to interact and destabilize the membranes (cell and endosomal membrane) owing to their amphiphilic characters. In the present work, we synthesized novel nonviral vectors for gene delivery by conjugating the cholic acid to cationic polymers and explored their performance for in vitro transfection in two cell lines (293T cells and human cord blood mesenchymal stem cells). Cholic acid (CA) was conjugated with polyethylenimine (PEI, 2kDa) and polyallylamine (PAA, 15kDa) using carbodiimide chemistry at 4 different substitution ratios and characterized by 1H-NMR spectra along with their DNA binding capacities. The binding results showed a reduced binding as a result of cholic acid substitution. In vitro transfection results in 293T cells are promising as PEI 2K-CA shows several hundred times more transfection efficiency as compared to native PEI 2kDa and significantly more than PEI 25 kDa, whereas PAA-CA did not show significant increase in transfection efficiency as compared to native PAA. On the other hand, the synthesized polymer conjugates showed less cytotoxicity than PEI 25kDa in the same cell line as evaluated by MTT assay.

Based on our findings it can be concluded that mere conjugation of bile acid to cationic polymer is not only responsible for increased transfection efficiency but also the polymer should have some structural and functional qualifications.

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