

Novel inhalable formulation based on hybrid lipid-polymer nanoparticles for pulmonary siRNA delivery

Marta Cabibbo,¹ Emanuela Fabiola Craparo,¹ Emanuele Salvatore Drago,¹ Simone P. Carneiro,² Gaetano Giammona,¹ Gennara Cavallaro,¹ Olivia Merkel.²

¹Department of Biological, Chemical and Pharmaceutical Science and Technologies (STEBICEF), University of Palermo, Via Archirafi 32, 90123 Palermo, Italy

²Affiliation Department of Pharmacy, Pharmaceutical Technology and Biopharmacy, Ludwig-Maximilians-University, Butenandtstrasse 5-13, 81337 Munich, Germany

Inhalation-based siRNA delivery has a unique potential for the treatment of several lung diseases, by silencing the expression of the target gene in a post-transcriptional way. However, for optimum siRNA pulmonary delivery, an appropriately designed delivery system is required. [1] [2]

Here, novel hybrid lipid-polymer nanoparticles (LPHNPs), comprising a fluorescent cationic polyaspartamide-poly(lactic-co-glycolic) acid (PHEA-DY700-bAPAE-PLGA) core and a lipid shell of 1,2-distearoyl-sn-glycero-phosphoethanolamine-N-(polyethyleneglycol)₂₀₀₀ (DSPE-PEG₂₀₀₀), were prepared by emulsion/solvent diffusion for pulmonary delivery of siRNA. These exhibit colloidal size, a negative ζ potential, 50 wt % phospholipid, and spherical shape. They are able to incorporate successfully siGFP and interact slightly with mucin. LPHNPs are well internalized into lung cancer cells and do not exert any cytotoxic effect. Moreover, to achieve an inhalable formulation, the nano-into-micro strategy was applied.

References:

[1] Ling Ding, Siyuan Tang, Todd A. Wyatt, Daren L. Knoell, David Oupický, Pulmonary siRNA delivery for lung disease: Review of recent progress and challenges, *Journal of Controlled Release*, 330 (2021) 977–991.

[2] Jenny Ka-Wing Lam, Wanling Liang, Hak-Kim Chan, Pulmonary delivery of therapeutic siRNA, *Advanced Drug Delivery Reviews* 64 (2012) 1–15.

marta.cabibbo@unipa.it |

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