Ln(III) Doped Nanoparticles Interaction With Human Blood Plasma. DLS and Luminescent Spectroscopy Study

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Extended Abstract

Nanoparticles are suitable platforms for cancer targeting and diagnostic applications. Typically, less than 10% of all systemically administered nanoparticles accumulate in the tumour. Here we explore the interactions of blood components with nanoparticles and describe how these interactions influence solid tumour targeting. In the blood, serum proteins adsorb onto nanoparticles to form a protein corona in a manner dependent on nanoparticle physicochemical properties. These serum proteins can block nanoparticle tumour targeting ligands from binding to tumour cell receptors. Additionally, serum proteins can also encourage nanoparticle uptake by macrophages, which decreases nanoparticle availability in the blood and limits tumour accumulation. The formation of this protein corona will also increase the nanoparticle hydrodynamic size or induce aggregation, which makes nanoparticles too large to enter into the tumour through pores of the leaky vessels, and prevents their deep penetration into tumours for cell targeting. Recent studies have focused on developing new chemical strategies to reduce or eliminate serum protein adsorption, and rescue the targeting potential of nanoparticles to tumour cells. An in-depth and complete understanding of nanoparticle-blood interactions is key to designing nanoparticles with optimal physicochemical properties with high tumour accumulation (J. Lazarovits et al., 2015).

The silica nanoparticles doped with Tb-thiacalix[4]arene complex synthesised according to Stober and microemulsion (A.R. Mukhametshina et al., 2014) strategies as well as Tb-betadiketone substituted calix[4]arene (N.A. Shamsutdinova et al., 2014) and Eu(TTA)3-phosphine oxide polyelectrolyte coated nanoparticles' (A. Mustafina et al., 2011) size, zeta potential and photophysical properties were studied in the presence of BSA and blood plasma. The impact of the nature of polyelectrolyte exterior layer and phospholipid bilayer formation on the nanoparticles surface on the studied parameters are discussed.

- Lazarovits, J., Chen, Y.Y., Sykes, E.A., & Chan, W.C.W. (2015). Nanoparticle-Blood Interactions: The Implications On Solid Tumour Targeting. *Chem. Commun.*, 51, 2756-2767.
- Mukhametshina, A.R., Mustafina, A.R., Davydov, N.A., Nizameev, I.R., Kadirov, M.K., Gorbatchuk, V.V., & Konovalov, A.I. (2014). The Interfacial Interactions Of Tb-Doped Silica Nanoparticles With Surfactants And Phospholipids Revealed Through The Fluorescent Response. *Colloids Surf. B*, 115, 93-99.
- Mustafina, A., Zairov, R., Gruner, M., Ibragimova, A., Tatarinov, D., Nizameyev, I., Nastapova, N., Kadirov, M., Mironov, V., & Konovalov, A. (2011). Synthesis And Photophysical Properties Of Colloids Fabricated By The Layer-By-Layer Polyelectrolyte Assembly Onto Eu(III) Complex As A Core. Coll. Surf. B: Biointerfaces, 88, 490-496.

Shamsutdinova, N.A., Podyachev, S.N., Sudakova, S.N., Mustafina, A.R., Zairov, R.R., Burilov, V.A., Nizameev, I.R., Rizvanov, I.K., Syakaev, V.V., Gabidullin, B.M., Katsuba, S.A., Gubaidullin, A.T., Safiullin, G.M., & Dehaen, W. (2014). A Facile Synthetic Route To Convert Tb(III) Complexes Of Novel Tetra-1,3-Diketone Calix[4]Resorcinarene Into Hydrophilic Luminescent Colloids. *New J. Chem.*, 38, 4130-4140.