Abstract

Modern diagnostic method magnetic resonance imaging (MRI) usually uses contrast agents T_1 -type, which are based on Gd^{3+} complexes. Due to severe toxicity of free Gd^{3+} , it is desired to have thermodynamically stable and kinetically inert complexes with fast elimination from the body. This work summarizes information about a novel contrast agent based on ligand DO3A^P (1,4,7,10-tetraazacyclododecane-1-methyl(alkyl)phosphinic-4,7,10triacetic acid) with pendant hydrophobic dibenzylamino group which is able to interact hydrophobically with the macromolecule of serum albumin. The stability of supracomplex is dependent on pH value, i.e. on the protonation of the pendant amino group of the complex (p K_A = 5.6) and this interaction was confirmed from 1H -NMRD profile and fluorescent analysis. The compound was tested for its angiographic properties in vivo on rat model. Furthermore, other complexes of the ligand with trivalent lanthanides (Nd³⁺, Eu³⁺, Tb³⁺, Dy³⁺, Er³⁺) were characterized by various methods (XRD, luminescence, UV-VIS, ¹H-, ¹⁷O- and ³¹P-NMR). The cleavage of the benzyl groups affords ligand whose Ln³⁺ complexes possess pH dependent PARACEST effect. These complexes were characterized by XRD, luminescence and ¹H- and ³¹P-NMR. Moreover, the novel ligands with modified length of pendant arm and with dibenzylamino group were synthesized. The altered value of pK_A was determined by NMR titration method.

Keywords

MRI, gadolinium, macrocyclic ligands, macrocyclic complexes, serum albumin