

Supporting Information

Manuscript title: A T_1 , T_2 magnetic resonance imaging (MRI)-fluorescent imaging (FI) by using ultrasmall mixed gadolinium-europium oxide nanoparticles

Authors: Wenlong Xu, Ja Young Park, Krishna Kattel, Badrul Alam Bony, Woo Choul Heo, Seonguk Jin, Jang Woo Park, Yongmin Chang, Ji Yeon Do, Kwon Seok Chae, Tae Jeong Kim, Ji Ae Park, Young Woo Kwak, and Gang Ho Lee

(1) XRD pattern

An XRD pattern of the as-prepared powder sample is shown in Fig. S1. Among the known three phases (*i.e.*, $\text{Ln}(\text{OH})_3$, LnOOH , and Ln_2O_3 , $\text{Ln} = \text{Gd}$ and Eu),¹ the obtained XRD pattern is close to that of Ln_2O_3 . The broad XRD pattern is due to ultrasmall particle diameters of nanoparticles, similarly to that observed in ultrasmall Gd_2O_3 nanoparticles.² After TGA analysis up to 900 °C, however, sharp peaks (222), (400), (440), and (622) corresponding to a highly crystallized form of Ln_2O_3 with a cubic structure were observed (Fig. S1). Note that the peaks from both Gd_2O_3 and Eu_2O_3 occur at very similar positions and thus, can not be distinguished from each other in the XRD pattern. The estimated cell constant ($a = 10.84 \text{ \AA}$) is consistent with the weighted average of those of Gd_2O_3 ($a = 10.81 \text{ \AA}$) and Eu_2O_3 ($a = 10.86 \text{ \AA}$) given by JCPDS-International Center for Diffraction Data, PCPDFWIN, Version 1.30.³

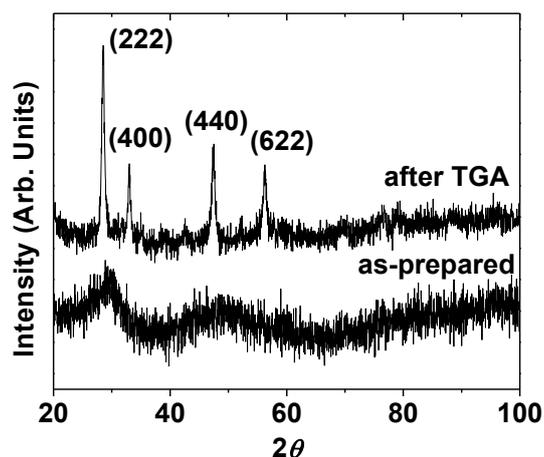


Fig. S1 XRD patterns of powder samples of lactobionic acid coated ultrasmall mixed gadolinium-europium oxide nanoparticles: (bottom) as-prepared and (top) after TGA analysis up to 900 °C. The assignments are the Miller indices (hkl).

(2) 3 tesla T_1 and T_2 MR images

A series of 3 tesla coronal T_1 MR images of a mouse kidney and artery are shown in Fig. S2. Positive contrast enhancements can be clearly seen after injection of an aqueous sample solution into a mouse tail vein. After 30 minutes, however, signal decayed away due to excretion of nanoparticles through kidneys. A series of 3 tesla axial T_2 MR images of a mouse liver are also shown in Fig. S3. Negative contrast enhancements can be clearly seen after injection. The negative contrast enhancement in liver was kept longer than the positive contrast enhancement in kidney because clearance of nanoparticles through liver takes a longer time than through kidneys.

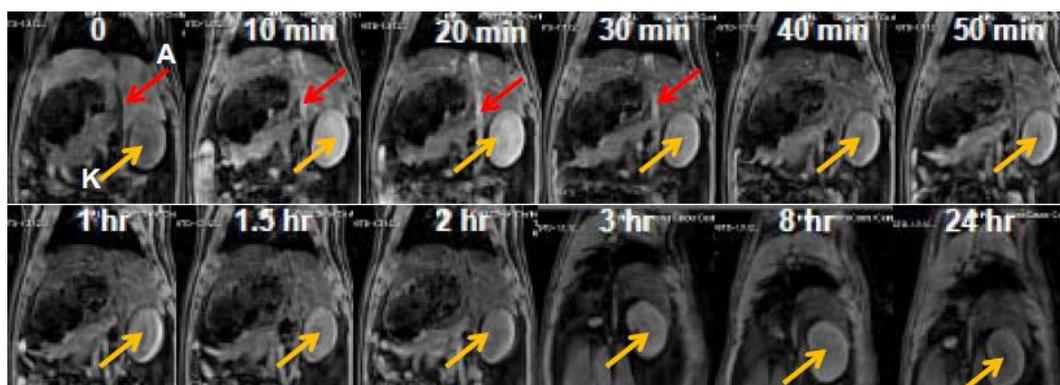


Fig. S2 A series of 3 tesla coronal T_1 MR images of kidney (indicated as K) and artery (indicated as A) before and after injection of a sample solution into a mouse tail vein.

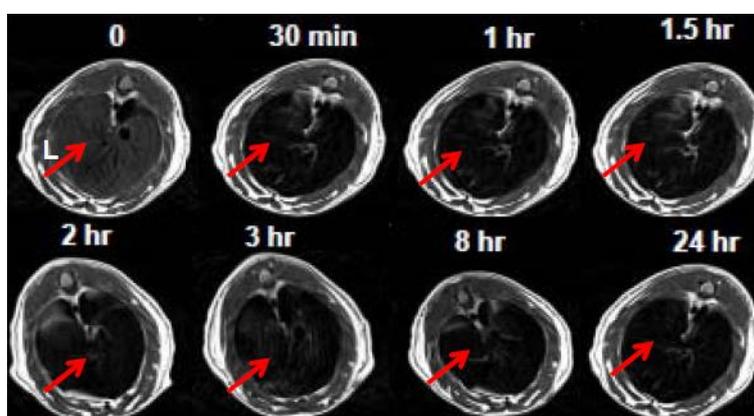


Fig. S3 A series of 3 tesla axial T_2 MR images of liver (indicated as L) before and after injection of a sample solution into a mouse tail vein.

(3) Solubility and stability of nanoparticles in sample solutions

Sample solutions of lactobionic acid coated nanoparticles dispersed in PBS solution, DMEM solution used for incubation of NCTC1469 cell, and RPMI1640 solution used for incubation of DU145 cell were tested for solubility and stability of lactobionic acid coated nanoparticles in them. The lactobionic acid coated nanoparticles were completely dispersed in all solutions, implying that they are soluble in all solutions. In case of stability, we measured the precipitation times of lactobionic acid coated nanoparticles at various Gd&Eu concentrations after they were dispersed in

solutions (Fig. S4). The stability of lactobionic acid coated nanoparticles, depending on the solution type, was decent. Pictures of sample solutions (Gd&Eu concentration = 11 mM) after dispersion and after precipitation of lactobionic acid coated nanoparticles in solutions are provided in Fig. S5. The less stability in PBS solution than those in both DMEM and RPMI1640 solutions is likely related to the salt effect of PBS solution.

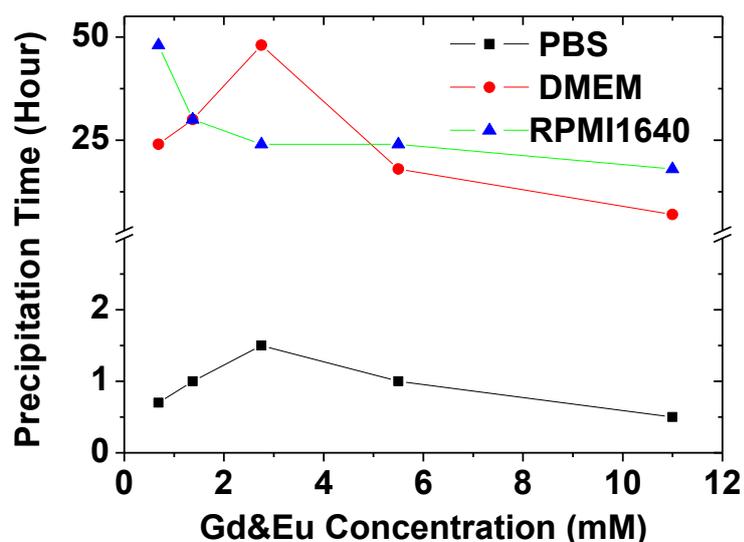


Fig. S4 Stability tests of sample solutions of lactobionic acid coated nanoparticles dispersed in three solutions (PBS, DMEM, and RPMI1640).

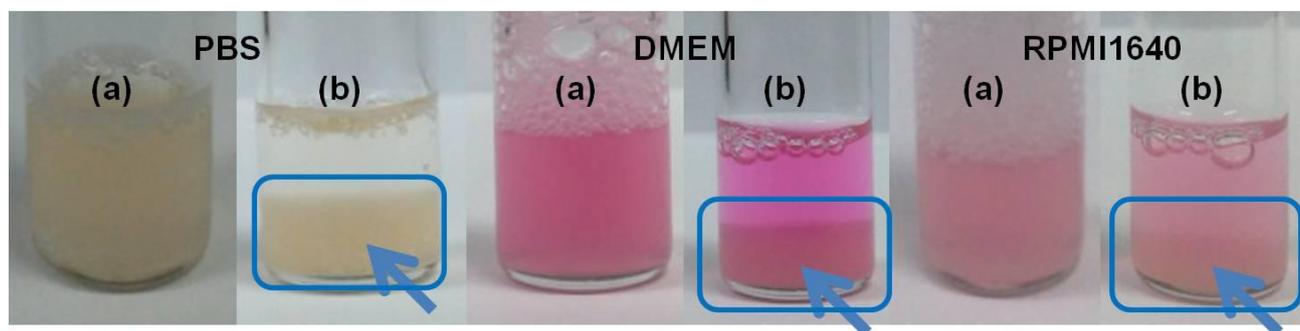


Fig. S5 Pictures of sample solutions (a) after dispersion and (b) after precipitation of lactobionic acid coated nanoparticles in solutions. Square and arrow indicate the area of precipitation.

References

- 1 C. Chang and D. Mao, *Int. J. Chem. Kinet.*, 2007, **39**, 75-81.
- 2 F. Söderlind, H. Pedersen, R. M. Petoral Jr., P. -O. Käll and K. Uvdal, *J. Colloid Interface Sci.*, 2005, **288**, 140-148.
- 3 JCPDS - International Centre for Diffraction Data, card no. 43-1008 ($a = 10.86 \text{ \AA}$) and 43-1014 ($a = 10.81 \text{ \AA}$) for Eu_2O_3 and Gd_2O_3 , respectively.