## **Supporting Information**

## Sequential growth of CaF<sub>2</sub>:Yb,Er@CaF<sub>2</sub>:Gd nanoparticles for efficient magnetic resonance angiography and tumor diagnosis

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(a)	Spectrum	1 Area 1		(c)	Spectrum	2	Area 2
600 nm				600 nm			
(b)	Element	Weight%	Atomic%	(d)	Element	Weight%	Atomic%
	F	16.04	39.42		F	0	0
	Ca	8.43	9.82		Са	0	0
	Cu	65.2	47.91		Cu	100	100
	Gd	2.41	0.71		Gd	0	0
	Er	0	0		Er	0	0
	Yb	7.93	2.14		Yb	0	0
	Totals	100	100		Totals	100	100

**Fig. S1.** The detailed elemental compositions of the as-prepared CaF<sub>2</sub>:Yb,Er@CaF<sub>2</sub>:Gd nanoparticles at different transmission electron microscopy (TEM) image area (a, b were at nanoparticles area 1 and c, d were at blank area 2).



**Fig. S2.** The X-ray photoelectron spectra (XPS) of the as-prepared CaF<sub>2</sub>:Yb,Er@CaF<sub>2</sub>:Gd nanoparticles: (a) Survey spectrum, (b) Ca 2p spectrum, (c) Gd 4d spectrum, (d) F 1s spectrum.



**Fig. S3.** FT-IR spectra of the CaF<sub>2</sub>:Yb,Er@CaF<sub>2</sub>:Gd-OA (black line), the hydrophilic CaF<sub>2</sub>:Yb,Er@CaF<sub>2</sub>:Gd nanoparticles(red line) and the PEG-PAA di-block copolymer (blue line).



**Fig. S4.** TEM images of (a) the as-prepared CaF<sub>2</sub>:Yb,Er@CaF<sub>2</sub>:Gd-OA nanoparticles dispersed in cyclohexane and (b) the hydrophilic CaF<sub>2</sub>:Yb,Er@CaF<sub>2</sub>:Gd nanoparticles dissolved in water.



**Fig. S5** Absorption spectra of arsenazo III at different filtrates time points obtained after dialysis of 1 mg Gd/mL (a) GdCl<sub>3</sub> aqueous solution and (b) the hydrophilic CaF<sub>2</sub>:Yb,Er@CaF<sub>2</sub>:Gd nanoparticles aqueous solution to study the leaching of Gd ions from the original aqueous solution .Water and GdCl<sub>3</sub> aqueous solution (1  $\mu$ g Gd/mL, 10  $\mu$ g Gd/mL and 100  $\mu$ g Gd/mL)were used as negative and positive controls, respectively.



**Fig. S6.** Hydrodynamic size of the hydrophilic CaF<sub>2</sub>:Yb,Er@CaF<sub>2</sub>:Gd nanoparticles aqueous solution at different storage time points measured by dynamic light scattering (DLS).



Fig. S7. Zeta potential of the hydrophilic  $CaF_2$ :Yb,Er@CaF\_2:Gd nanoparticles aqueous solutions at different storage time points.



Scheme S1. Schematic illustration of the detail structure difference between as-prepared  $CaF_2$ : Yb, Er@CaF\_2: Gd nanoparticles and CaF\_2: Gd nanoparticles.



**Fig. S8**. (a)  $T_1$  maps of the hydrophilic CaF<sub>2</sub>:Gd nanoparticles; (b) Comparison of longitudinal relaxivity  $r_1$  of the hydrophilic CaF<sub>2</sub>:Gd nanoparticles under 3.0 T.



**Fig. S9** The other variations of indices of hepatic function, renal function and blood routine tests (n=3), including aspartate transaminase / alanine transaminase ratio (AST/ALT), albumin (ALB), blood urea

nitrogen / creatinine ratio (BUN/CREA), uric acid (UA), lactate dehydrogenase (LDH), white blood cells (WBC), red blood cells (RBC), hemo-globin (HGB), hematocrit (HCT), plateletcrit (PCT), mean corpuscular volume (MCV), mean corpuscular hemoglobin (MCH), red blood cell distribution width measured by standard deviation (RDW-SD), red blood cell distribution width measured by variation coefficient (RDW-CV), mean corpuscular hemoglobin concentration (MCHC), platelet distribution width (PDW), lymphocyte (LYMPH), mean platelet volume (MPV), platelet-large contrast ratio(P-LCR), platelets (PLT).

**Table S1**. The detailed elements composition of the hydrophilic CaF<sub>2</sub>:Yb,Er@CaF<sub>2</sub>:Gd nanoparticles measured by means of ICP-AES.

Element	ICP-AES data (μg/mL)	Weight%	Atomic%
Ca	206.2	22.06	54.31
Yb	491.7	52.59	29.94
Er	34.5	3.69	2.17
Gd	202.5	21.66	13.57
Totals		100	100