Supplementary Materials

Simultaneous enhancement of T_1 and T_2 magnetic resonance imaging of liver tumor at respective low and high magnetic fields

Huan Li,^{#1} Zijuan Hai,^{#2} Liwei Zou,¹ Lele Zhang,² Lulu Wang,⁴ Longsheng Wang^{*1} and Gaolin Liang^{*3}

- Department of Radiology, the Second Hospital of Anhui Medical University, Hefei, Anhui 230601, China
- Key Laboratory of Structure and Functional Regulation of Hybrid Materials, Ministry of Education, Institutes of Physical Science and Information Technology, Anhui University, Hefei, Anhui 230601, China
- State Key Laboratory of Bioelectronics, School of Biological Science and Medical Engineering, Southeast University, Nanjing, Jiangsu 210096, China
- High Magnetic Field Laboratory, Hefei Institutes of Physical Science, Chinese Academy of Sciences, Hefei, Anhui 230031, China

[#]These authors contributed equally to this work.

E-mail: wanglongsheng@ahmu.edu.cn (L.-S. W.), gliang@seu.edu.cn (G.-L. L.).

Contents:

1. Supplementary figures and tables

1. Supplementary figures and tables



Figure S1. Statistics of size distribution of DOTA-Gd-CBT-NP in Figure 1B.



Figure S2. Time-course DLS measurement of DOTA-Gd-CBT-NP in PBS buffer.



Figure S3. Cell viability of HepG2 cells incubated with Glu-DOTA-Gd-CBT at different concentrations for 4 h, and 8 h. Each error bar represents the standard deviation of three independent experiments.



dynamic T₁-weighted transverse Figure *S4*. In vivo MR images of Glu-DOTA-Gd-CBT-injected **DON-pretreated** mice (top row), and then Glu-DOTA-Gd-CBT-injected mice (middle row), and Gd-DTPA-injected mice (bottom row) at low magnetic field (1.0 T). White circles indicate the liver tumors.



T₂-weighted Figure *S*5. In vivo dynamic transverse MR images of Glu-DOTA-Gd-CBT-injected row), DON-pretreated mice (top and then Glu-DOTA-Gd-CBT-injected mice (middle row), and Gd-DTPA-injected mice (bottom row) at low magnetic field (1.0 T). White circles indicate the liver tumors.



Figure S6. Normalized time course relative tumor-to-liver (T/L) contrast ratios of T_2 values at low magnetic field (1.0 T) in Figure S5. Each error bar represents the standard deviation of three independent experiments.



dynamic T₁-weighted MR images Figure *S*7. In vivo transverse of Glu-DOTA-Gd-CBT-injected mice (top row), DON-pretreated and then Glu-DOTA-Gd-CBT-injected mice (middle row), and Gd-DTPA-injected mice (bottom row) at high magnetic field (9.4 T). White circles indicate the liver tumors.



Figure S8. Normalized time course relative tumor-to-liver (T/L) contrast ratios of T_1 values at high magnetic field (9.4 T) in Figure S7. Each error bar represents the standard deviation of three independent experiments.



Figure *S9*. dynamic T₂-weighted images In vivo transverse MR of Glu-DOTA-Gd-CBT-injected mice (top DON-pretreated row), and then Glu-DOTA-Gd-CBT-injected mice (middle row), and Gd-DTPA-injected mice (bottom row) at high magnetic field (9.4 T). White circles indicate the liver tumors.



Figure S10. The contents of Gd (μ g/g, determined with ICP-MS) in tumors and

organs of three groups after MRI at 2.5 h.

Table S1. The r_2/r_1 ratio values of **DOTA-Gd-CBT-NP**, **Glu-DOTA-Gd-CBT** and Gd-DTPA at low (1.0 T) and high (9.4 T) magnetic field.

	r_2/r_1 ratio value at 1.0 T	r ₂ /r ₁ ratio value at 9.4 T
DOTA-Gd-CBT-NP	0.91	11.8
Glu-DOTA-Gd-CBT	0.96	0.55
Gd-DTPA	1.58	2.16

Table S2. The r_2/r_1 ratio values of cells in Group **Glu-DOTA-Gd-CBT**, Group "DON + **Glu-DOTA-Gd-CBT**", and Group Gd-DTPA at low (1.0 T) and high (9.4 T) magnetic field.

	r_2/r_1 ratio value at 1.0 T	r_2/r_1 ratio value at 9.4 T
Glu-DOTA-Gd-CBT	0.90	7.83
DON + Glu-DOTA-Gd-CBT	1.58	2.13
Gd-DTPA	1.06	2.15

Table S3. GGT activity in liver tumor lysates.

Diameter of Tumor	Increased O.D. Value at 405 nm	GGT activity (U/L)	
Size (mm)	between 1 min		
10	0.037 ± 0.015	313.3 ± 45.3	

	Heart	Liver	Spleen	Lung	Kidney	Tumor
Glu-DOTA-Gd -CBT	0.32 ± 0.24	6.67 ± 1.37	0.55 ± 0.35	0.78 ± 0.39	8.23 ± 1.78	5.35 ± 1.37
DON+Glu-DO TA-Gd-CBT	0.39 ± 0.16	5.41 ± 1.52	0.62 ± 0.27	0.63 ± 0.31	8.44 ± 1.63	1.91 ± 1.65
Gd-DTPA	0.33 ± 0.23	6.52 ± 1.41	0.72 ± 0.39	0.84 ± 0.41	8.65 ± 1.24	1.22 ± 1.03

Table S4. The contents of Gd (μ g/g, determined with ICP-MS) in tumors and main organs of three groups after MRI at 2.5 h.