Ultra-small manganese dioxide nanoparticles with high T_1 relaxivity for magnetic resonance angiography

Yuting Jiang, ^a Haojie Gu, ^a Zhongyuan Cai, ^a Shengxiang Fu, ^b Yingzi Cao, ^a Lingling Jiang, ^a Changqiang Wu, ^c Wei Chen, ^c Chunchao Xia, ^b Su Lui, ^b Bin Song, ^b Qiyong Gong, ^{d,e} and Hua Ai* ^{a,b}

^a National Engineering Research Center for Biomaterials, Sichuan University, Chengdu, 610064, China

^b Department of Radiology, West China Hospital, Sichuan University, Chengdu, 610041, China

^c Sichuan Key Laboratory of Medical Imaging, School of Medical Imaging, North Sichuan Medical College, Nanchong, 637000, China

^d Huaxi MR Research Center (HMRRC), Department of Radiology, West China Hospital of Sichuan University, Chengdu, 610041, China

^e Psychoradiology Research Unit of Chinese Academy of Medical Sciences, Sichuan University, Chengdu, 610064, China



Figure S1. XPS spectra of MnO₂/PAA50 NPs and Mn 2p (A), MnO₂/PAA100 NPs and Mn 2p (B), MnO₂/PAA150 NPs and Mn 2p (C), MnO₂/PAA300 NPs and Mn 2p (D), MnO₂/PAA400 NPs and Mn 2p (E).

Table S1. TGA results for MnO₂/PAA NPs.

Sample	Weight loss of PAA (%)	
MnO ₂ /PAA50	25.6	
MnO ₂ /PAA100	25.8	
MnO ₂ /PAA150	30.3	
MnO ₂ /PAA200	31.1	
MnO ₂ /PAA300	31.4	
MnO ₂ /PAA400	31.5	

Table S2. The r_1 , r_2 and r_2/r_1 values of MnO₂/PAA NPs under a 0.5 T NMR Analyzing system at 32°C.

Sample	$r_1 (\mathrm{mM}^{-1}\mathrm{s}^{-1})$	r_2 (mM ⁻¹ s ⁻¹)	r_2/r_1
MnO ₂ /PAA50	7.6	10.0	1.3
MnO ₂ /PAA100	9.9	14.8	1.5
MnO ₂ /PAA150	23.1	33.6	1.4
MnO ₂ /PAA200	29.8	40.6	1.3
MnO ₂ /PAA300	28.9	39.5	1.3
MnO ₂ /PAA400	28.5	41.2	1.4

Sample	$r_1 (\mathrm{mM}^{-1}\mathrm{s}^{-1})$	r_2 (mM ⁻¹ s ⁻¹)	r_2/r_1
MnO ₂ /PAA50	6.7	11.9	1.8
MnO ₂ /PAA100	9.6	17.2	1.8
MnO ₂ /PAA150	23.3	42.1	1.8
MnO ₂ /PAA200	29.0	53.3	1.8
MnO ₂ /PAA300	27.9	51.9	1.9
MnO ₂ /PAA400	27.4	51.1	1.9

Table S3. The r_1 , r_2 and r_2/r_1 values of MnO₂/PAA NPs under a 1.5 T MRI scanner at 25°C.

Table S4. The r_1 , r_2 and r_2/r_1 values of MnO₂/PAA NPs under a 3.0 T MRI scanner at 25°C.

Sample	$r_1 (\mathrm{mM}^{-1}\mathrm{s}^{-1})$	r_2 (mM ⁻¹ s ⁻¹)	r_2/r_1
MnO ₂ /PAA50	4.7	11.5	2.4
MnO ₂ /PAA100	6.0	16.5	2.7
MnO ₂ /PAA150	13.9	40.2	2.9
MnO ₂ /PAA200	16.7	51.4	3.1
MnO ₂ /PAA300	16.0	50.2	3.1
MnO ₂ /PAA400	15.6	49.1	3.1



Figure S2. (A) Cell viabilities of Raw264.7 incubated with different concentrations of MnO₂/PAA200 NPs for 24 h; (B) The hemolysis percentage of MnO₂/PAA200 NPs; (C) The UV absorption spectra of blood incubated with deionized water, PBS and MnO₂/PAA200 NPs at different concentrations for 2 h. (D) Biochemical indicators and (E) H&E staining of main issues from SD rats 24 h after administration (0.05 mmol Mn kg/BW). Scale bar: 100 μm.



Figure S3. (A) Pseudocolor MR images of heart and neck of SD rats after injection of Gadovist[®] (Gd-DO3A-Butrol, dosage: 0.1 mmol Gd kg/BW) and MnO₂/PAA200 NPs (dosage: 0.05 or 0.1 mmol Mn kg/BW), respectively at different time points; (B) Percentage-injected dosage per gram of manganese (% ID/g) in blood at different time points with the dosage of 0.05 mmol Mn kg/BW; (C) Percentage-injected dosage of manganese in the feces and urine of SD rats at different time (6 h, 12 h and 24 h) after injection of MnO₂/PAA200 NPs (0.05 mmol Mn kg/BW).