Supporting Information

MR-visible and pH-ultrasensitive micellar nanodrugs for cancer

theranostic application

Bo Li,#,a,b Mingyue Cai,#,a Liteng Lin,#,a Weitong Sun,c Zhimei Zhou, a Shiyin Wang,b

Yong Wang,^b Kangshun Zhu,*,^a and Xintao Shuai**,^b

^aLaboratory of Interventional Radiology, Department of Minimally Invasive Interventional Radiology, and Department of Radiology, the Second Affiliated Hospital of Guangzhou Medical University, Guangzhou, 510260, China.

^bPCFM Lab of Ministry of Education, School of Materials Science and Engineering, Sun Yat-sen University, Guangzhou, 510275, China.

^cPharmaceutical college of Jiamusi University, Jiamusi University, Jiamusi, 154007, China.

*Correspondence to:

Kangshun Zhu, Laboratory of Interventional Radiology, Department of Minimally Invasive Interventional Radiology, and Department of Radiology, the Second Affiliated Hospital of Guangzhou Medical University, Guangzhou, 510260, China. E-mail: zhksh010@163.com.

**Correspondence to:

Xintao Shuai, PCFM Lab of Ministry of Education, School of Materials Science and Engineering, Sun Yat-Sen University, Guangzhou, 510275, China. E-mail: shuaixt@mail.sysu.edu.cn. Telephone: 86-20-84110365, Fax: 86-20-84112245.

[#]These authors contributed equally.



Fig. S1. Acid–base titration curve of PPAP_{0%} (A), PPAP25_% (B), PPAP_{50%} (C), PPAP_{75%} (D), PPAP_{100%} (E)and NaCl solution (F). (NaOH concentration: 0.2 N).

Composition	PPAPSD _{0%}	PPAPSD _{25%}	PPAPSD _{50%}	PPAPSD _{75%}	PPAPSD _{100%}
Concentration of	600	2500	2450	2750	3500
Sample (ppm)					
DOX (ppm)	6.72	98.00	101.43	157.45	82.95
Concentration of	1 20	62 50	95 55	180 75	164 50
SPIONs (ppm)	1.20	02.50	95.55	107.75	104.50
Drug Loading	1.12	3.92	4.14	5.72	2.37
SPIO Londing					
content (%)	0.20	2.50	3.90	6.90	4.70

Table S1. The laoding contents of DOX and SPIONs of each micelle.

 $ppm,\,\mu g/mL$

Formulations	IC ₅₀ (μg/mL)	
DOX·HCl	0.22	
PPAPSD _{75%}	0.76	
PPAPSD _{75%} (pH 5.0)	0.29	

Table S2. IC₅₀ values of DOX against HepG2 cells in different formations

IC₅₀, half-maximal inhibitory concentration.



Fig. S2. Particle size of PPAPSD_{75%} against time in PBS containing 10% FBS. Data were presented as mean \pm standard deviation (n = 3).



Fig. S3. Signal intensity (on T2WI images) and T2 value (on T2-map images) change of the tumors before and after the injection (means \pm SD; n = 5).



Fig. S4. H&E staining of heart tissue sections from the mice treated with different formulations. The DOX HCl group exhibited obvious myocyte loss and matrix disorganization in the cardiac tissues, whereas the PPAPS_{75%} and PPAPSD_{75%} groups displayed no significant cardiotoxicity (scale bar: 20 μm).