Supplementary Information for

pH-responsive nanoparticle based on Ibuprofen prodrug as drug carrier for inhibition of primary tumor growth and metastasis

Zhi Zeng,^a Zeliang Wei,^b Limei Ma,^a Yao Xu,^a Zhihua Xing,^a Hai Niu,^{bc} Haibo Wang,^{*d} and Wen Huang^{*ab}

^a Laboratory of Ethnopharmacology, Regenerative Medicine Research Center, West China Hospita, West China Medical School, Sichuan University, Chengdu 610041, China. *E-mail: huangwen@scu.edu.cn

^b Institute for Nanobiomedical Technology and Membrane Biology, Sichuan University, Chengdu 610041, China. *E-mail: niuhai@scu.edu.cn; Fax: +86-028-85164075; Tel: +86-028-85164073

^c College of Mathematics, Sichuan University, Chengdu, 610045, China

^d Textile Institute, College of Light Industry, Textile and Food Engineering, Sichuan University, Chengdu, 610065, China



Figure S1 ¹H-NMR spectra of HEI (A) and PEG-CBA (B) in CDCl₃.

Table S1 Characterization data of the polymers		
	$Mn(GPC)^a$	PDI
MPEG-PHEI5	3714	1.15
MPEG-PHEI ₁₀	6229	1.24

^{*a*} measured by GPC (THF, PS standards).



Figure S2 Size distribution of blank nanoparticles in physiological and acidic conditions (37 °C, pH 7.4 and 5.0).





Figure S4 Ibuprofen release profiles of MPEG-PHEI in physiological and acidic conditions (37 °C, pH 7.4 and 5.0).



Figure S5 *In vivo* antitumor effcacy of DOX/NPs compared to ibuprofen, as well as the combination of DOX and ibuprofen. (A) Body weights of mice, (B) tumor growth curves, (C) tumor morphologies after the mice were sacrificed. Error bars indicate SD (n = 6). *P < 0.01, treated group versus saline group. The scale bar is 500 µm.