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## **Supplementary Information**

## "Sheddable" PEG-Lipid to Balance the Contradiction of PEGylation between Long Circulation and Poor Uptake

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**Scheme S1** Hydrolysis route of DSPE-Hy-PEG copolymer at acidic condition.

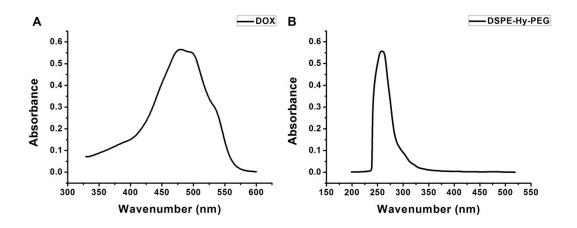


Fig. S1 (a) UV-vis absorption spectra of DOX; (b) UV-vis absorption spectra of DSPE-Hy-PEG.

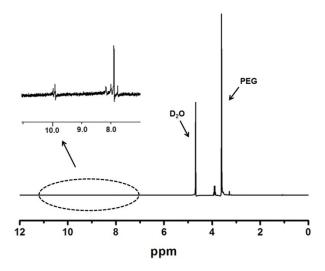


Fig. S2  $^1\mbox{HNMR}$  spectra of DSPE-Hy-PEG micelles incubated at pH 6.8 for 24 h.

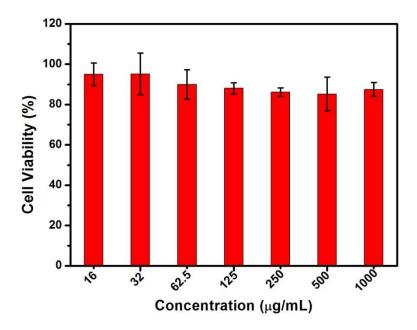


Fig. S3 Viability of MCF-7 cells after incubation for 24 h with empty DSPE-Hy-PEG micelles at various concentrations.

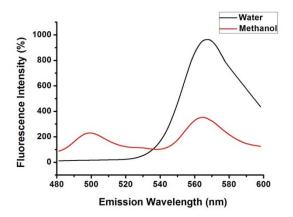


Fig. S4 Emission spectra of DiO/DiI-loaded DSPE-Hy-PEG micelles diluted by water or methanol.

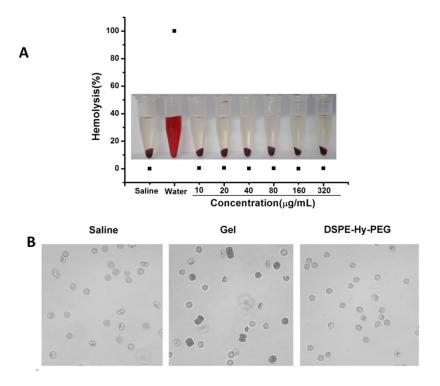
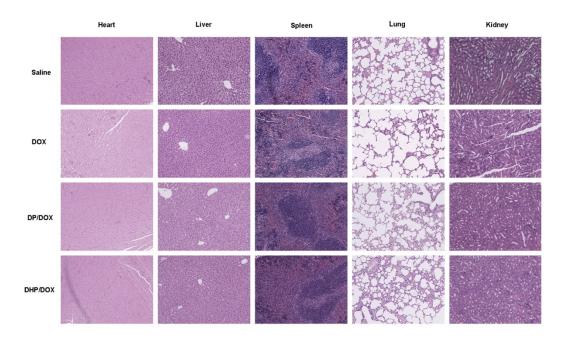


Fig. S5 (A) Hemolysis assay of empty micelles, using water as a positive control and saline as a negative control;
(B) CLSM images of Red blood cells incubated with empty micelles, using gel as a positive control and saline as a negative control.



**Fig. S6** Representative H&E sections of various organs of tumor-bearing nude mice after treatment with various drug formulations (DSPE-PEG represented by DP, DSPE-Hy-PEG represented by DHP).