Supporting Information for

## Tumor-specific activatable biopolymer nanoparticles stabilized by hydroxyethyl starch prodrug for self-amplified cooperative cancer therapy

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## **Supplementary Figures**



**Figure S1.** (A) Absorption spectra of P(Cu-DA) obtained in different ratios. (B) Absorption spectra of P(HSD-Cu-DA) obtained in different ratios. (C) DLS results for P(HSD-Cu-DA) NPs acquired with different ratios.



**Figure S2.** (A) XPS survey spectrum of P(HSD-Cu-DA) NPs. (B) Zeta potential of P(Cu-DA) and P(HSD-Cu-DA) NPs.



**Figure S3.** (A) Stability test of P(HSD-Cu-DA) NPs in different media (n=3). (B) Longterm stability of P(Cu-DA) and P(HSD-Cu-DA) NPs. (C) Digital photo of P(HSD-Cu-DA) and P(Cu-DA) NP solutions after storage in PBS for one month.



**Figure S4**. (A) Temperature rising curves for different concentrations of P(HSD-Cu-DA) NPs solution. (B) Temperature rise and fall curves for P(HSD-Cu-DA) and PDA NPs. (C) Linear relationship between -Ln ( $\theta$ ) and time (s) obtained from the cooling curves.



**Figure S5.** (A) UV-Vis spectra of P(Cu-DA) and P(HSD-Cu-DA) with same amount of DA and Cu<sup>2+</sup>. Temperature changes (B) and corresponding photothermal images (C) of P(Cu-DA) and P(HSD-Cu-DA) solution under NIR irradiation (808nm, 1W/cm<sup>2</sup>).



**Figure S6.** (A) Size distribution and TEM image of P(HSD-Cu-DA) NPs. (B) Size distribution and TEM image of P(HSD-Cu-DA) NPs treated with GSH (10mM) for 48h.



**Figure S7** FL emission spectra of P(HSD-Cu-DA) NPs under different conditions (excitation wavelength = 490nm).



**Figure S8**. In vitro release profiles of P(HSD-Cu-DA) NPs with/without NIR irradiation under pH5 condition (A) or in the presence of GSH (B).



**Figure S9**. (A) UV-Vis spectra of P(HSD-Cu-DA) NPs after incubation at different pH conditions for 48h. (B) Temperature changes and corresponding photothermal images of P(HSD-Cu-DA) solution upon NIR irradiation (808nm, 1W/cm<sup>2</sup>) after incubation at different pH conditions.



**Figure S10.** CLSM images (A) and semi-quantification (B) of 4T1 cells with ROS probe in different treatments.



Figure S11. Relative intracellular GSH content in 4T1 cells after different treatments.



**Figure S12.** Cytotoxicity of P(HSD-Cu-DA) NPs on Panc02 cells (A) and B16-F10 cells (B).



Figure S13. Cu contents in different organs and tumor of blank mice (n=3).



**Figure S14.** The plateau temperatures obtained by NIR irradiation (1W/cm<sup>2</sup>) after different times of intravenous administration.



Figure S15. The tumor inhibition rate (TIR) of different groups after 2-week therapy.



Figure S16. Masson staining images with or without NIR exposure.



**Figure S17.** UV-Vis spectra of P(HSD-Cu-DA) NPs after incubation with different concentrations of  $H_2O_2$  for 24 h.