Electronic Supplementary Material (ESI) for Journal of Materials Chemistry B. This journal is © The Royal Society of Chemistry 2019

## **Supplementary Information**

for

Dual-targeting Selenium Nanoparticles Antagonize Hyperinsulinemia-Promoted Tumor Growth via Activating Cell Autophagy

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**Figure S1.** Promotion of HepG2 cells migration by different concentrations (0.0005 mg  $L^{-1}$ , 0.05 mg  $L^{-1}$ , 5 mg  $L^{-1}$ ) of insulin for 36 h.



**Figure S2.** Promotion of HepG2 cells invasion different concentrations (0.0005 mg  $L^{-1}$ , 0.05 mg  $L^{-1}$ , 5 mg  $L^{-1}$ ) of insulin for 36h.



**Figure S3. A)** Promotion of HepG2 cells proliferation by insulin (0.0005 mg L<sup>-1</sup>, 0.05 mg L<sup>-1</sup>, 5 mg L<sup>-1</sup>); **B)** HepG2 cells drug resistance on DOX by insulin (0.0005 mg L<sup>-1</sup>, 0.05 mg L<sup>-1</sup>, 5 mg L<sup>-1</sup>).



**Figure S4.** HepG2 cells viability incubated by EBSS (2 mL), Insulin (0.0005 mg  $L^{-1}$ , 0.05 mg  $L^{-1}$ , 5 mg  $L^{-1}$ ) and 3-MA (5 mM).



**Figure S5.** Stability of different nanoparticles. Size changes (A) and zeta potential (B) changes in different solution for 12 h.



Figure S6. Inhibition of HepG2 cells migration by different nanoparticles (0.5  $\mu$ M) for 36 h.



Figure S7. Inhibition of HepG2 cells invasion by different nanoparticles (0.5  $\mu$ M) for 36 h.



**Figure S8.** Cell cycle of HepG2 cells treated by different concentrations of CS-SeNPs, uPA-SeNPs and ACPP-SeNPs.



Figure S9. Quantitative analysis of cell cycle.



Figure S10. Representative photos of ROS produced by CS-SeNPs and u/A-SeNPs.



Figure S11. Images of HepG2 cells after MDC staining with 10  $\mu$ M u/A-SeNPs and other solution.



Figure S12. Blue spots in HepG2 after MDC staining and 10  $\mu$ M u/A-SeNPs treatment.