

## Supporting Information

### **Polarity Control of DNA Adsorption Enabling Surface Functionalization of CuO**

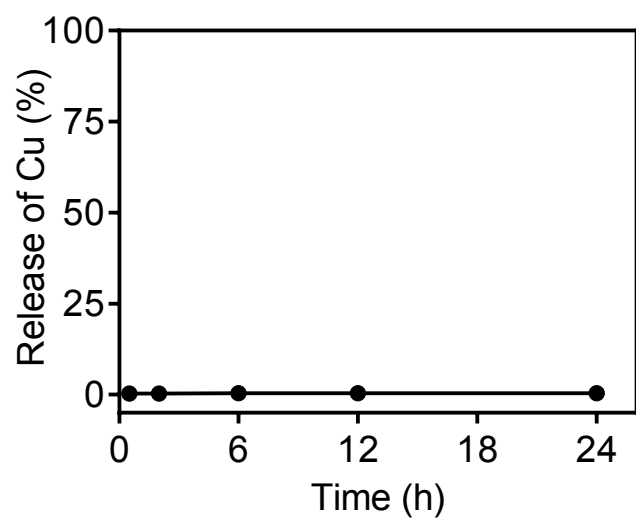
#### **Nanozyme for Targeted Tumor Therapy**

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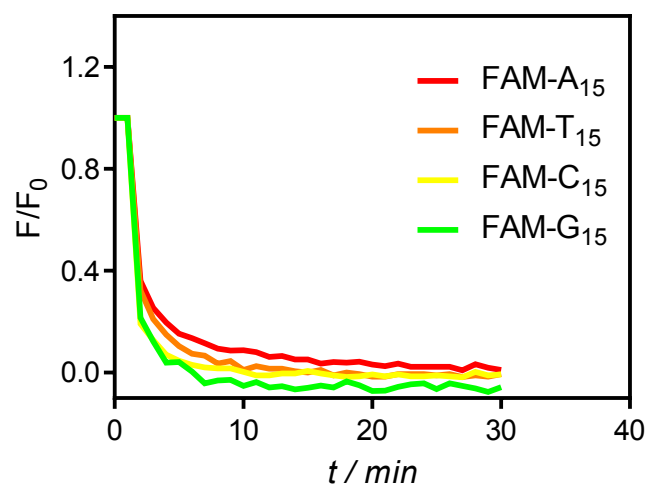
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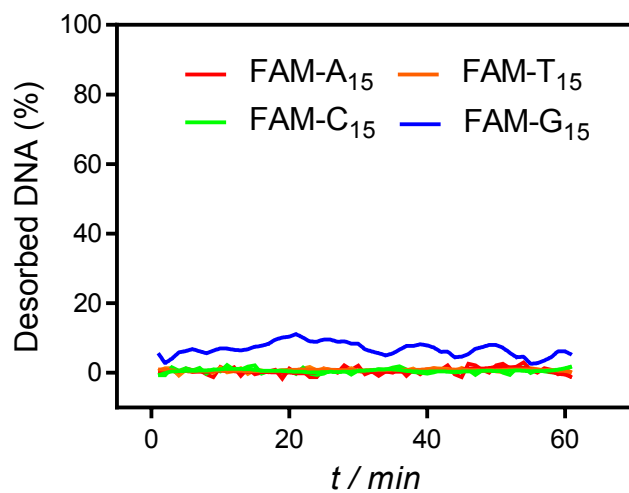
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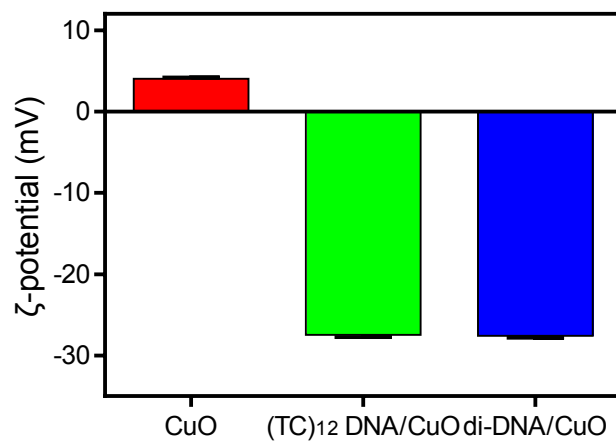
**Figure S1.** The release kinetics of Cu ions from CuO NPs.



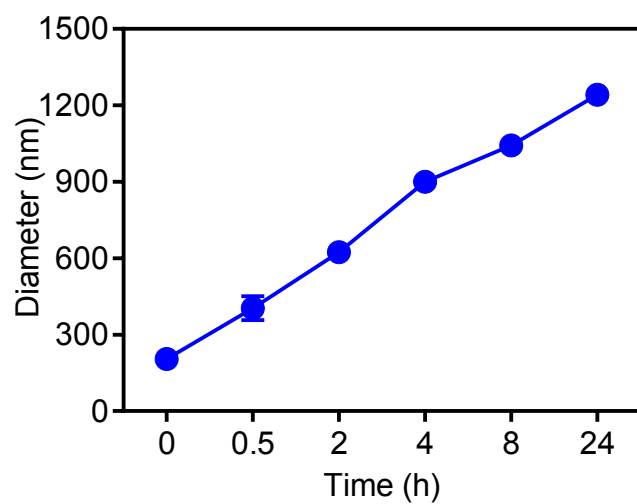
**Figure S2.** The adsorption of FAM-labeled 15-mer DNA homo-polymers (20 nM) by 50 µg/mL CuO NPs in HEPES buffer (10 mM, pH 7.6, 150 mM NaCl).



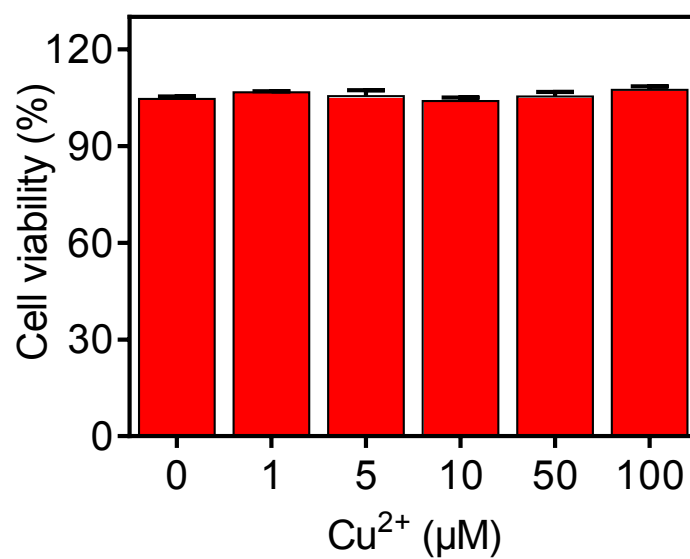
**Figure S3.** DNA desorption kinetics from CuO NPs surface in presence of 500 mM NaCl.



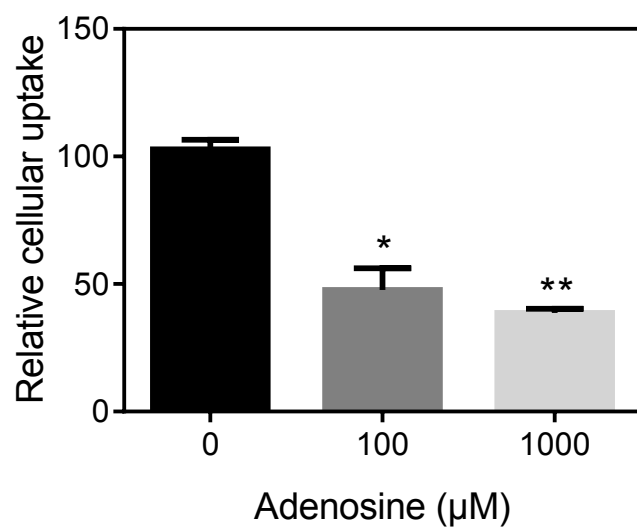
**Figure S4.** The  $\zeta$  potential of CuO NPs, (TC)<sub>12</sub> DNA/CuO and di-DNA/CuO.



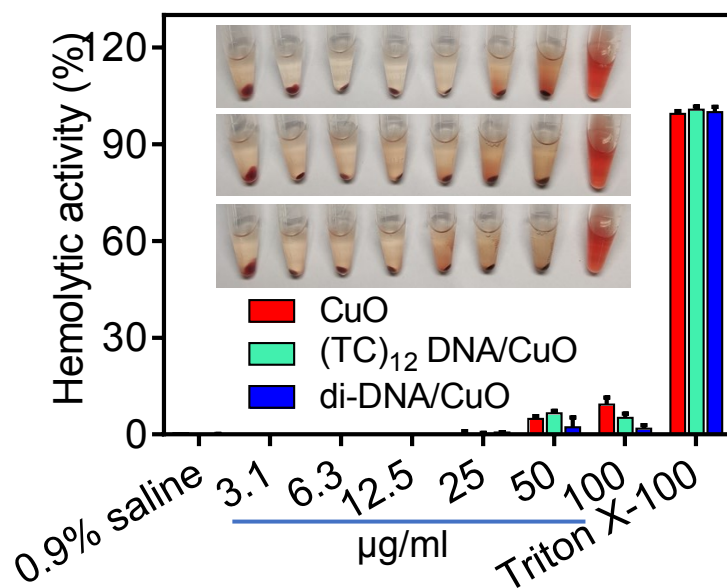
**Figure S5.** Colloidal stability of bare CuO NPs in water.



**Figure S6.** Cell viability of MDA-MB-231 cells after incubation with different  $\text{Cu}^{2+}$  concentrations for 24 h.

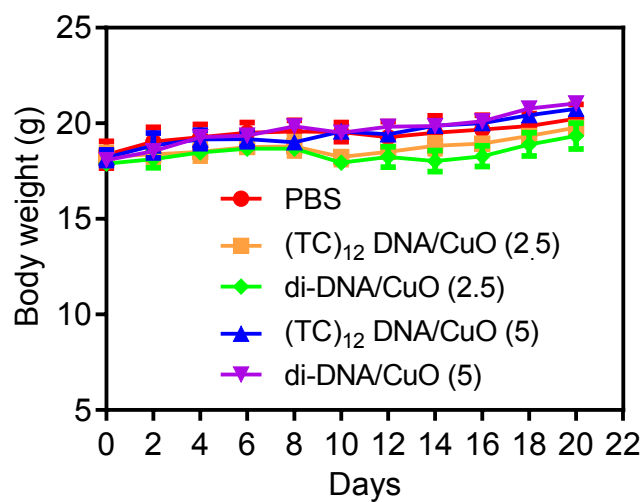


**Figure S7.** The cellular uptake of di-DNA/CuO by MDA-MB-231 cells without or with pretreatment of different concentrations of free adenosine. \* $P < 0.05$ , \*\* $P < 0.01$ .

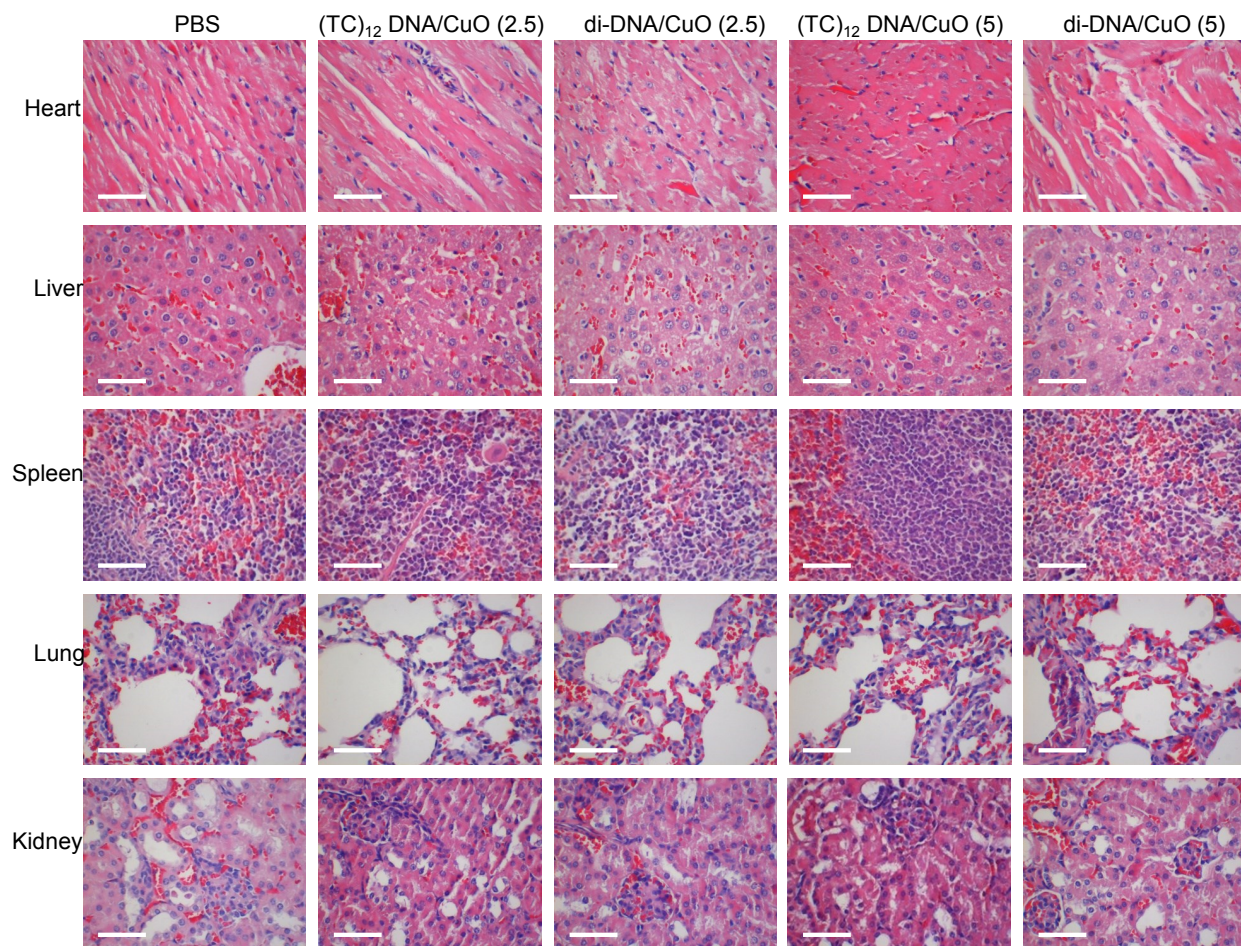


**Figure S8.** Hemolysis test of CuO, (TC)<sub>12</sub> DNA/CuO and di-DNA/CuO at different concentrations. Negative control and positive control were 0.9% saline and Triton X-100,

respectively. All these formulations showed minimal hemolysis as compared to the positive control, demonstrating the biosafety of the nano-systems for intravenous injection.



**Figure S9.** Dynamic monitoring the body weight for 20 days with different treatments.



**Figure S10.** Representative H&E staining images of main organs (heart, liver, spleen, lung and kidney) after the treatments, scale bar = 50  $\mu\text{m}$ .