

Supplementary Material

Table S1. List of miRNAs used in this study.

Name	Sequences (5' to 3')	
miRNA-204 mimic (5' FAM-tag)	Sense	5'-UUC CCU UUG UCA UCC UAU GCC U-3'
	Antisense	5'-A GGC AUA GGA UGA CAA AGG GAA-3'
random miRNA (Negative Control)	Sense	5'-UUC UCC GAA CGU GUC ACG UTT-3'
	Antisense	5'-ACG UGA CAC GUU CGG AGA ATT-3'

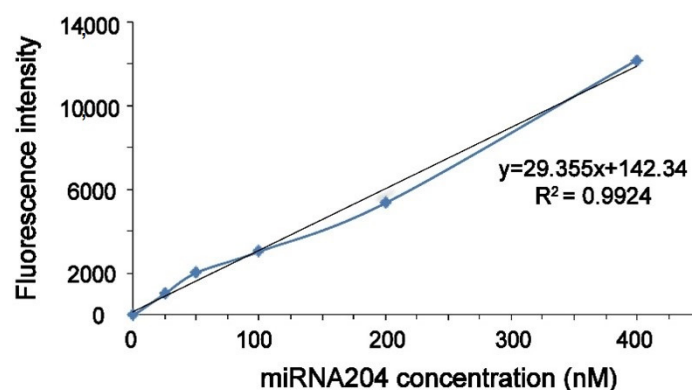


Figure S1. Standard curve of miR-204 concentration versus the corresponding fluorescence intensity.

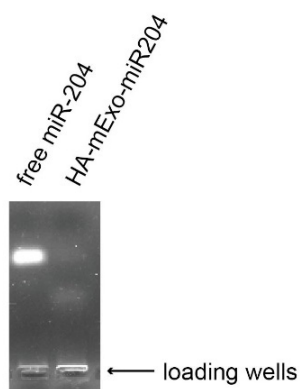


Figure S2. Agarose gel electrophoresis of free miRNA-204 and HA-mExo-miR204.

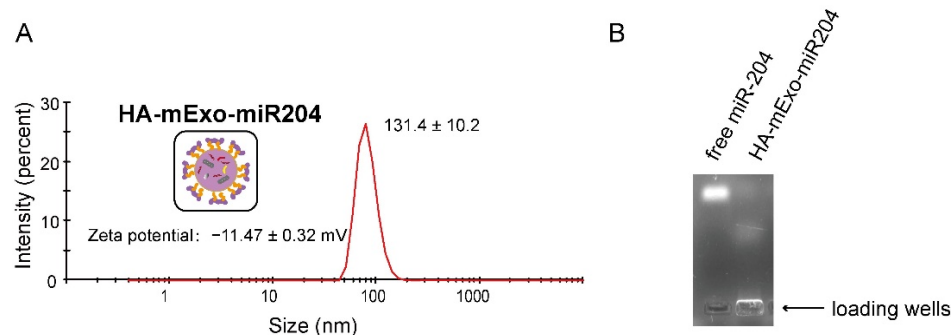


Figure S3. Storage stability of HA-mExo-miR204. DLS analysis (A) and agarose gel electrophoresis (B) of HA-mExo-miR204 after storage at -20°C for 7 days.

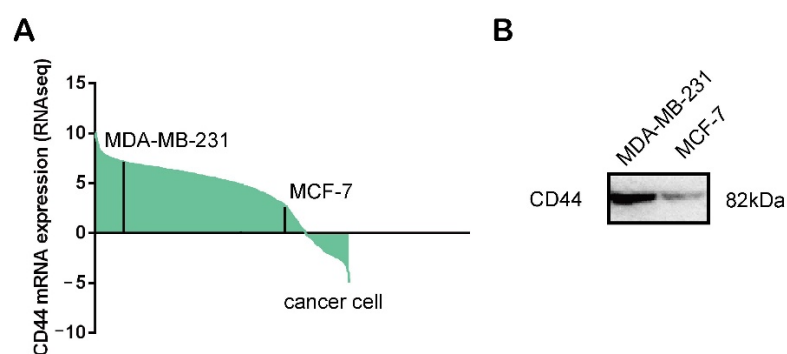


Figure S4. CD44 expression in cancer cells. (A) Relative expression of CD44 mRNA in MDA-MB-231 and MCF-7 cancer cell lines. (B) Western blot analysis of CD44 protein expression in MDA-MB-231 and MCF-7 cancer cell lines.

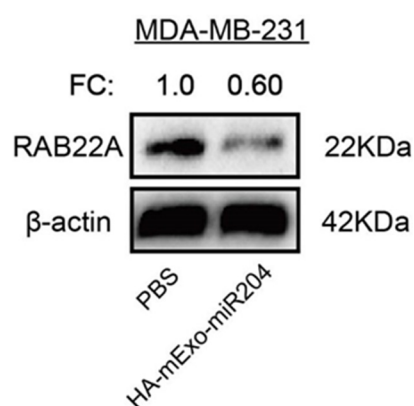


Figure S5. Western blotting analysis of RAB22A expression. MDA-MB-231 cells were treated with PBS or HA-mExo-miR204 for 12 h.

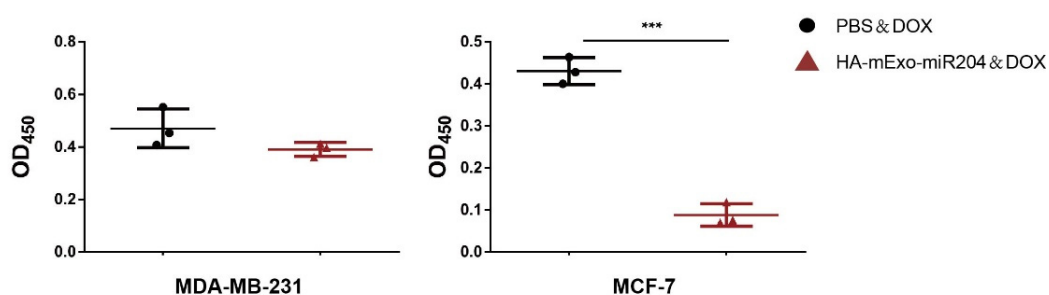


Figure S6. The effect of HA-mExo-miR204 on the chemosensitivity of Dox. Cancer cells were treated with PBS or HA-mExo-miR204 in combination with Dox for 12 h and the effect of HA-mExo-miR204 on Dox chemosensitivity was determined by CCK-8 cell viability assay. *** $p < 0.001$.

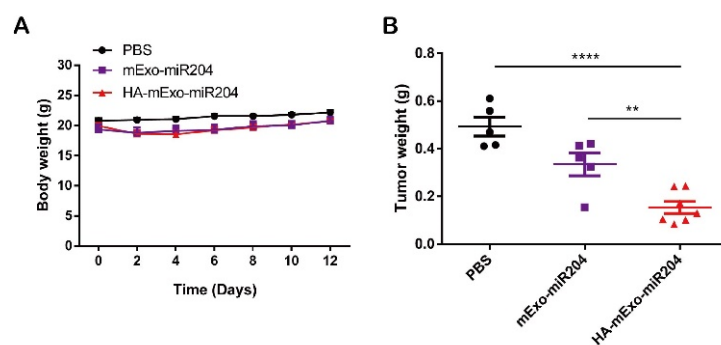


Figure S7. Body weight and tumor weight of mice. (A) Body-weight of tumor-bearing mice during the therapeutic course. (B) Endpoint tumor weight the mice after 12-days treatments. ** $p < 0.01$, **** $p < 0.0001$.

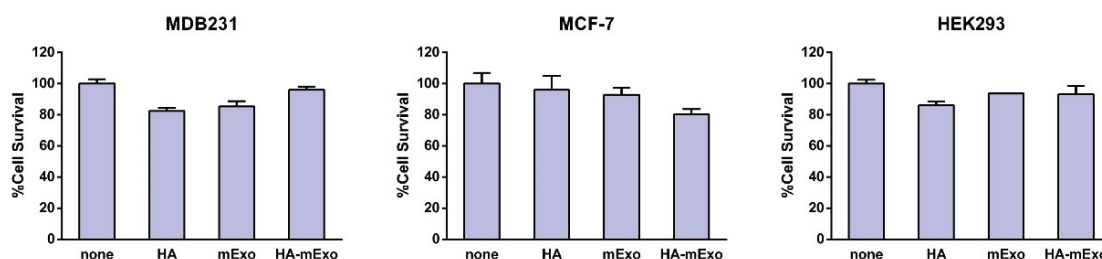


Figure S8. Cytotoxicity of DSPE-PEG₂₀₀₀-HA, mExos and HA-mExos. MDA-MB-231, MCF-7 and HEK293 cells were incubated with DSPE-PEG₂₀₀₀-HA, mExos or HA-mExos, respectively. After 48 h, cell viability was determined by CCK-8 assay.

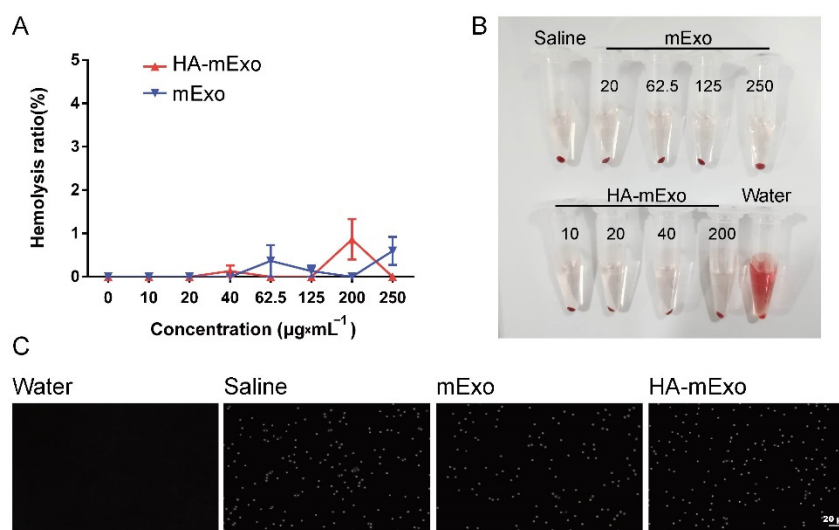


Figure S9. In vitro hemocompatibility of mExos and HA-mExos. (A) Hemolysis ratio of mExos and HA-mExos. (B) The images of red blood cells after incubation with mExos and HA-mExos. (C) Microscopy analysis of red blood cell membrane integrity after incubation with mExos and HA-mExos. Scale bars: 20 µm.

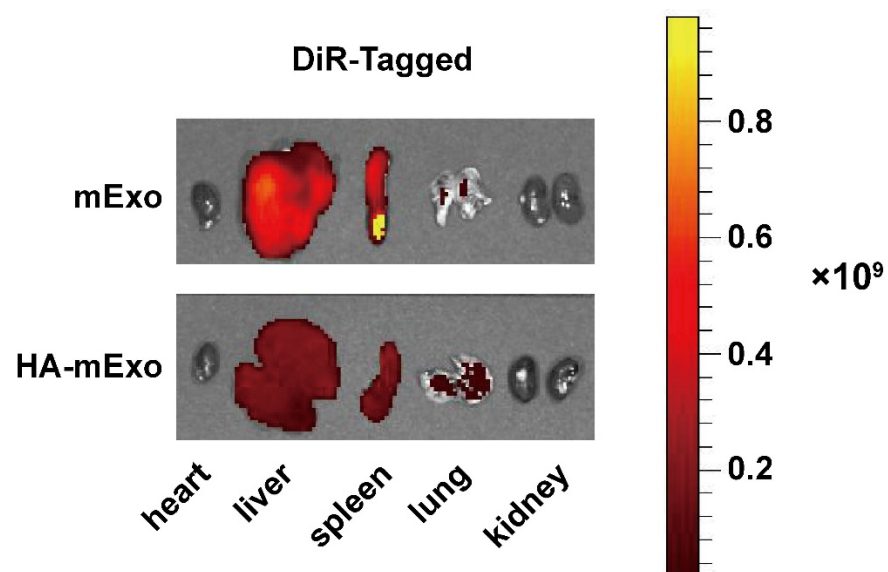


Figure S10. In vivo distribution of mExos and HA-mExos. The visualization of mExos and HA-mExos was achieved by labeling the exosomes with DiR dyes. Images of mice organs were recorded using an in vivo Fluorescence Image System (IVIS SPECTRUM, PerkinElmer, Waltham, MA, USA).