

*Supplementary Information*

**Lung-targeted feedback regulation of the mitochondrial ATP  
synthetic pathway for orthotopic tumors suppression**

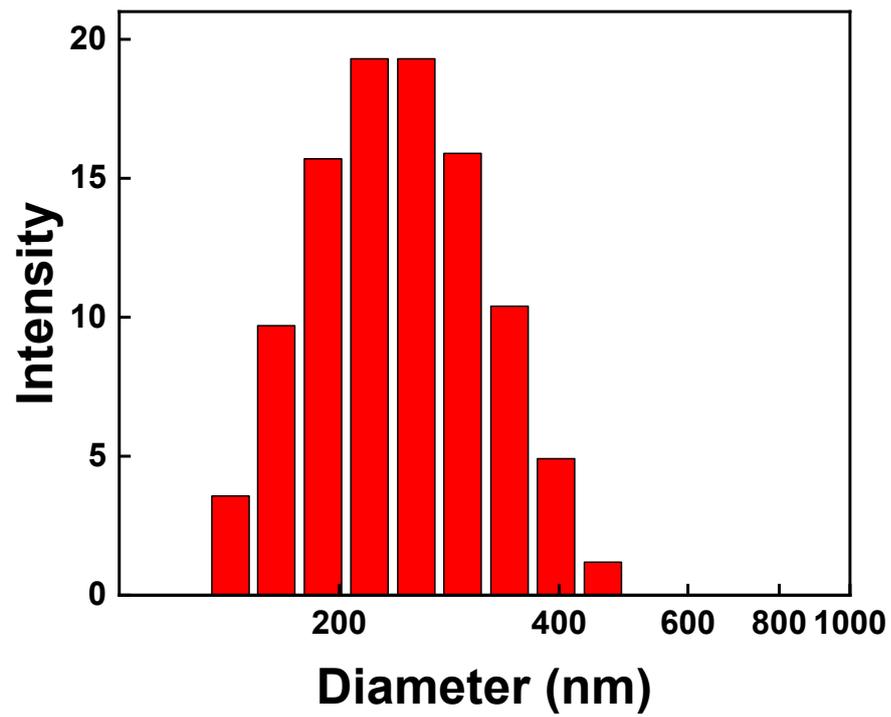
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**Fig. S1** Hydrodynamic diameter of ZIF-90/AIPH/BE.

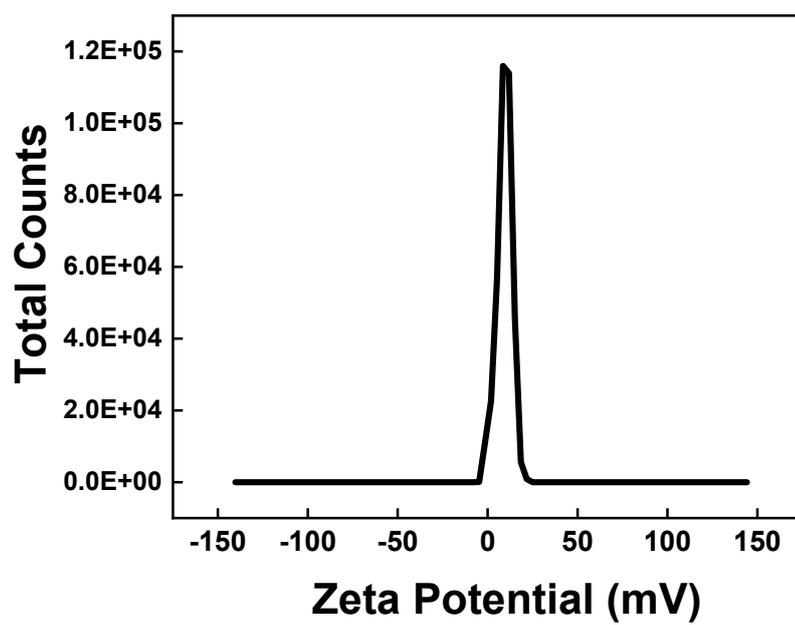
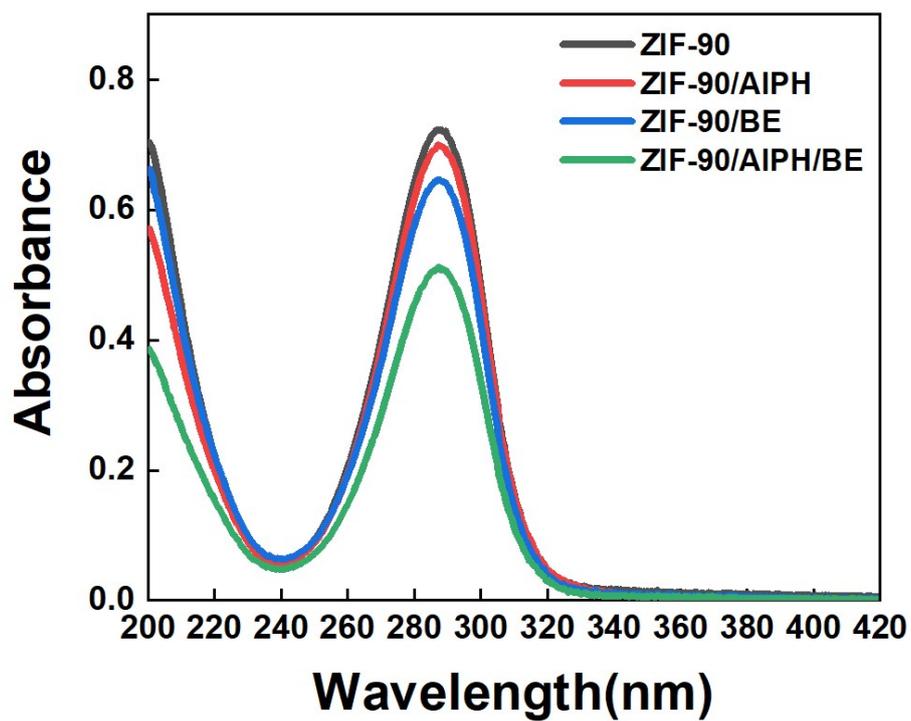
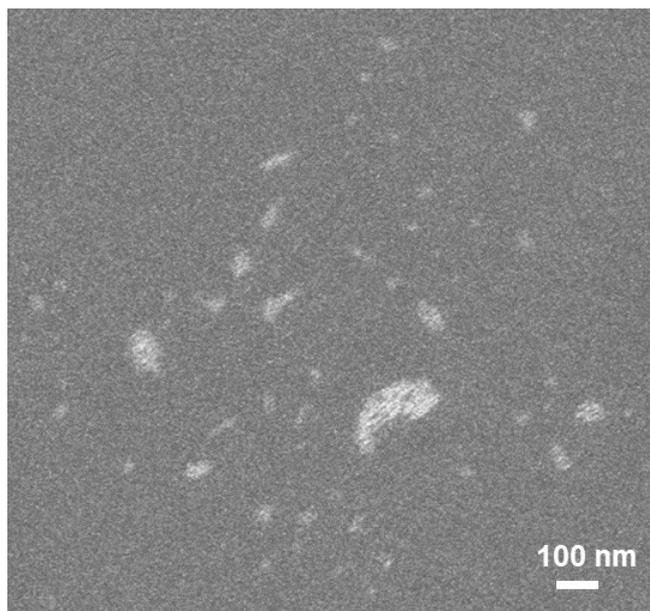


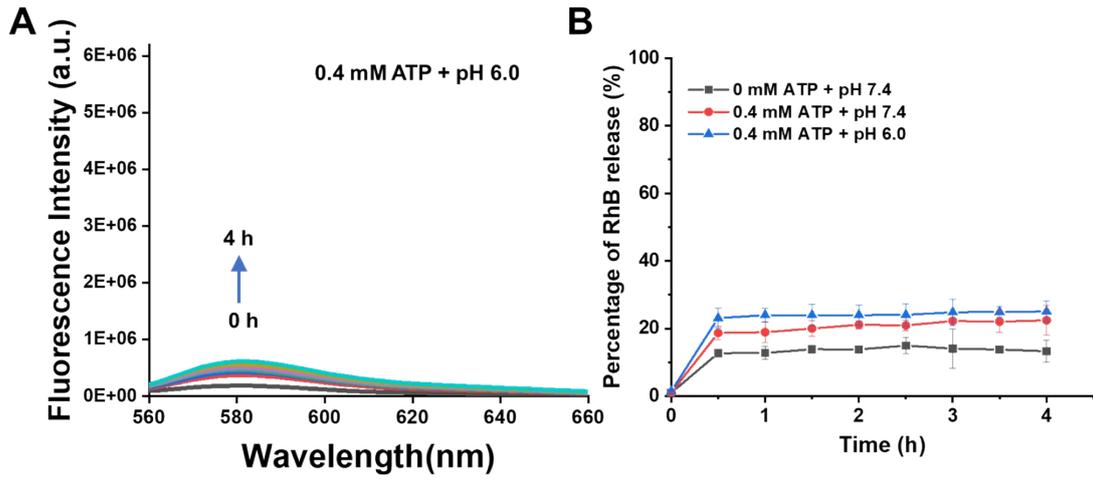
Fig. S2 Zeta potential diagram of ZIF-90/AIPH/BE.



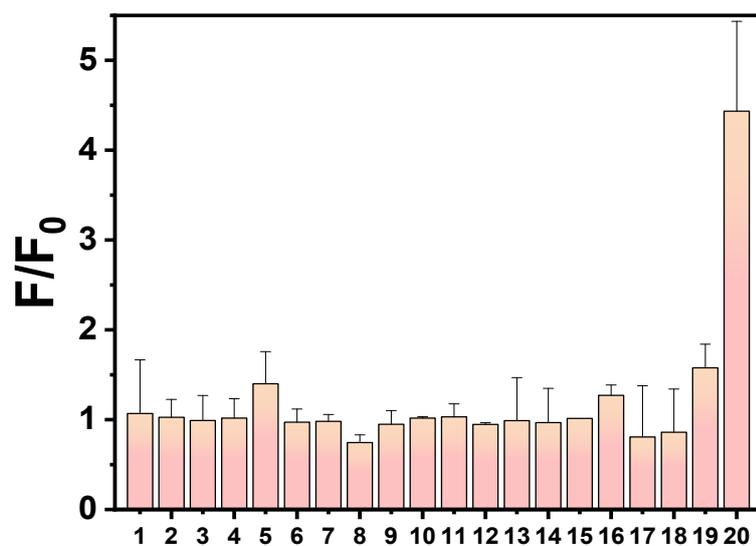
**Fig. S3** UV spectra of ZIF-90, ZIF-90/AIPH, ZIF-90/BE, ZIF-90/AIPH/BE.



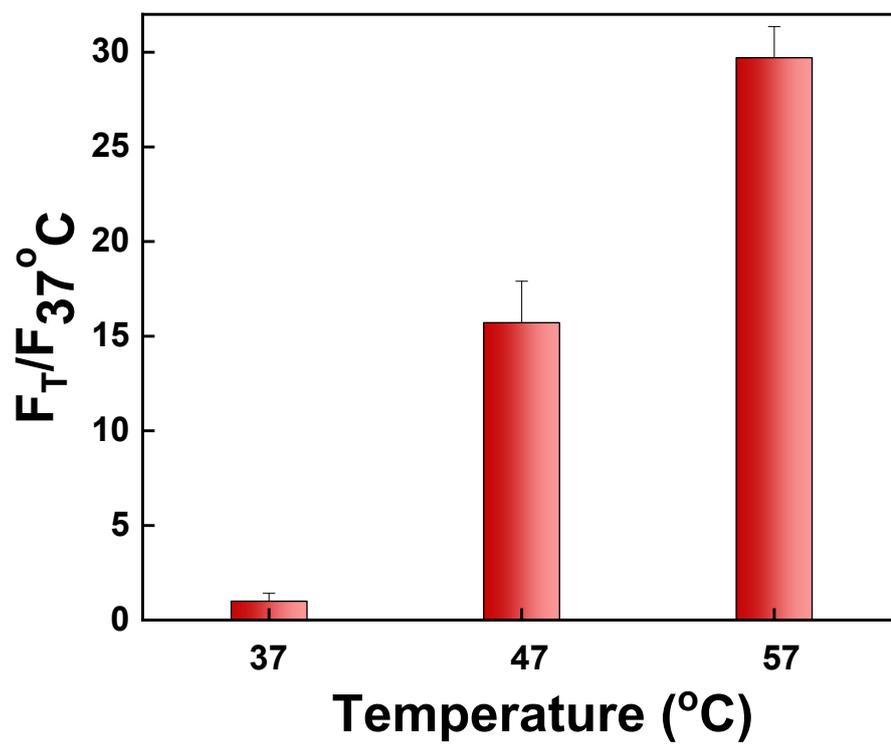
**Fig. S4** SEM of ZIF-90/AIPH/BE after incubation with 2 mM ATP.



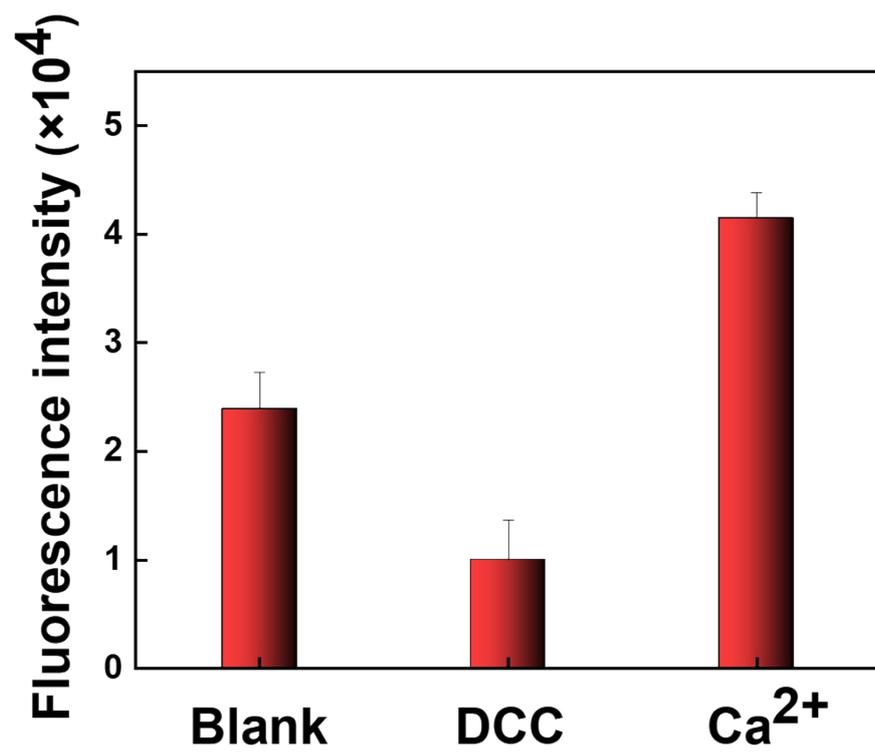
**Fig. S5** Release profiles of RhB when subjected to the mimic extratumoral conditions: (A) the fluorescence spectra; (B) the amount of released RhB.



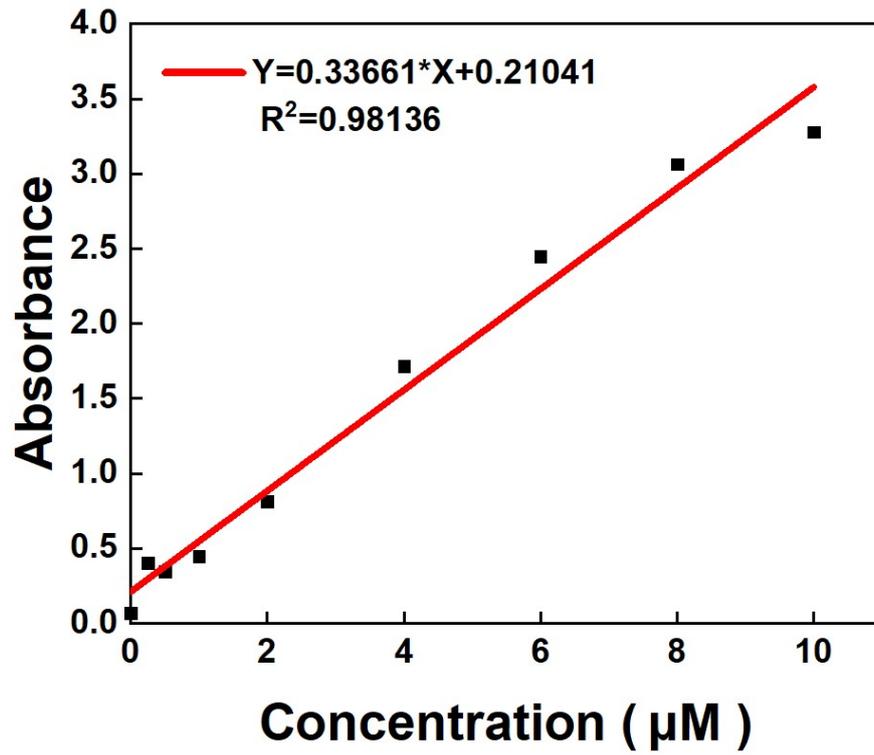
**Fig. S6** The selectivity of ZIF-90/RhB nanoprobe for ATP. (1:  $K^+$ ; 2:  $Na^+$ ; 3:  $Ca^{2+}$ ; 4:  $Mg^{2+}$ ; 5:  $Cu^{2+}$ ; 6:  $Ba^{2+}$ ; 7:  $NH_4^+$ ; 8:  $NO_3^-$ ; 9:  $PO_3^{3-}$ ; 10:  $SO_4^{2-}$ ; 11:  $SO_3^{2-}$ ; 12:  $CO_3^{2-}$ ; 13:  $HCO_3^-$ ; 14:  $Br^-$ ; 15:  $Cl^-$ ; 16: Gly; 17: Glu; 18: GMP; 19: AMP; 20: ATP).



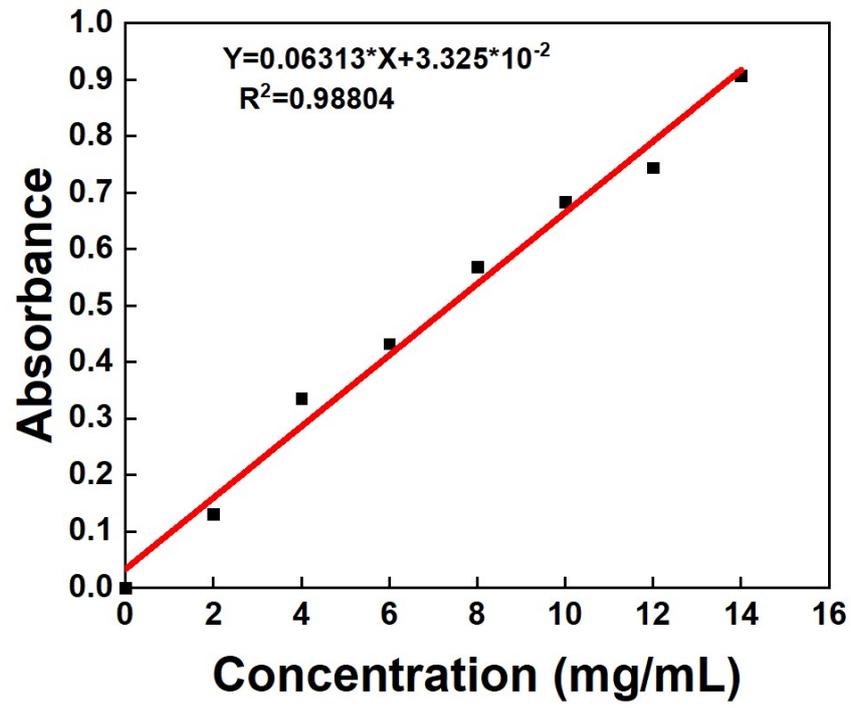
**Fig. S7** Fluorescence ratio of DCF through a reaction between DCFH-DA and alkyl radicals from AIPH at different temperature.



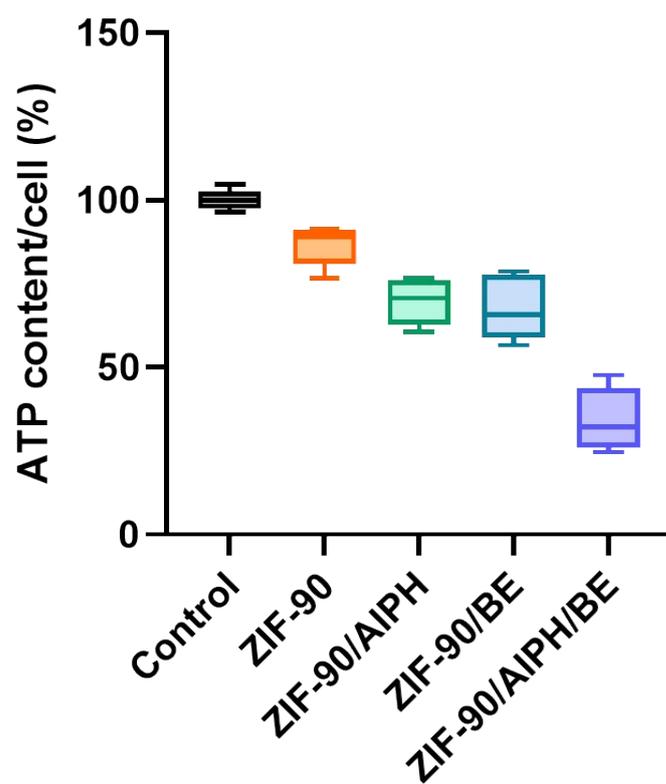
**Fig. S8** Fluorescence intensity of ZIF-90/RhB in A549 cells upon treatment with DCC and Ca<sup>2+</sup>.



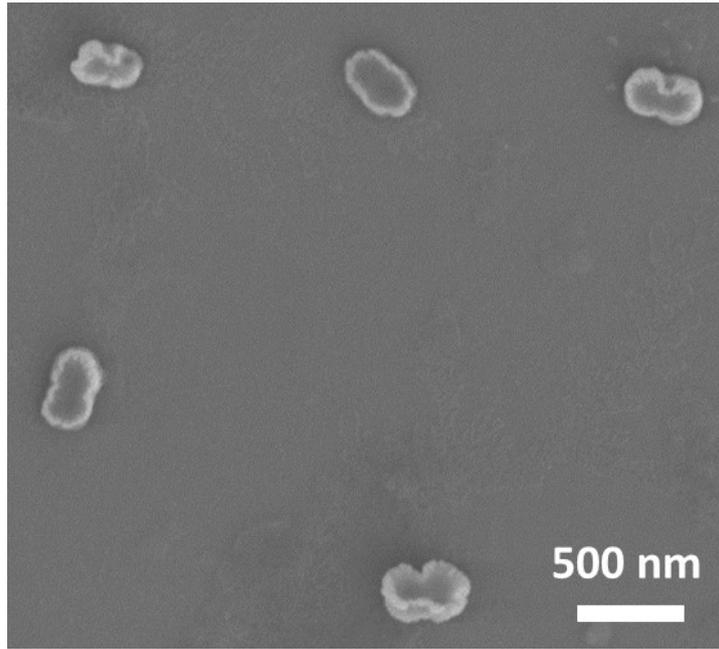
**Fig. S9** Standard curve of NADH concentrations detected by Amplitude Colorimetric Total NAD and NADH Assay Kit.



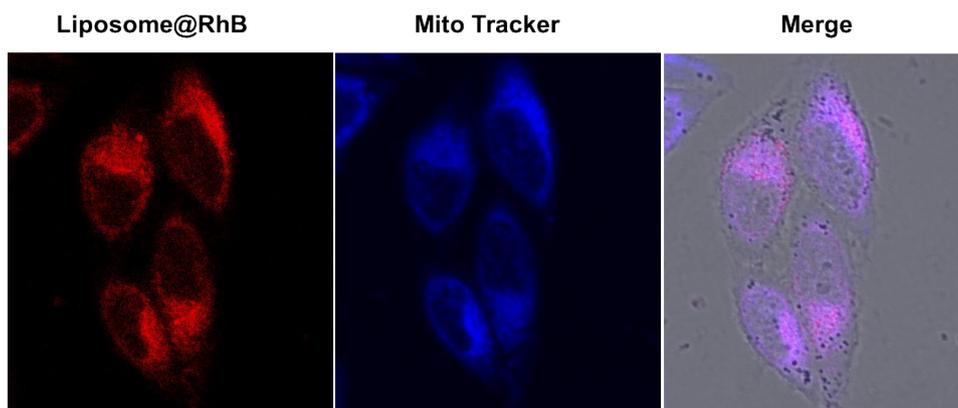
**Fig. S10** Standard curve of ATP contents in A549 cells detected by ATP assay Kit.



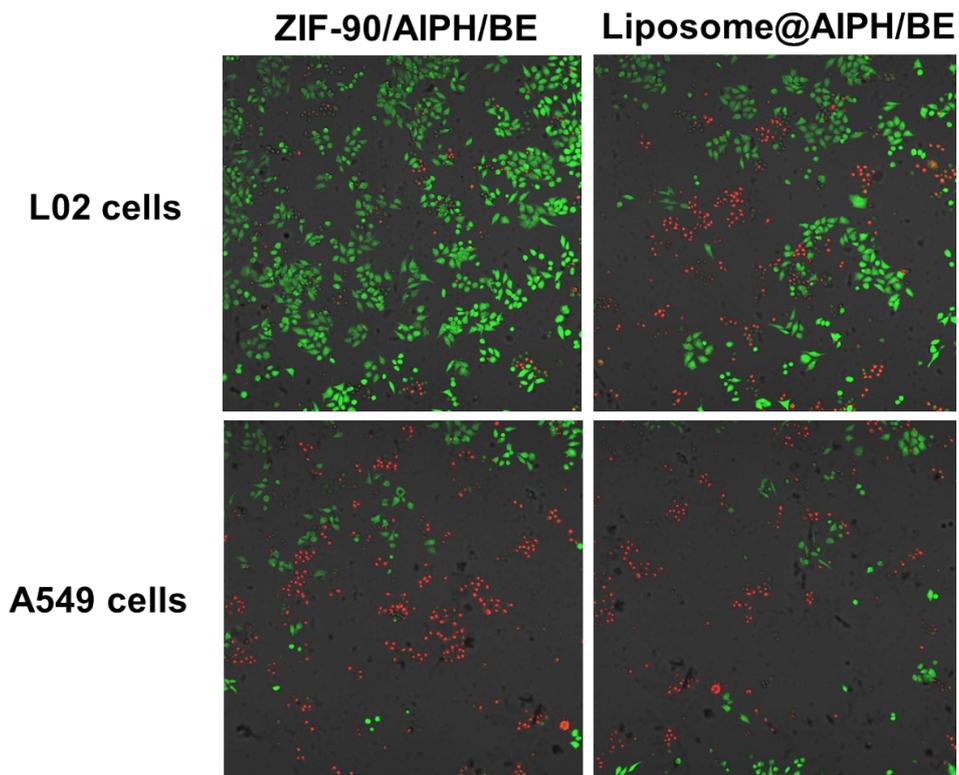
**Fig. S11** ATP content in per cell after A549 cells were treated with different formulations for 24 h. The values of the control group were all set as 100%.



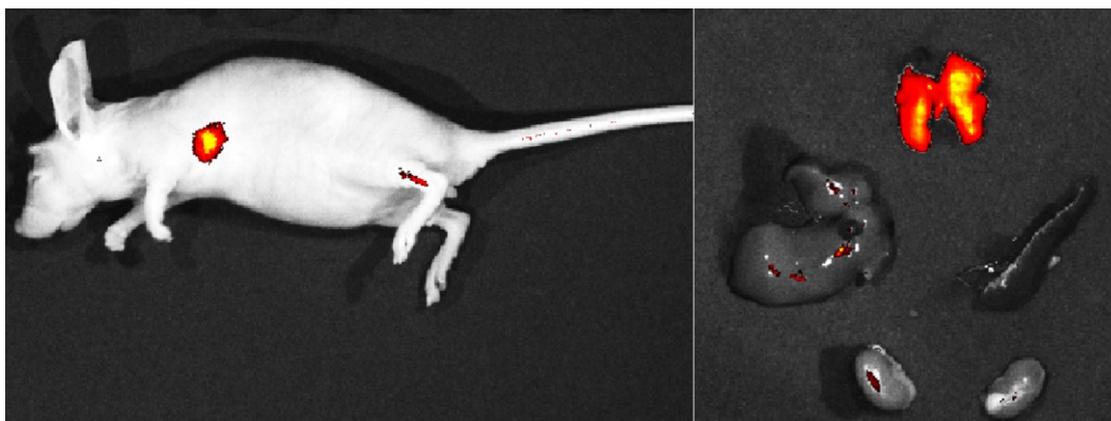
**Fig. S12** SEM of mitochondrion-orientated liposome.



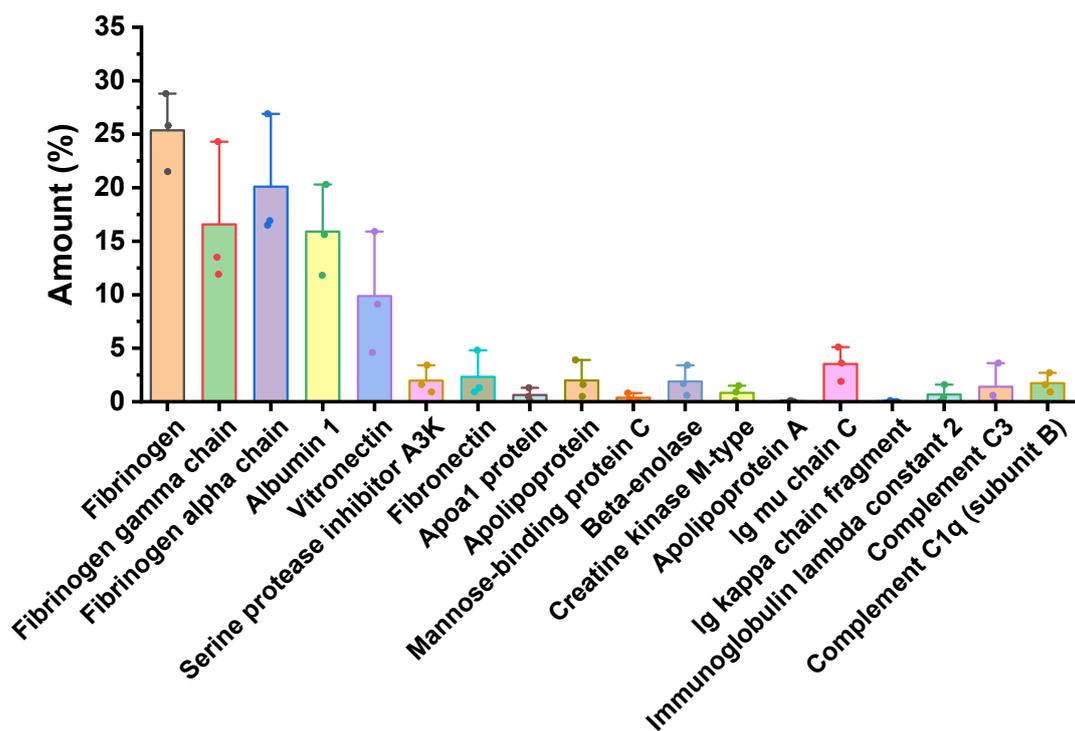
**Fig. S13** Co-localization experiments involving mitochondrion-orientated liposome@RhB, MitoTracker in A549 cells.



**Fig. S14** AM/PI staining confocal microscopy images of A549 or LO2 cells with different treatments.



**Fig. S15** *In vivo and ex vivo* fluorescence imaging after ZIF-90/Cy5.5 NPs was induced in normal Balb/c mice by intravenous injection for 2 h.



**Fig. S16** The top 18 most abundant proteins in the protein corona of different nanoparticles determined. Values were calculated from the molar masses of each protein identified by LC-MS.