

## Supporting Information

### **Tumor cell-activated “Sustainable ROS Generator” with homogeneous intratumoral distribution property for improved anti-tumor therapy**

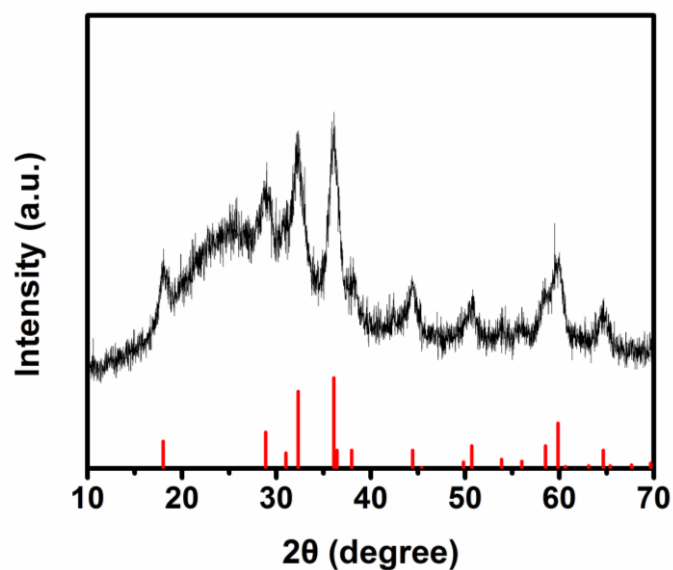
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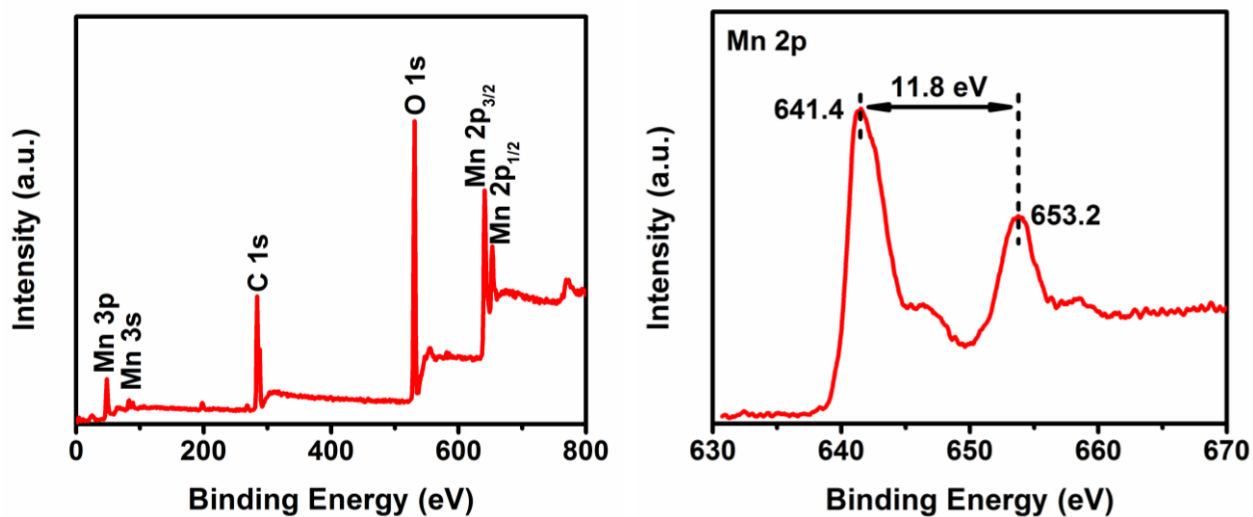
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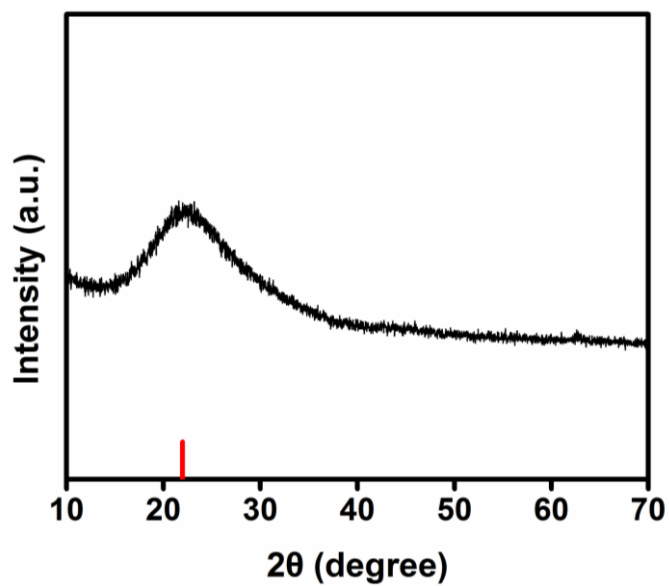
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shijinyxy@zzu.edu.cn



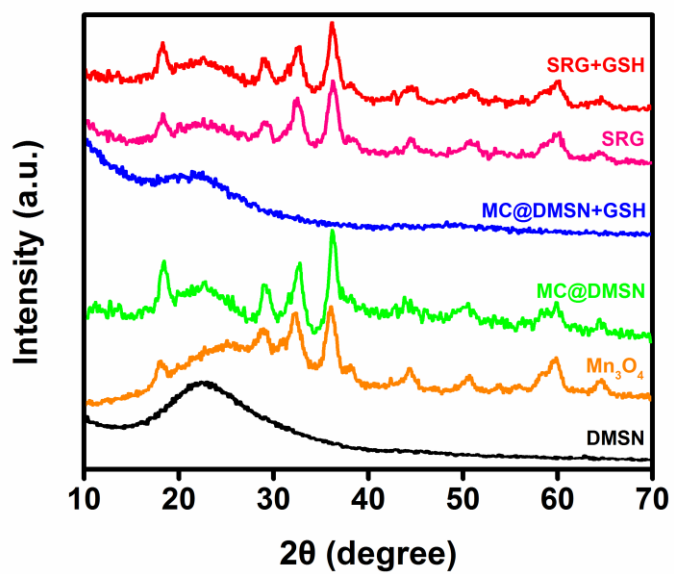
**Figure S1.** XRD pattern of obtained Mn<sub>3</sub>O<sub>4</sub> nanoparticles (the red lines at the bottom mark the reference pattern of hausmannite Mn<sub>3</sub>O<sub>4</sub> from the JCPDS database, card no.24-0734).



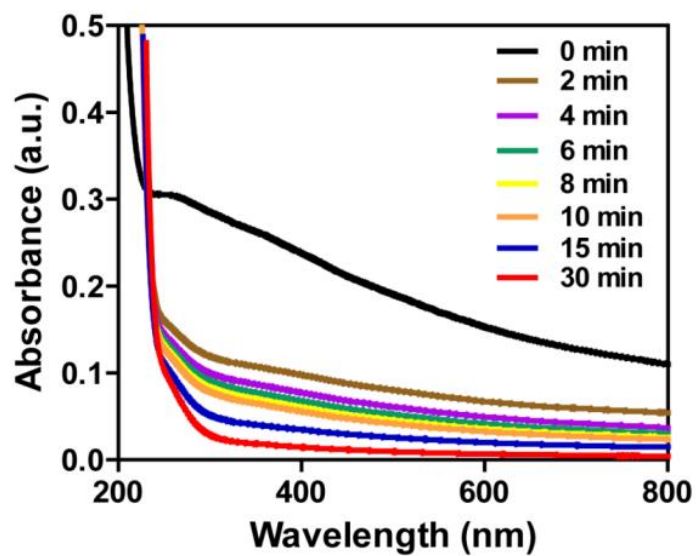
**Figure S2.** XPS spectrum of Mn<sub>3</sub>O<sub>4</sub> and XPS spectrum of Mn 2p.



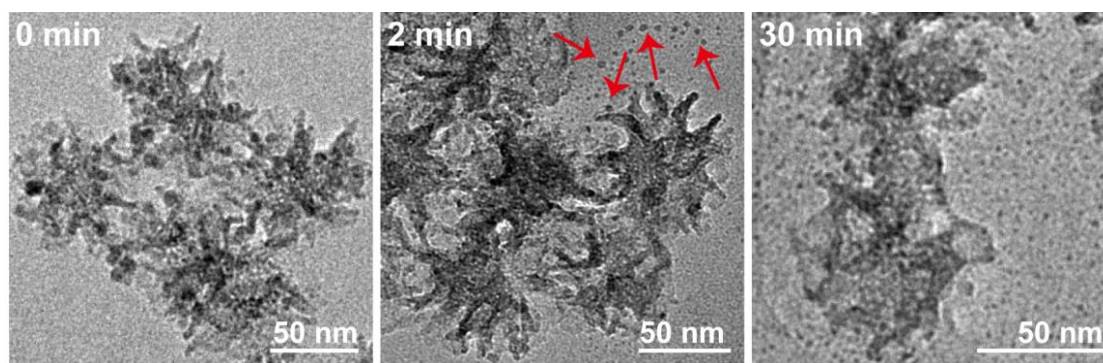
**Figure S3.** XRD pattern of obtained DMSN nanoparticles (the red lines at the bottom mark the reference pattern of silicon oxide from the JCPDS database, card no. 29-0085).



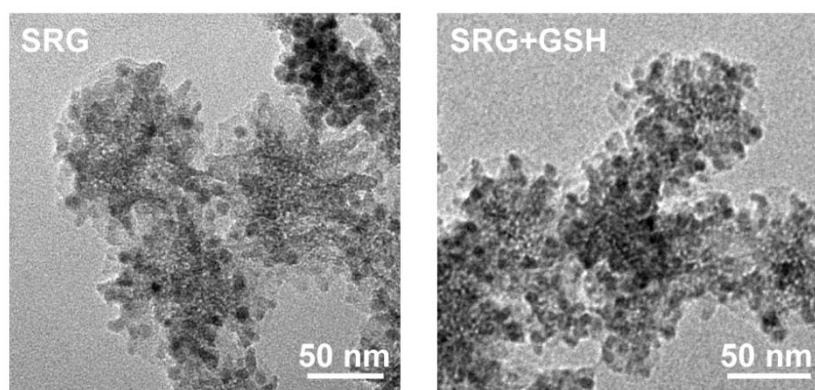
**Figure S4.** XRD pattern of different preparations.



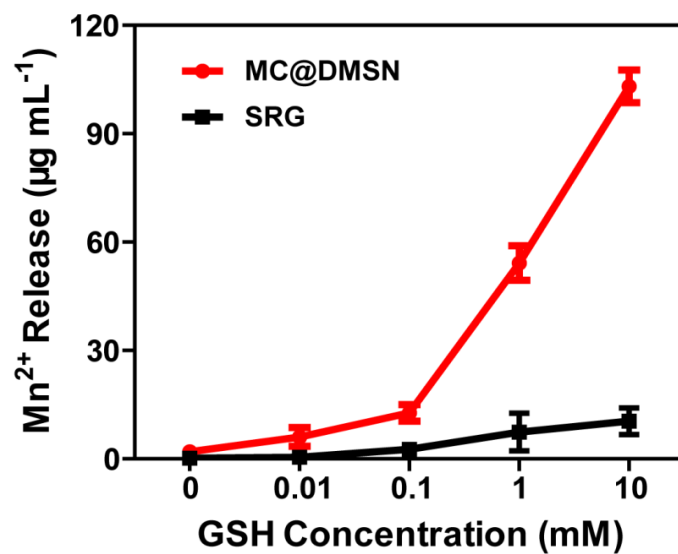
**Figure S5.** UV-vis absorption spectra of  $\text{Mn}_3\text{O}_4$  treated with GSH (1 mM) for different time.



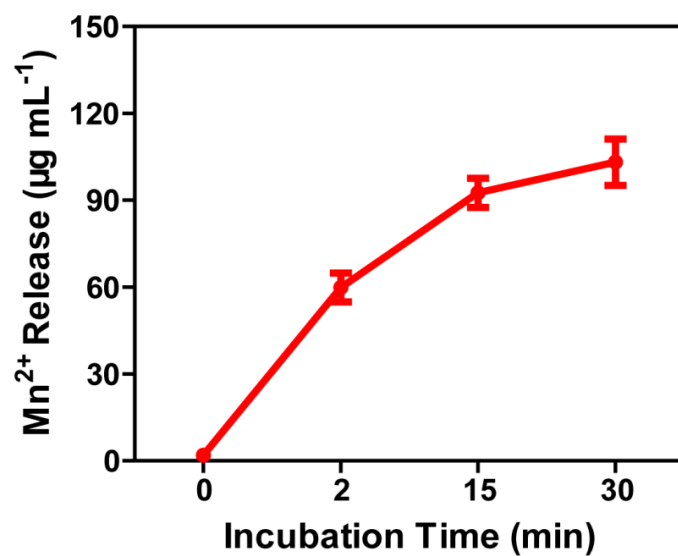
**Figure S6.** TEM images of MC@DMSN treated with 10 mM GSH for 0, 2 and 30 min, the red arrows represent part of un-degraded  $\text{Mn}_3\text{O}_4$  NPs.



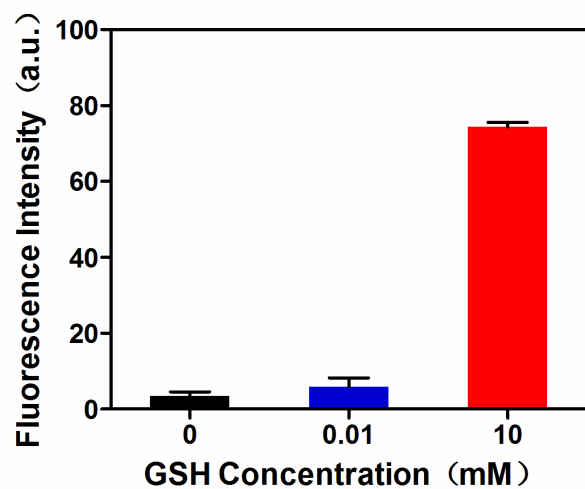
**Figure S7.** TEM images of SRG and SRG treated with 10 mM GSH for 30 min.



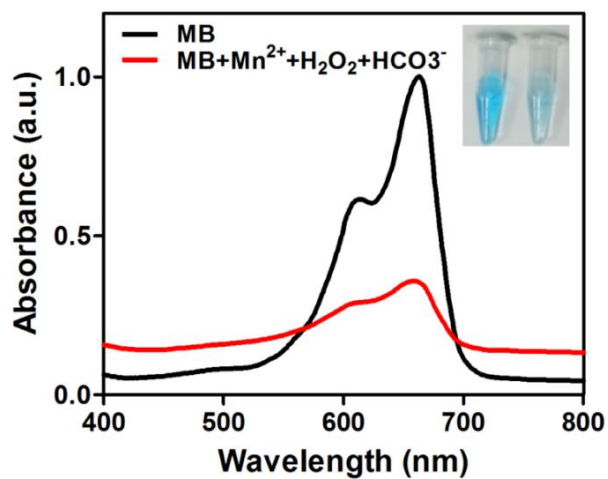
**Figure S8.** GSH-dependent release profiles of Mn<sup>2+</sup> from MC@DMSN and SRG as measured by ICP-MS.



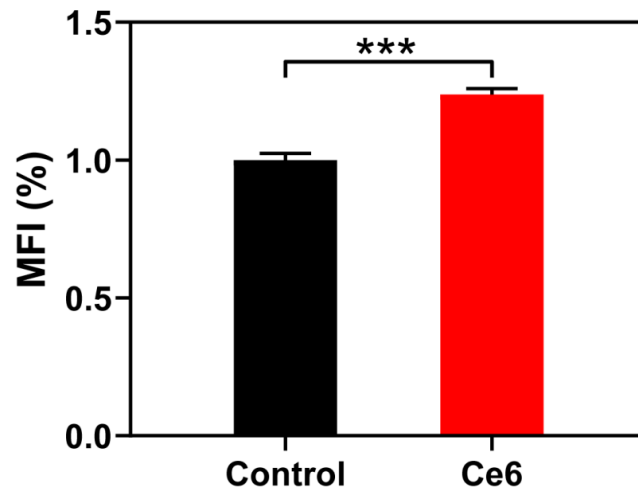
**Figure S9.** Time-dependent release profiles of Mn<sup>2+</sup> from MC@DMSN as measured by ICP-MS.



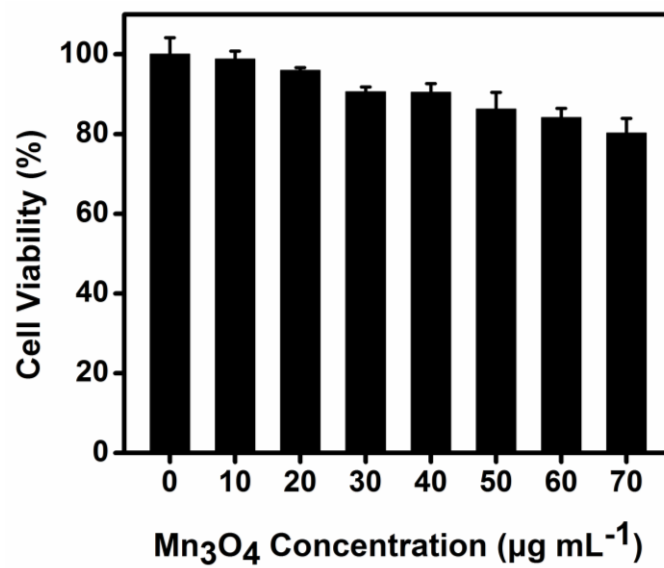
**Figure S10.** Response of Ce6 fluorescence intensity of MC to different concentrations of GSH (n=3).



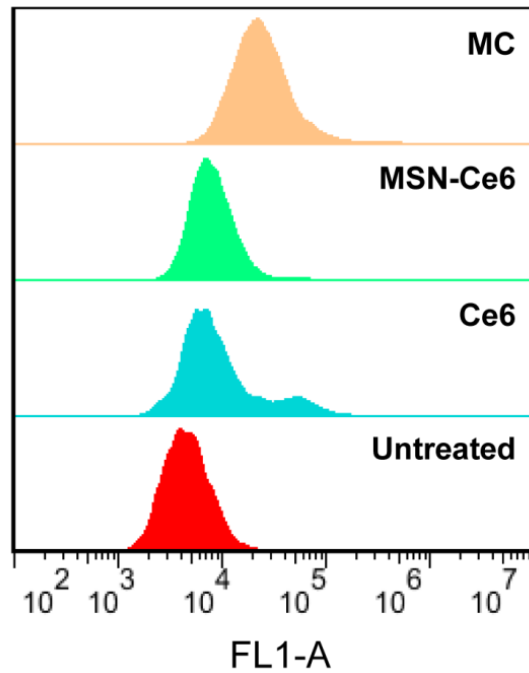
**Figure S11.** UV-vis absorption spectrum and photo (inset) of MB after degraded by the Mn<sup>2+</sup>-mediated Fenton-like reaction.



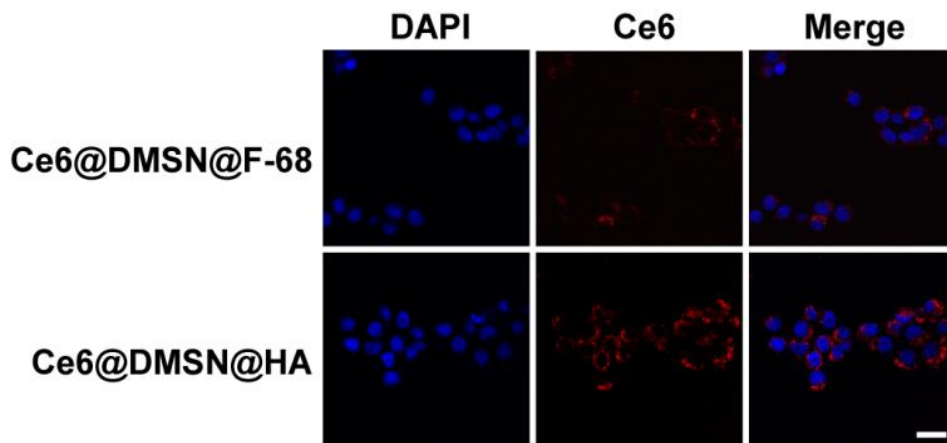
**Figure S12.** Mean fluorescence intensity (%) of H<sub>2</sub>O<sub>2</sub> in 4T1 cells after different treatments (n=3). Results are presented as means  $\pm$  s.d. \**P* < 0.05, \*\**P* < 0.01, and \*\*\**P* < 0.001 determined by Student's t test.



**Figure S13.** Cytotoxicity of Mn<sub>3</sub>O<sub>4</sub> to 4T1 cells at different concentrations as indicated (n=6).

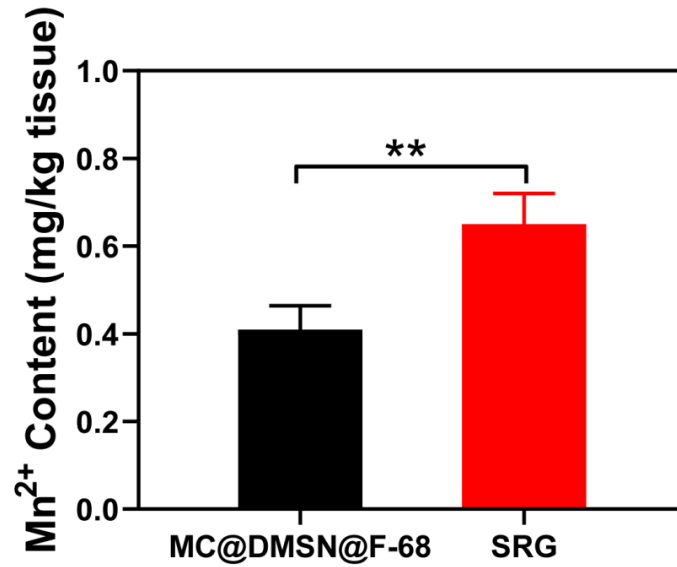


**Figure S14.** Flow cytometer analysis of ROS production after 4T1 cells were treated with different preparations.

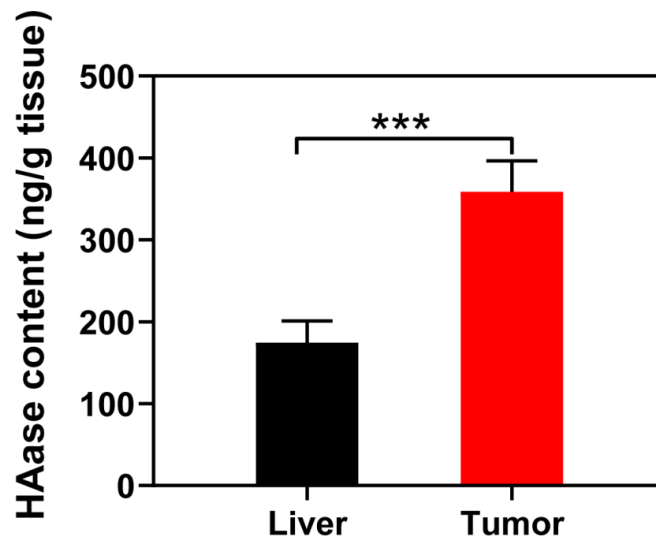


**Figure S15.** CLSM images of 4T1 cells incubated with Ce6@DMSN@F-68 and Ce6@DMSN@HA, respectively (scale bar: 25  $\mu$ m).





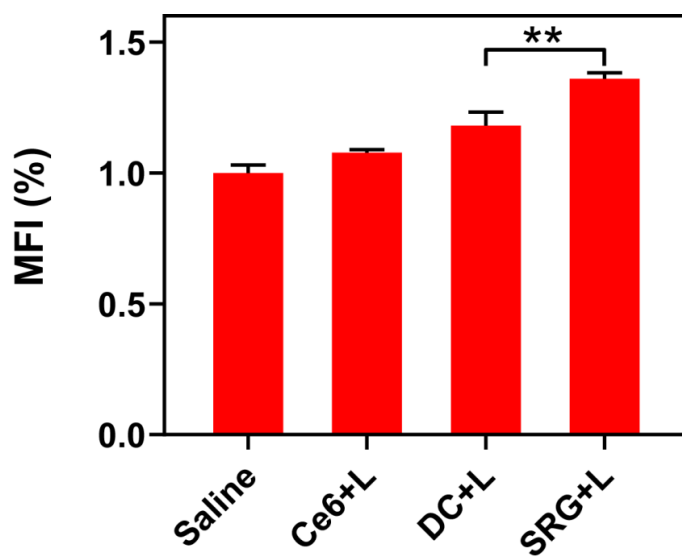
**Figure S16.** Mn<sup>2+</sup> content in tumor tissues of tumor-bearing mice after treated with different preparations (n=3). Results are presented as means  $\pm$  s.d. \**P* < 0.05, \*\**P* < 0.01, and \*\*\**P* < 0.001 determined by Student's t test.



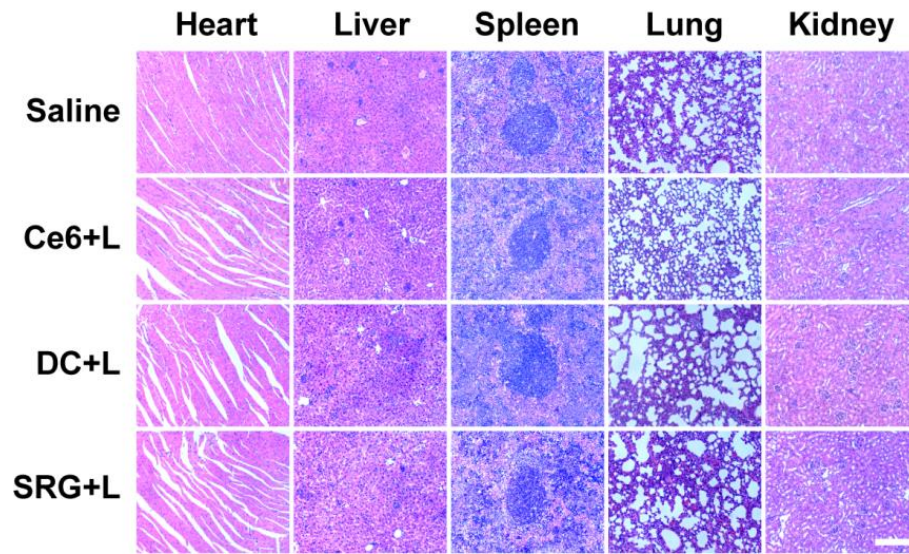
**Figure S17.** HAase expression in liver and tumor tissues of tumor-bearing mice (n=4). Results are presented as means  $\pm$  s.d. \**P* < 0.05, \*\**P* < 0.01, and \*\*\**P* < 0.001 determined by Student's t test.



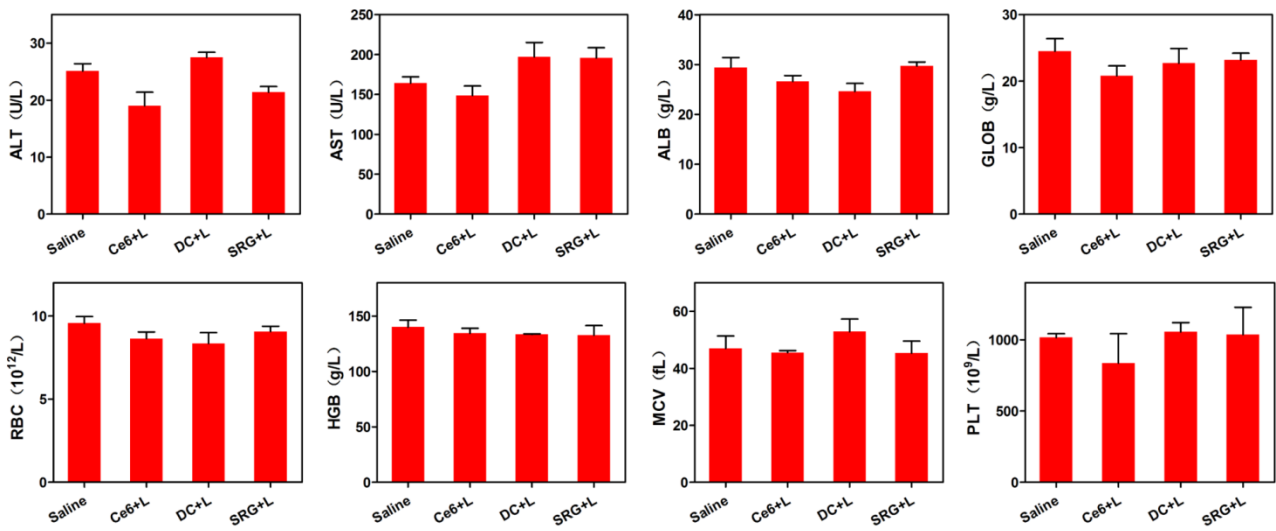
**Figure S18.** Images of tumor-bearing mice exposed to laser.



**Figure S19.** Mean fluorescence intensity (%) of ROS in tumor tissue (n=3). Results are presented as means  $\pm$  s.d. \* $P < 0.05$ , \*\* $P < 0.01$ , and \*\*\* $P < 0.001$  determined by Student's t test.



**Figure S20.** H&E staining of main organs from mice at 14 days post-treatment. (scale bars: 200  $\mu$ m).



**Figure S21.** Blood biochemistry and hematology analysis of the mice after treatment with different preparations for 13 days.