## **Supporting information**

## "Electron transport chain interference" strategy of amplified bacterial ferroptosis and defect-engineered nanozyme for diabetic wound healing

Yanlan Xie<sup>1#</sup>, Huan Wang<sup>1#</sup>, Yalan Wang<sup>1</sup>, Jiajie Liu<sup>1</sup>, Jinming Tong<sup>2</sup>, Tao Wu<sup>2</sup>, Li Yin<sup>2</sup>, Chuan Zhang<sup>2,3\*</sup>, Long Zhao<sup>2,4\*</sup>, Yuan Yong<sup>1,2\*</sup>

- 1 Key Laboratory of Pollution Control Chemistry and Environmental Functional Materials for Qinghai-Tibet Plateau of the National Ethnic Affairs Commission, School of Chemistry and Environment, Southwest Minzu University, Chengdu 610041, China 2 Nanomedicine Innovation Research and Transformation Institute, Affiliated Hospital of North Sichuan Medical College, Nanchong 637000, China
- 3 Biotechnology Innovation Drug Application and Transformation Key Laboratory of Sichuan Province, North Sichuan Medical College, Nanchong 637000, China
- 4 Department of Neurosurgery, Affiliated Hospital of North Sichuan Medical College, Nanchong 637000, China
- # Yanlan Xie and Huan Wang contributed equally to the work.

Corresponding Authors: <u>yongy1816@163.com</u>; <u>cbyzhaolong@163.com</u>; zhangchuanforever@yeah.net

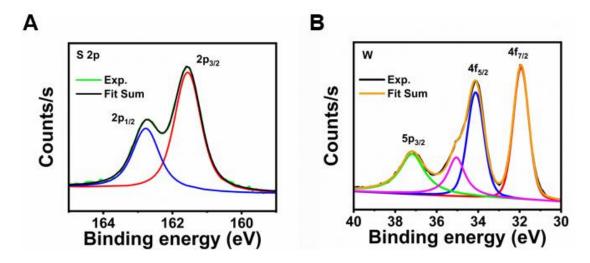
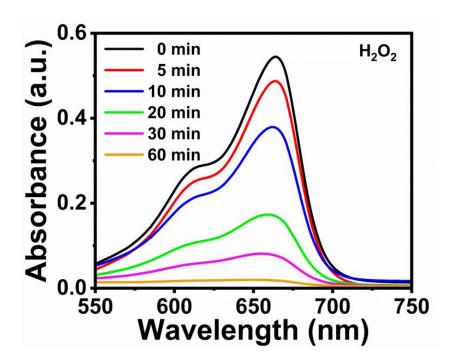
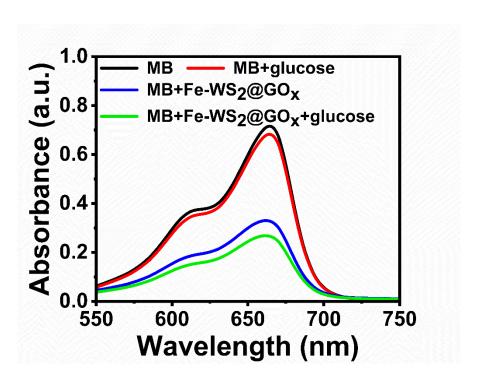


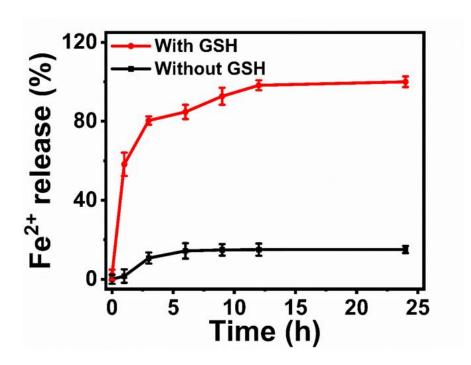
Figure S1. XPS fine spectrum of elements W and S in Fe-WS<sub>2</sub>.



**Figure S2.** The degradation of MB absorbance was measured at different reaction times.



**Figure S3.** GOx-like enzyme activity of Fe-WS<sub>2</sub>@GOx using MB chromogen.



**Figure S4.** Fe<sup>2+</sup> release was determined over time in the presence/absence of GSH.

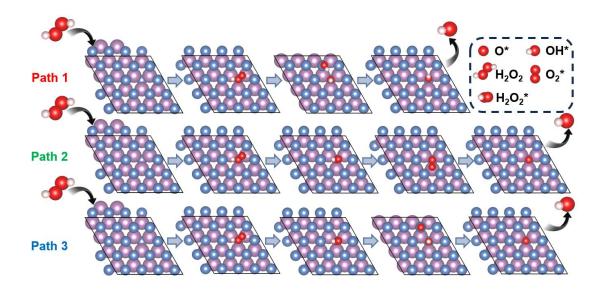
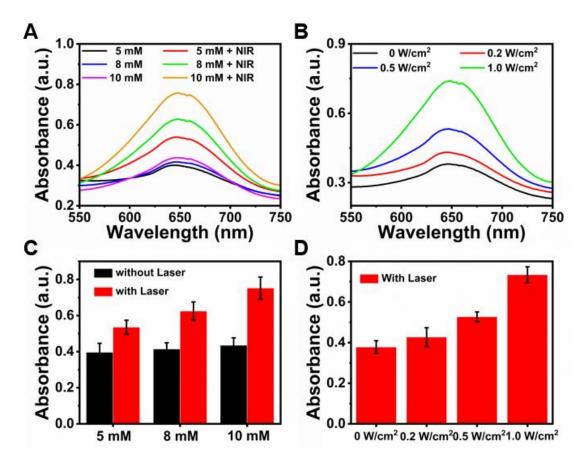
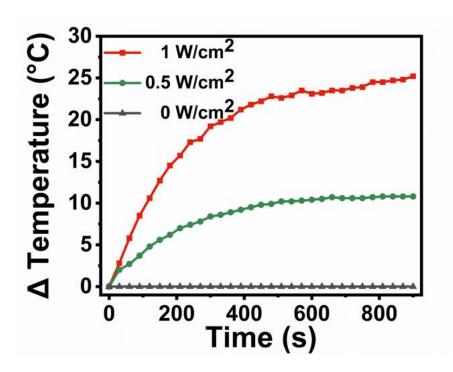


Figure S5. The corresponding reaction pathway for  $H_2O_2$ -catalyzed  $WS_2$ .



**Figure S6.** (A, C) UV absorption of Fe-WS<sub>2</sub> by varying H<sub>2</sub>O<sub>2</sub> concentration in the presence or absence of 808 nm NIR irradiation. (B, D) UV absorption of Fe-WS<sub>2</sub> by varying laser power density in the presence or absence of 808 nm NIR irradiation.



**Figure S7.** Photothermal effects of Fe-WS<sub>2</sub> nanozymes at different laser powers.

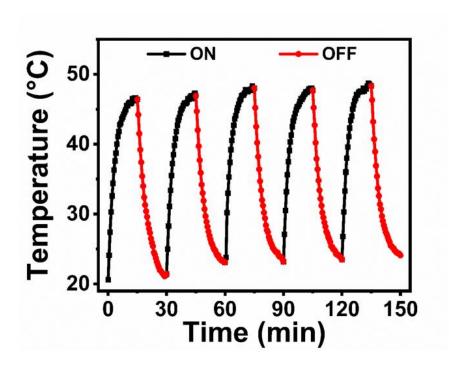
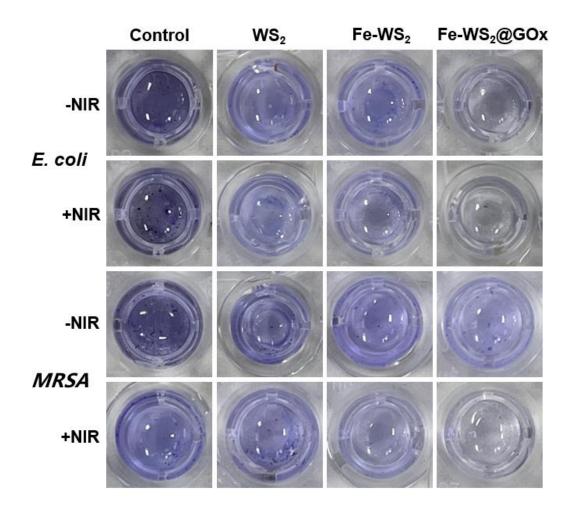
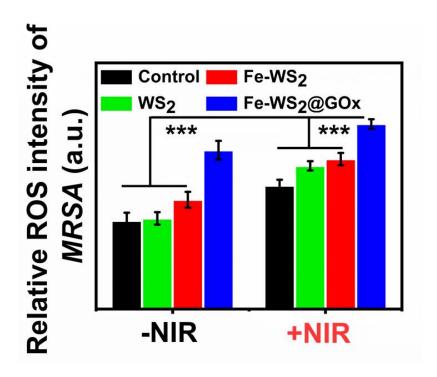


Figure S8. Five photothermal cycles of Fe-WS<sub>2</sub> at 100  $\mu$ g/mL in the NIR (808 nm, 1 W/cm<sup>2</sup>).



**Figure S9.** Bacterial biofilms stained with crystal violet dye after different treatments of *E. Coli* and MRSA.



**Figure S10.** Detection of ROS content in MRSA after different treatments using DCFH-DA probe. N = 3. (\* $P \le 0.05$ , \*\* $P \le 0.01$ , \*\*\* $P \le 0.001$ ).

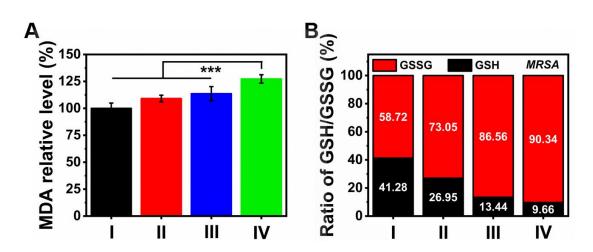
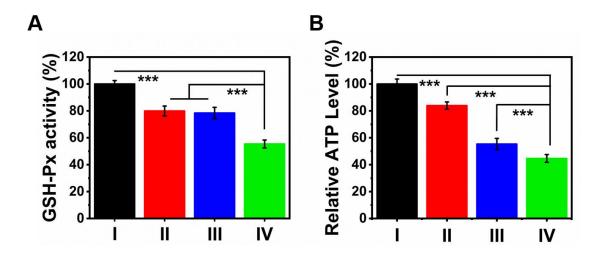
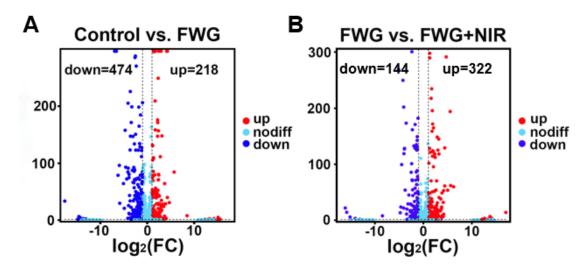


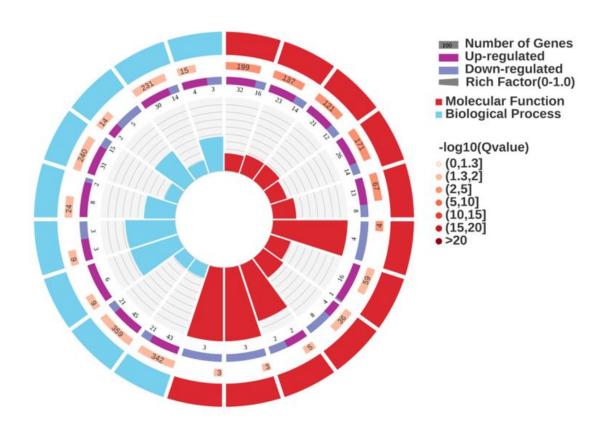
Figure S11. Detection of (A) MDA levels and (B) GSH/GSSG ratio within MRSA after different treatments. N = 3. (\*P  $\leq$  0.05, \*\*P  $\leq$  0.01, \*\*\*P  $\leq$  0.001). (I: Control; II: NIR; III: Fe-WS<sub>2</sub>@GOx; IV: Fe-WS<sub>2</sub>@GOx+NIR)



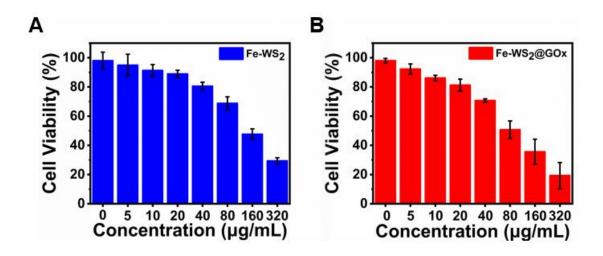
**Figure S12.** Detection of (A) GSH-Px and (B) ATP levels within MRSA by different treatments. N = 3. (\*P  $\leq$  0.05, \*\*P  $\leq$  0.01, \*\*\*P  $\leq$  0.001). (I: Control; II: NIR; III: Fe-WS<sub>2</sub>@GOx; IV: Fe-WS<sub>2</sub>@GOx+NIR)



**Figure S13.** Volcano plots of up-regulated and down-regulated gene expression in (A) Control vs. FWG and (B) FWG vs. FWG+NIR groups. (FWG+NIR=Fe-WS<sub>2</sub>@GOx+NIR; FWG=Fe-WS<sub>2</sub>@GOx)



**Figure S14.** Differential expression results of genes analyzed by Fe-WS<sub>2</sub>@GOx+NIR cyclic gene mapping.



**Figure S15.** Cells viability of L929 assayed with different concentrations of (A) Fe-WS<sub>2</sub> and (B) Fe-WS<sub>2</sub>@GOx. N = 3.

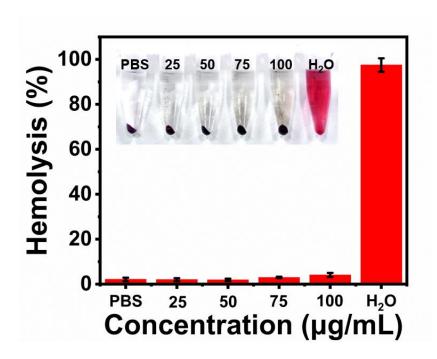
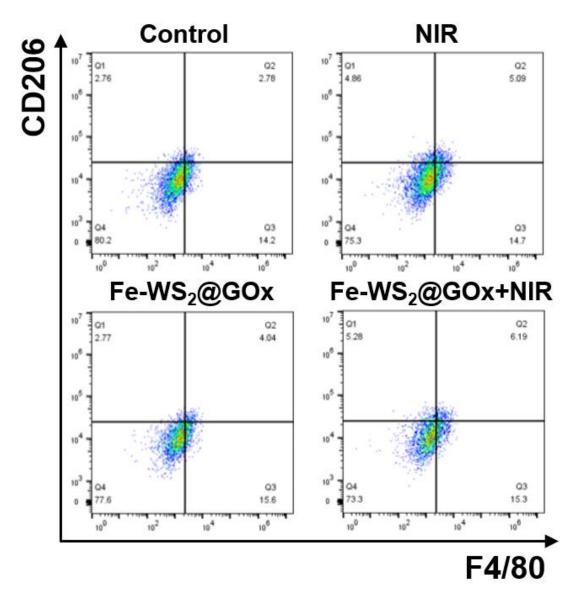
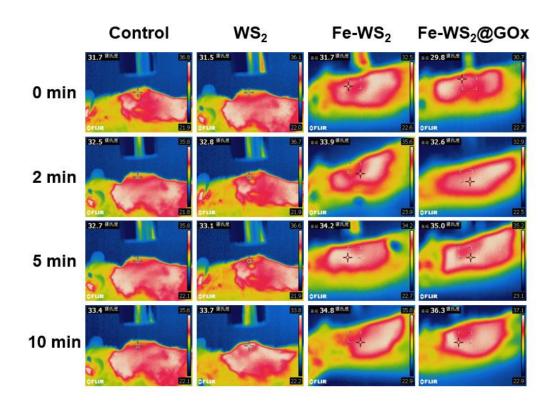


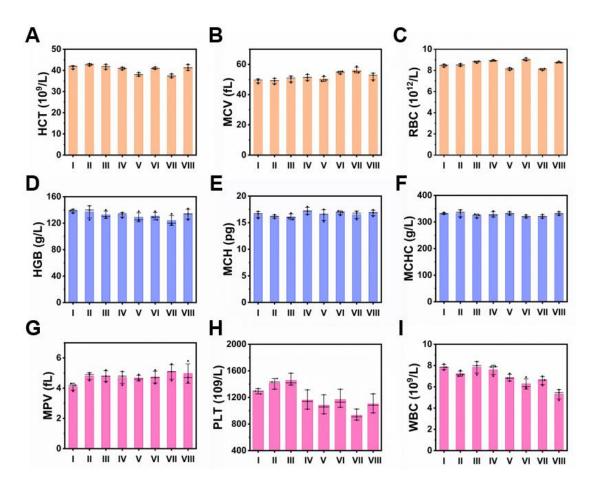
Figure S16. Hemolysis assay results after different treatments. N = 3.



**Figure S17.** Macrophages were first polarized to M1 using LPS, and then the M2 phenotype of macrophages was detected using flow cytometry after different treatments.



**Figure S18.** Mouse wounds were irradiated with 808 nm NIR (1.0 W/cm<sup>2</sup>) for 10 min and temperature changes were recorded.



**Figure S19.** Blood routine test of mice in each group was tested. N = 3. (I: Control; II: WS<sub>2</sub>; III: Fe-WS<sub>2</sub>; IV: Fe-WS<sub>2</sub>@GOx; V: NIR; VI: WS<sub>2</sub>+NIR; VII: Fe-WS<sub>2</sub>+NIR; VIII: Fe-WS<sub>2</sub>@GOx+NIR)

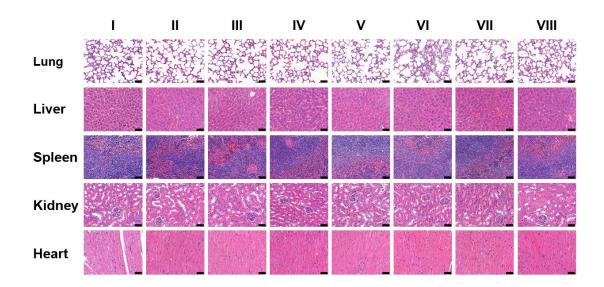


Figure S20. H&E staining of mice after different treatments. Scale bar = 50 μm. (I: Control; II: WS<sub>2</sub>; III: Fe-WS<sub>2</sub>; IV: Fe-WS<sub>2</sub>@GOx; V: NIR; VI: WS<sub>2</sub>+NIR; VII: Fe-WS<sub>2</sub>+NIR; VIII: Fe-WS<sub>2</sub>@GOx+NIR)