Supporting Information

Ultrasound-controllable ROS-responsive nanoplatform for O_2 and NO generation to enhance sonodynamic therapy against multidrug-resistant bacterial infection

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Figure S1. (A) Synthetic route and chemical structure of FOT. (B) ¹H NMR spectrum of synthesized FOT in CDCl₃ δ 8.20 (t, J = 4.0 Hz, 2H), 7.83 (dd, J = 16.0, 8.0 Hz, 1H), 7.66 (q, J = 8.0 Hz, 2H). (C) The MS of synthesized FOT.



Figure S2. Transmission electron microscope images of Fe₃O₄ NPs.



Figure S3. (A) UV–vis absorption spectra and (B) Fluorescence intensity of FOT/Fe₃O₄@Lipo-ICG at different concentration. (C) Fluorescence intensity of ICG in H₂O and DMSO.



Figure S4. Colloidal stability of FOT/Fe₃O₄@Lipo-ICG in PBS. Data are presented as mean ± s.d. (n = 3).



Figure S5. (A) The equations of the Fenton reaction. (B) The mechanism of GSH-mediated NO release from FOT.



Figure S6. (A) Consumption of DPBF due to $\bullet O^{2-}$, (B) Consumption of MB due to $\bullet OH$, (C) O_2 generation and (D) NO generation of H_2O_2 with US irradiation. US irradiation: 1.0 MHz, 1.0 W/cm2, 40% cycle.



Figure S7. Electron spin resonance spectra of ${}^{1}O_{2}$ (A) and $\cdot OH$ (B) in FOT/Fe₃O₄@Lipo-ICG and FOT/Fe₃O₄@Lipo-ICG+H₂O₂ dispersions after US irradiation or not. The concentration of H₂O₂ was 1 mM. US irradiation: 1.0 MHz, 1.0 W/cm², 40% cycle.



Figure S8. SEM images of MRSA after different treatments. US irradiation: 1.0 MHz, 1.0 W/cm², 40% cycle.



Figure S9. Photographs of the MRSA-infected skin during treatment with FOT/Fe $_3O_4@$ Lip-ICG without US irradiation at 0, 6 and 12 days.



Figure S10. Cell viability of HUVEC and 3T3 treated with different concentrations of FOT/Fe₃O₄@Lipo-ICG. Data are presented as mean \pm s.d. (n = 3).

Table S1. Properties of liposomes.

Entry	Liposomes	Size (nm)	PDIª	Zeta potential (mv)
1	Lipo-ICG	90.3 ± 7.4	0.18 ± 0.04	-22.5 ± 0.4
2	FOT@Lipo-ICG	140.2 ± 6.9	0.17 ± 0.02	-26.6 ± 0.2
3	FOT/Fe ₃ O ₄ @Lipo-ICG	165.1 ± 10.8	0.18 ± 0.02	-27.7 ± 0.5

^a Polydispersity index (PDI).

Table S2. LE and EE of FOT/Fe₃O₄@Lipo-ICG.

Linocomos	LEª (%)		EE ^b (%)					
Liposomes	FOT	Fe ₃ O ₄	FOT	Fe ₃ O ₄				
FOT@Lipo-ICG	5.4 ± 0.6	-	70.0 ± 4.4	-				
FOT/Fe ₃ O ₄ @Lipo-ICG	5.4 ± 0.5	8.0 ± 0.7	70.0 ± 3.2	80.0 ± 4.3				

^a Loading efficiency (LE).

^bEntrapment efficiency (EE).