

Supporting Information for

ORIGINAL ARTICLE

**Boosting 5-ALA-based photodynamic therapy by a liposomal nanomedicine through intracellular iron ion regulation**

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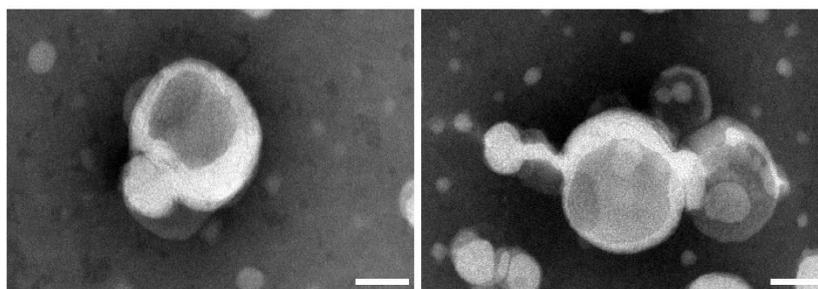
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Received 26 August 2020; received in revised form 8 November 2020; accepted 21 November 2020

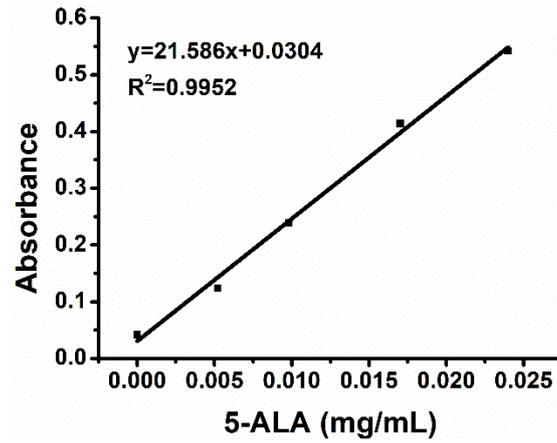
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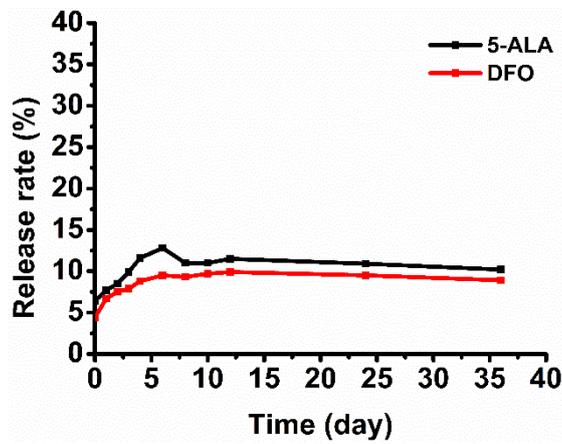
<sup>†</sup>These authors made equal contributions to this work.



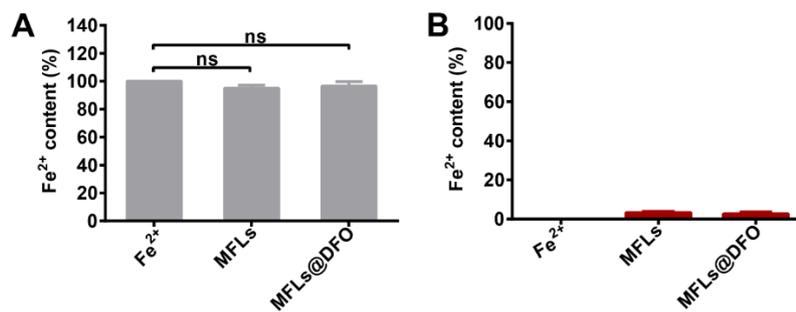
**Figure S1** The transmission electron microscope of MFLs. Scale bar = 50 nm.



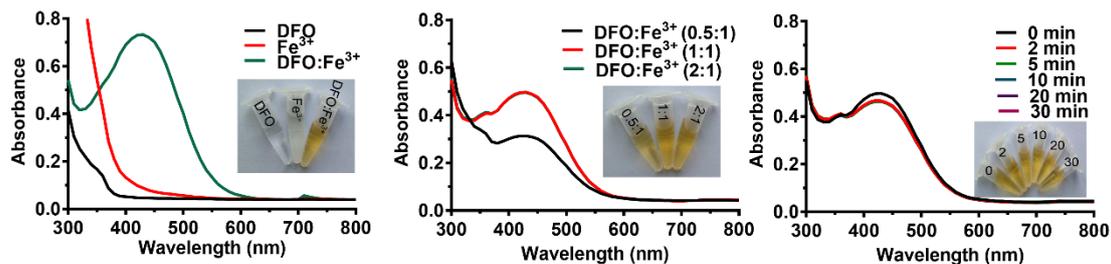
**Figure S2** The standard curve of 5-ALA.



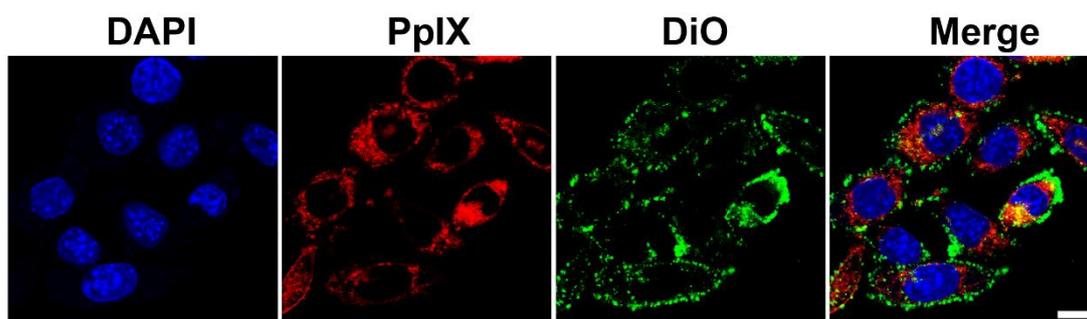
**Figure S3** The release profiles of 5-ALA and DFO from MFLs.



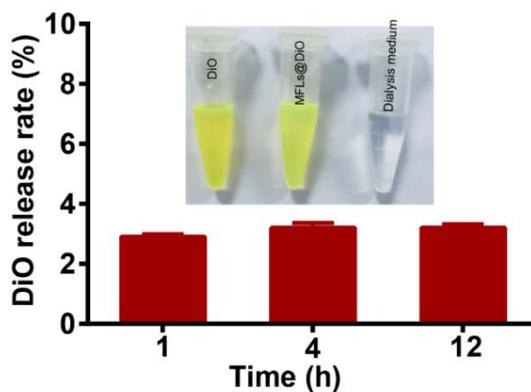
**Figure S4** The content of iron in supernatant (A) and MFLs, MFLs@DFO (B). Data are presented as means  $\pm$  SD ( $n = 3$ ). ns, not significant.



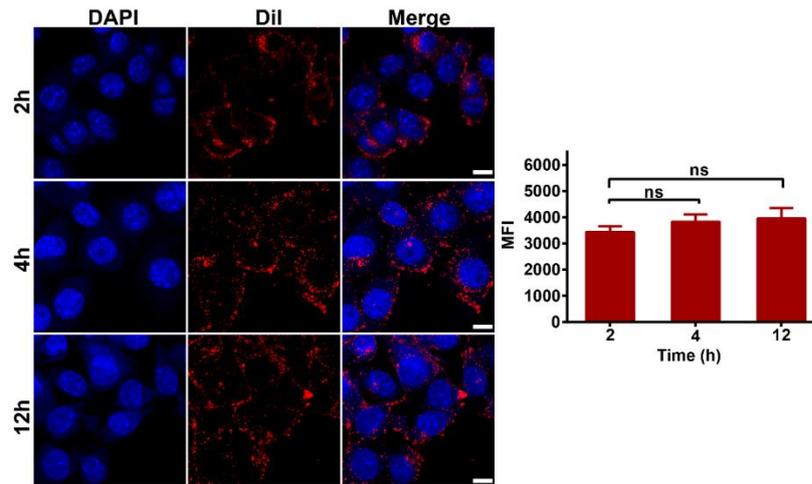
**Figure S5** The UV–Vis spectra of DFO complex  $\text{Fe}^{3+}$  at 430 nm and picture of complexing appearance (the inset).



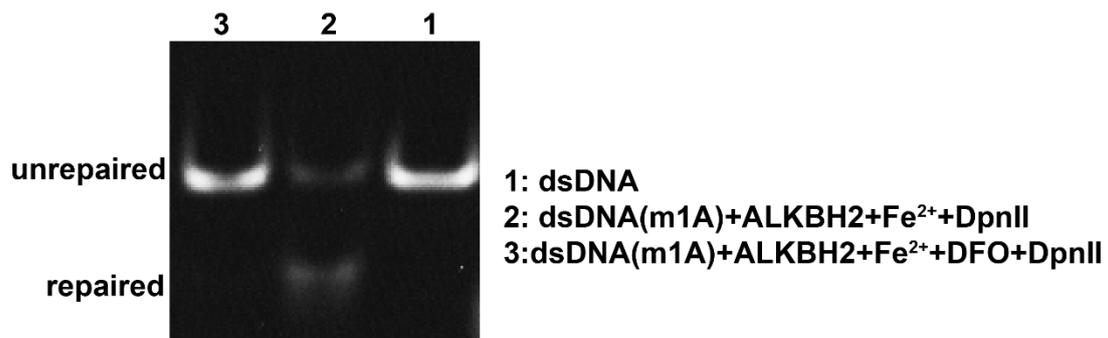
**Figure S6** The distribution of MFLs@5-ALA/DFO labeled with DiO in B16-F10 cells by CLSM. Scale bar = 10  $\mu\text{m}$ .



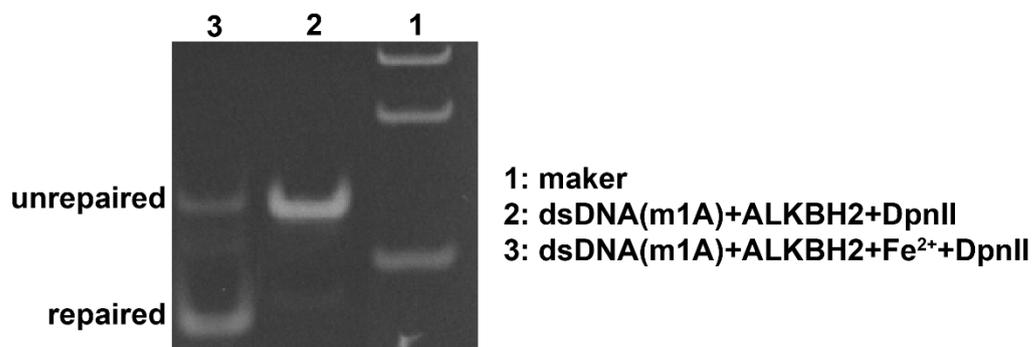
**Figure S7** The release rate of DiO from MFLs@DiO at 12 h and the appearance picture (the inset). Data are presented as means  $\pm$  SD ( $n = 3$ ).



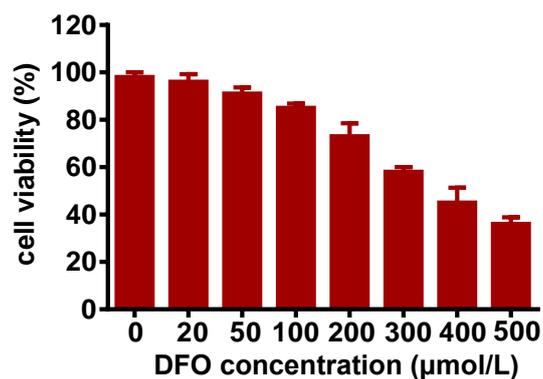
**Figure S8** The stability and semi-quantitative analysis of DiI-labeled cell membrane. Scale bar = 10  $\mu$ m. Data are presented as means  $\pm$  SD ( $n = 3$ ). ns, not significant.



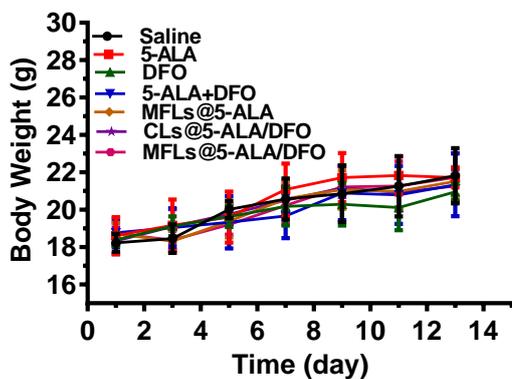
**Figure S9** DFO inhibits ALKBH2 repair of m1A in dsDNA by using the DpnII digestion assay.



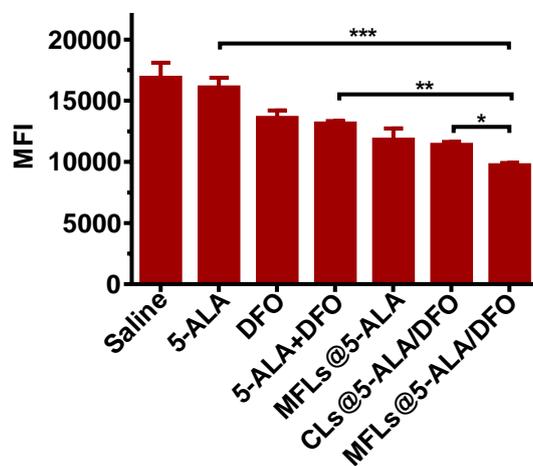
**Figure S10** The confirmation of Fe<sup>2+</sup> dependent ALKBH2 repair of m1A in dsDNA by using the DpnII digestion assay.



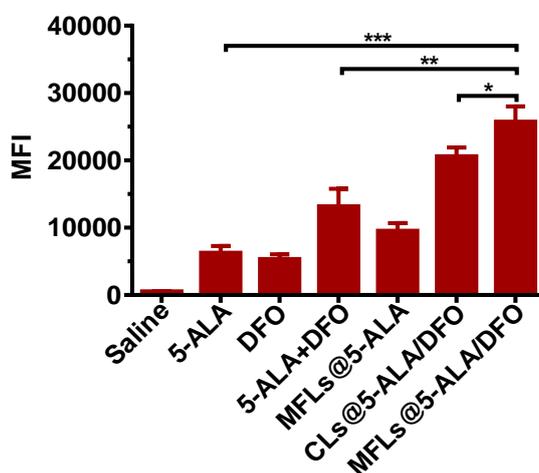
**Figure S11** The cell viability of DFO after incubated with B16-F10 cells. Data are presented as means  $\pm$ SD ( $n = 3$ ).



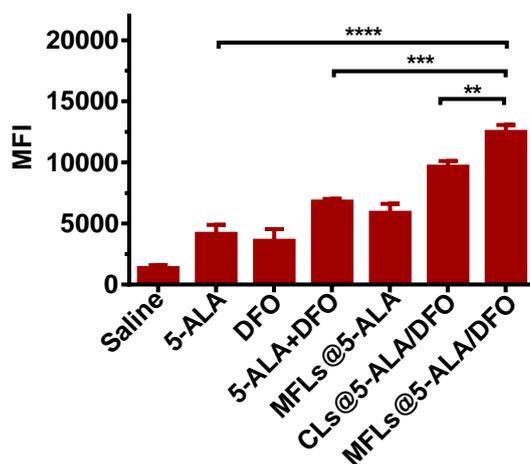
**Figure S12** The weight of tumor-bearing mice after treated 14 days with different Preparations. Data are presented as means  $\pm$ SD ( $n = 6$ ).



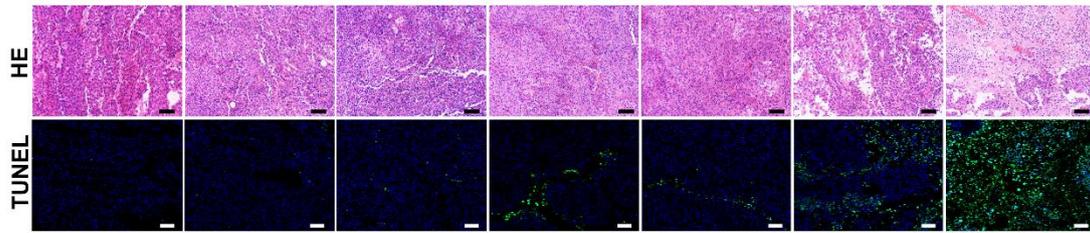
**Figure S13** The semi-quantitative analysis of  $\text{Fe}^{2+}$  in tumor tissues after treated 14 days with different preparations. Data are presented as means  $\pm$  SD ( $n = 3$ ). \*\*\* $P < 0.001$ , \*\* $P < 0.01$ , \* $P < 0.05$ .



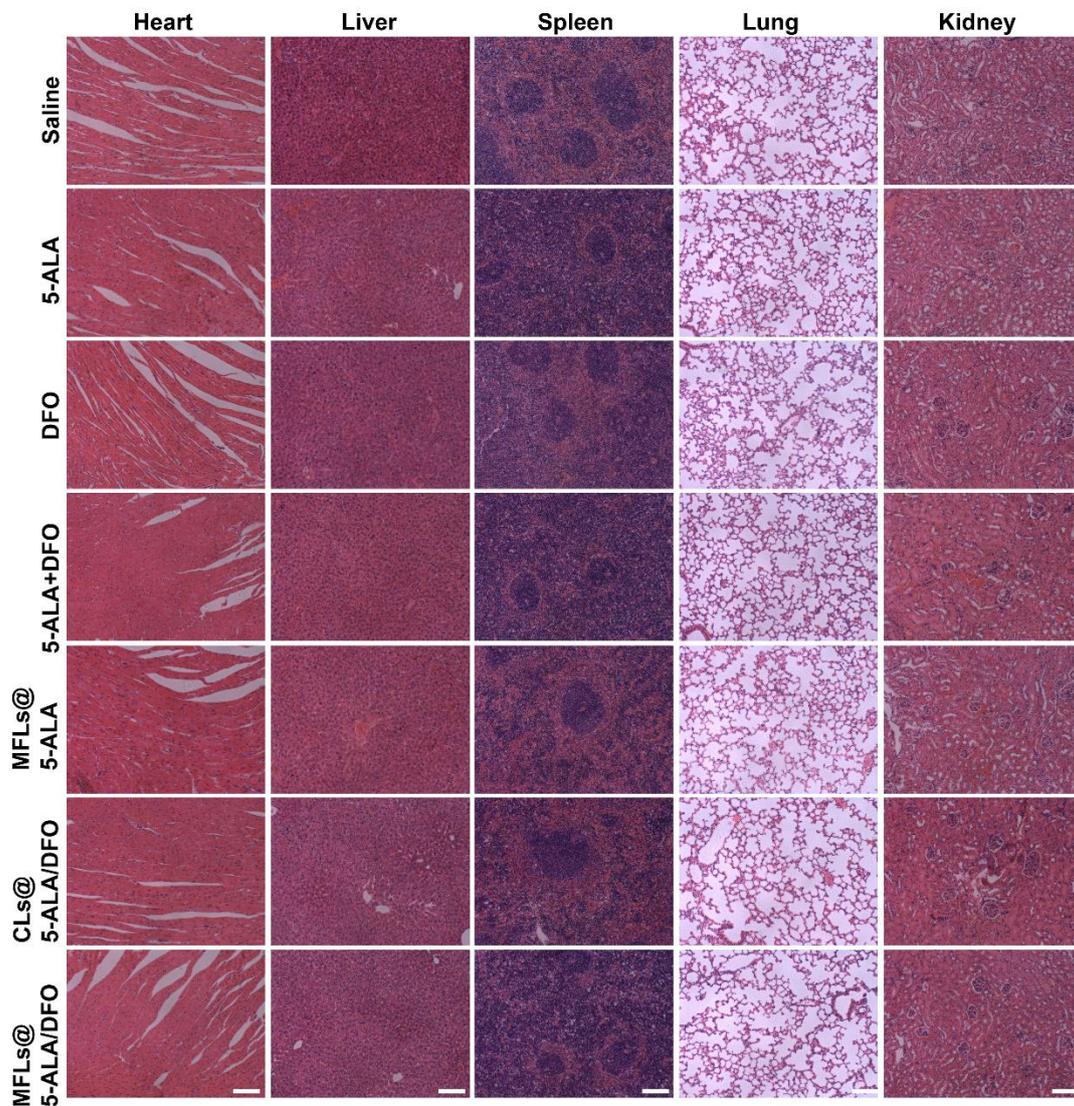
**Figure S14** The semi-quantitative analysis of PpIX in tumor tissues after treated 14 days with different preparations. Data are presented as means  $\pm$  SD ( $n = 3$ ). \*\*\* $P < 0.001$ , \*\* $P < 0.01$ , \* $P < 0.05$ .



**Figure S15** The semi-quantitative analysis of ROS in tumor tissues. Data are presented as means  $\pm$  SD ( $n = 3$ ). \*\*\*\* $P < 0.0001$ , \*\*\* $P < 0.001$ , \*\* $P < 0.01$ .



**Figure S16** H&E staining and TUNEL staining of tumor tissues. The tumor tissues were exfoliated from different groups after treated 14 days with different preparations. Scale bar = 50  $\mu$ m.



**Figure S17** The histologic assessments of major organs with H&E staining. Scale bar = 200  $\mu$ m.