## **Supporting information for**

## Bioinspired Multifunctional Melanin-Based Nanoliposome for Photoacoustic/Magnetic Resonance Imaging-Guided Efficient Photothermal Ablation of Cancer

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## **Supplementary Figures**



Figure S1. Schematic illustration of detailed synthetic procedure for multifunctional Lip-Mel.



Figure S2. Photographs of lyophilized powder of (A) Lip-Mel and (B) Lip; (C) Zeta potential of Lip-Mel and Lip.



Figure S3. In vitro release profile of melanin from Lip-Mel nanocomposite in PBS.



**Figure S4.** Serum biochemical indexes analysis of BALB/c mice from the control group and the experimental groups 1, 3, 7 and 14 days post intravenous injection of Lip-Mel, Lip and melanin.



**Figure S5.** Routine blood examination of BALB/c mice from the control group and the experimental groups 1, 3,7 and 14 days post intravenous injection of Lip-Mel, Lip and melanin.



Figure S6. H&E staining of major organs from the control group. All the scale bars are 100 µm.



Figure S7. H&E staining of major organs from the experimental group 1 day post intravenous injection of Lip-Mel. All the scale bars are  $100 \mu m$ .



**Figure S8.** H&E staining of major organs from the experimental group 1 day post intravenous injection of Lip. All the scale bars are  $100 \mu m$ .



Figure S9. H&E staining of major organs from the experimental group 1 day post intravenous injection of Mel. All the scale bars are  $100 \mu m$ .



**Figure S10.** H&E staining of major organs from the experimental group 3 days post intravenous injection of Lip-Mel. All the scale bars are 100  $\mu$ m.



Figure S11. H&E staining of major organs from the experimental group 3 days post intravenous injection of Lip. All the scale bars are  $100 \mu m$ .



Figure S12. H&E staining of major organs from the experimental group 3 days post intravenous injection of Mel. All the scale bars are 100  $\mu$ m.



**Figure S13.** H&E staining of major organs from the experimental group 7 days post intravenous injection of Lip-Mel. All the scale bars are 100  $\mu$ m.



**Figure S14.** H&E staining of major organs from the experimental group 7 days post intravenous injection of Lip. All the scale bars are  $100 \mu m$ .



Figure S15. H&E staining of major organs from the experimental group 7 days post intravenous injection of Mel. All the scale bars are  $100 \mu m$ .



**Figure S16.** H&E staining of major organs from the experimental group 14 days post intravenous injection of Lip-Mel. All the scale bars are 100 µm.



Figure S17. H&E staining of major organs from the experimental 14 days post intravenous injection of Lip. All the scale bars are  $100 \mu m$ .



Figure S18. H&E staining of major organs from the experimental group 14 days post intravenous injection of Mel. All the scale bars are  $100 \mu m$ .



**Figure S19.** In vivo bio-distribution of Lip-Mel. (A) In vivo fluorescence images of tumor after intravenous injection of Lip-Mel labeled with DiR at different time points. (B) Quantitative tumor fluorescence signal intensities at corresponding time points. (n = 3, mean  $\pm$  SD). (C) Ex vivo fluorescence images of major organs and tumor dissected from mice at 24 h post injection. (D) Quantitative bio-distribution of Lip-Mel in mice.



**Figure S20.** Magnetization hysteresis loops of (A) Lip-Mel and (B) melanin at 300 K in the range of -12 kOe < H < +12 kOe.



**Figure S21.** Plot of  $1/T_1$  as a function of melanin concentration in Lip-Mel. The slope of the curve is defined as the specific relativity of  $r_1$ .



**Figure S22.** (A)  $T_1$ -weighted MRI detection of Lip-Mel in living mice. Mice were injected subcutaneously (region enveloped by red dotted line) with Lip-Mel at concentrations of 0, 2, 4 (from left to right in upper layer), and 6, 8, 10 (from left to right in bottom layer) mg/mL. (B) Ratio of intensity (Lip-Mel/muscle) at different concentrations of Lip-Mel.



Figure S23. IR thermal images of pure water and Lip-Mel suspension at different concentrations (45.45, 90.90, 181.80, 363.60  $\mu$ g/mL) as a function of irradiation duration using an 808 nm laser (1.50 W/cm<sup>2</sup>).



Figure S24. (A) Plot of temperature change ( $\triangle$ T) and (B) IR thermal images of pure water and Lip-Mel suspension at different concentrations (45.45, 90.90, 181.80, 363.60 µg/mL) as a function of irradiation duration using an 808 nm laser (1.00 W/cm<sup>2</sup>).



**Figure S25.** IR thermal images of Lip-Mel suspension at different power densities of an 808 nm laser (0.75, 1.00, 1.25 and 1.50 W/cm<sup>2</sup>) as a function of irradiation duration (melanin concentration:  $363.60 \mu g/mL$ ,  $100 \mu L$ ).



**Figure S26.** Recycling heating profiles of Lip-Mel (melanin concentration: 363.60  $\mu$ g/mL, 100  $\mu$ L) as irradiated by an 808 nm laser (1.00 W/cm<sup>2</sup>) for five laser on/off cycles.



**Figure S27.** IR thermal images of Lip-Mel (melanin concentration:  $363.60 \ \mu\text{g/mL}$ ), melanin ( $363.60 \ \mu\text{g/mL}$ ) and Lip suspension as a function of irradiation duration using an 808 nm laser ( $1.50 \ \text{W/cm}^2$ ).



**Figure S28.** Temperature changes of MDA-MB-231 tumor regions of the four groups (control, Lip-Mel only, Laser only and Lip-Mel + Laser group) as a function of irradiation duration using an 808 nm laser (1.50 W/cm<sup>2</sup>, 10 min).