

## Supporting Information

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Ultrasound-Responsive Oxygen-Carrying Pollen for Enhancing Chemo-Sonodynamic Therapy of Breast Cancer

Baojie Wen, Danqing Huang, Chuanhui Song, Jingyang Shan and Yuanjin Zhao\*

## **Supporting Information**

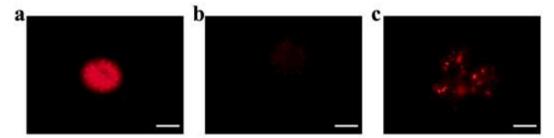
## Ultrasound-responsive oxygen-carrying pollen for enhancing chemo-sonodynamic therapy of breast cancer

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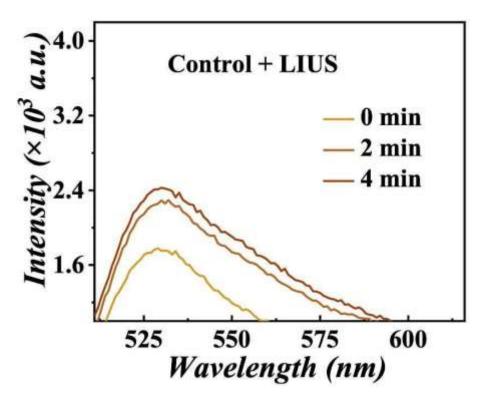
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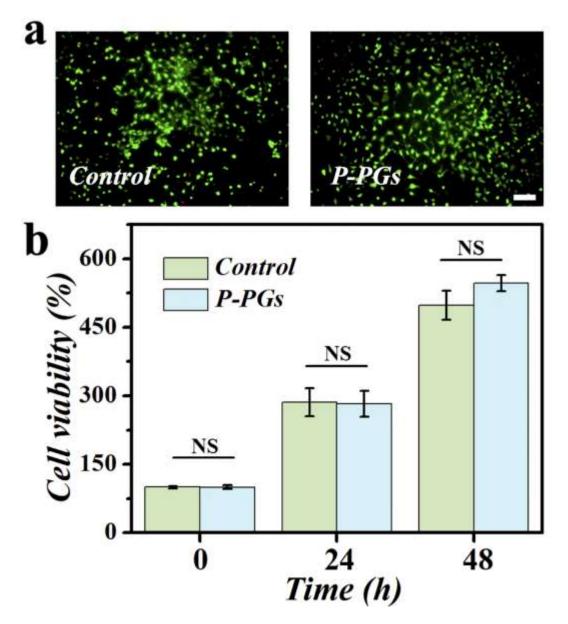
## **Supporting Figures:**



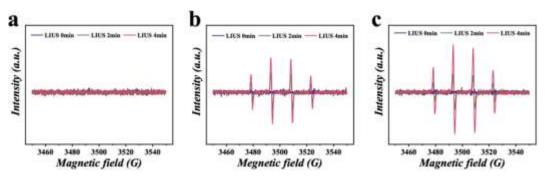
**Figure S1.** Fluorescence microscopy of fluorescent bleached and DOX-loaded carbonized pollen. (a) Fluorescence microscopy of carbonized pollen. (b) Fluorescence microscopy of fluorescent bleached carbonized pollen. (c) Fluorescence microscopy of DOX loaded with fluorescent bleached pollen. Scale bar is  $20 \mu m$  in (a-c).



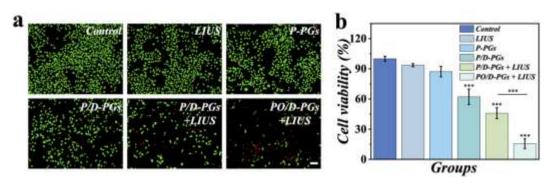
**Figure S2.** Fluorescence intensity curves of SOSG and pure water mixed solutions after different intervals of LIUS irradiation.



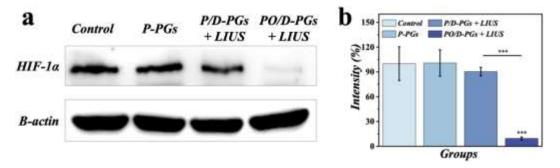
**Figure S3.** Biocompatibility study of P-PGs on 3T3 cells. (a) Live/dead staining images of 3T3 cells cultured in blank medium and co-cultured with P-PGs for 48h. Scale bar is  $100 \mu m$ . (b) CCK-8 results of 3T3 cells at different time periods; error bars indicate standard deviation.



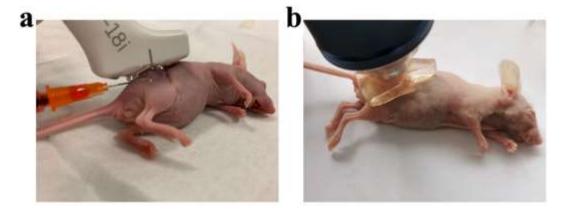
**Figure S4.** Detection of •OH via ESR analysis. (a) The •OH of the control group was determined after ultrasonic irradiation at different time intervals by ESR. (b) The •OH of the P/D-PGs solution was determined after ultrasonic irradiation at different time intervals by ESR. (c) The •OH of the PO/D-PGs solution was determined after ultrasonic irradiation at different time intervals by ESR.



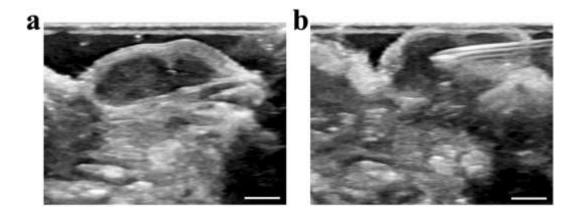
**Figure S5.** Cytotoxicity of MDA-MB-231 cells. (c) Live/dead staining images of MDA-MB-231 cells in different cell groups. (d) The relative cell viability of MDA-MB-231 cells in different groups (n = 6). The scale bar is 100  $\mu$ m.



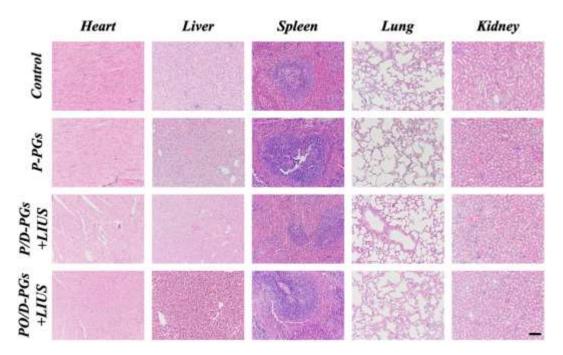
**Figure S6.** Western blotting assay of HIF-1 $\alpha$  in different groups. (a) Western Blotting results of HIF-1 $\alpha$  expression. (b) Quantification of HIF-1 $\alpha$  expression (n = 3).



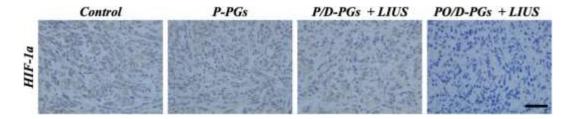
**Figure S7.** The recording images of the treatment. (a) US-guided intratumoral injection of pollen grains. (b) Irradiation with LIUS after injection.



**Figure S8.** Ultrasound imaging of the process of injecting pollen microcarriers. (a) Grayscale ultrasound image of the tumor during ultrasound examination. (b) Ultrasound-guided needle insertion into the tumor. Scale is 2.5 mm in (a) and (b).



**Figure S9.** H&E staining of major organs of mice in each experimental group. The scale bar represents  $100 \ \mu m$ .



**Figure S10.** Immunohistochemical results of HIF-1 $\alpha$  in different groups of breast cancer. Scale bar is 40  $\mu m$ .