

## Supporting Information for

## ORIGINAL ARTICLE

# Localized light-triggered release macrophage cytopharmaceuticals containing *O*-nitrobenzyl group for enhanced solid tumor cell-chemotherapy

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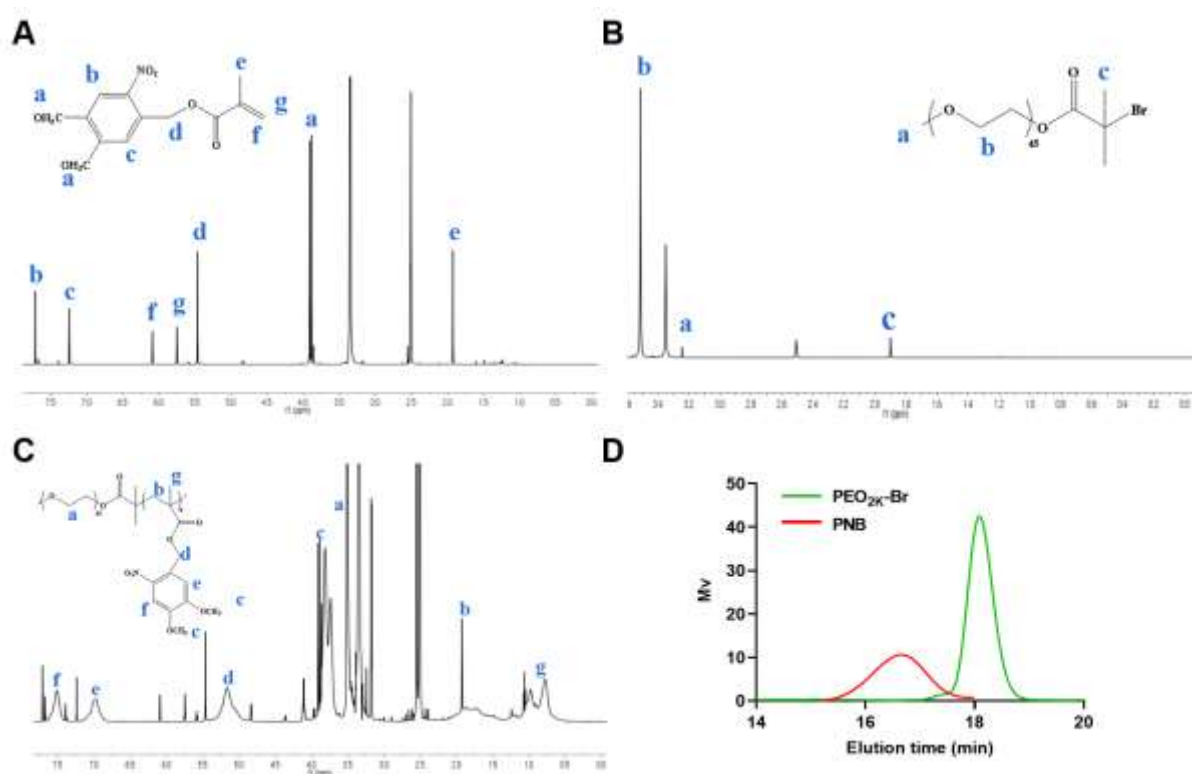
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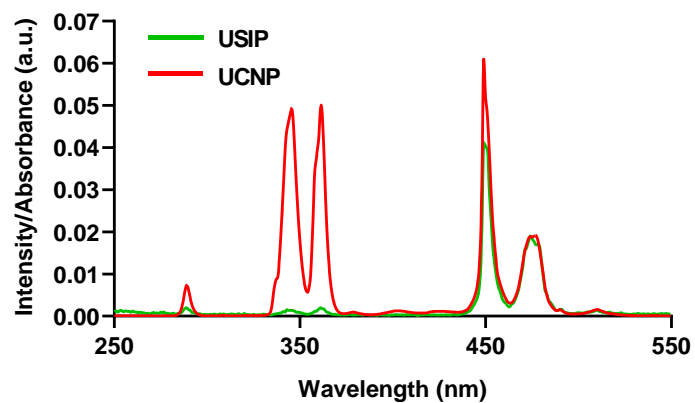
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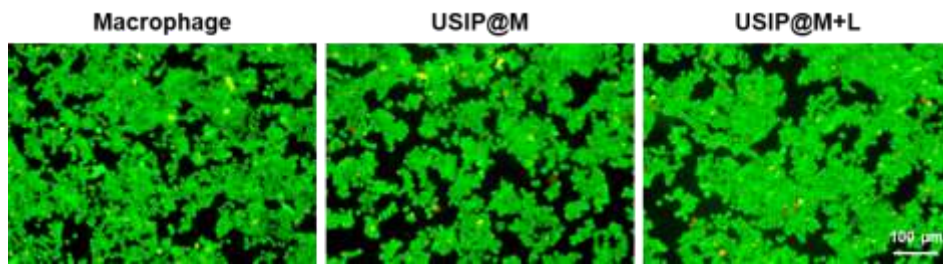
<sup>†</sup>These authors made equal contributions to this work.



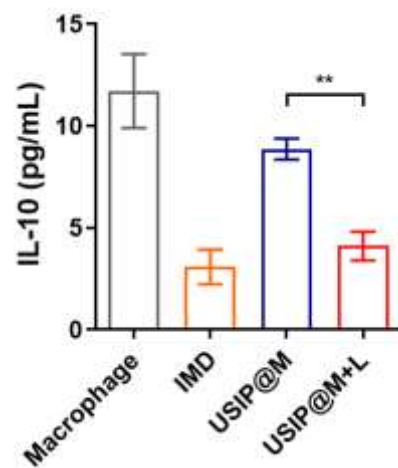
**Figure S1** (A-C) The <sup>1</sup>H-NMR spectrum of (A) the monomer NBMA, (B) PEO<sub>2K</sub>-Br and (C) PNB. (D) Gel permeation chromatography spectrums of PEO<sub>2K</sub>-Br and PNB.



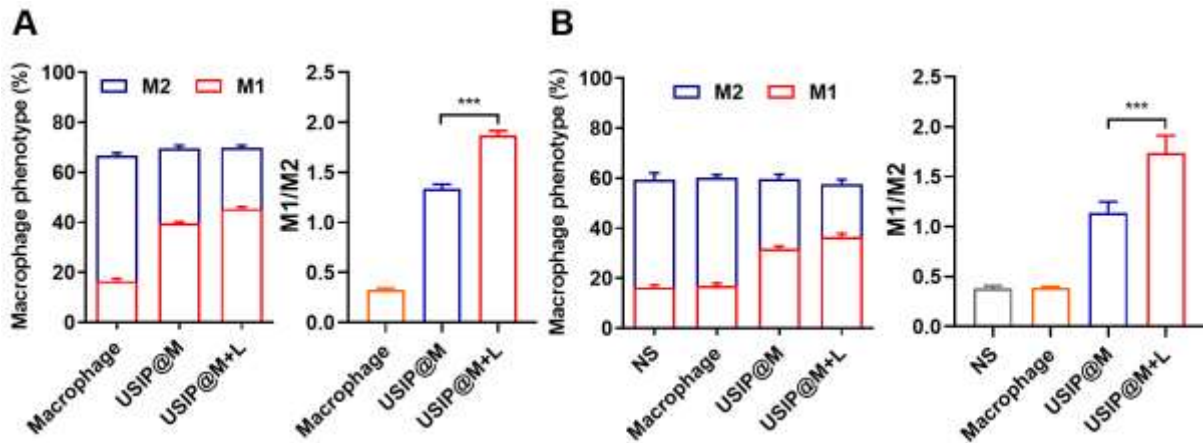
**Figure S2** The emission spectra of the USIPs (green line) and the UCNP (red line).



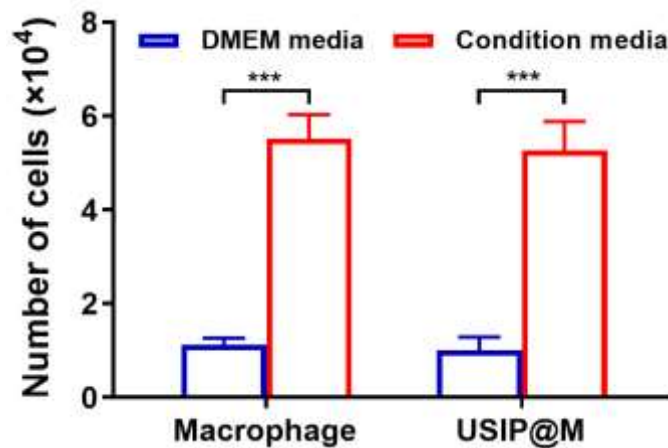
**Figure S3** Phototoxicity investigation of USIP@M after NIR irradiation. USIP@M was irradiated with NIR light (980 nm, 1 W, 10 min), and then detected by Calcein-AM/PI live cell/dead cell double staining kit, scale bar = 100  $\mu\text{m}$ .



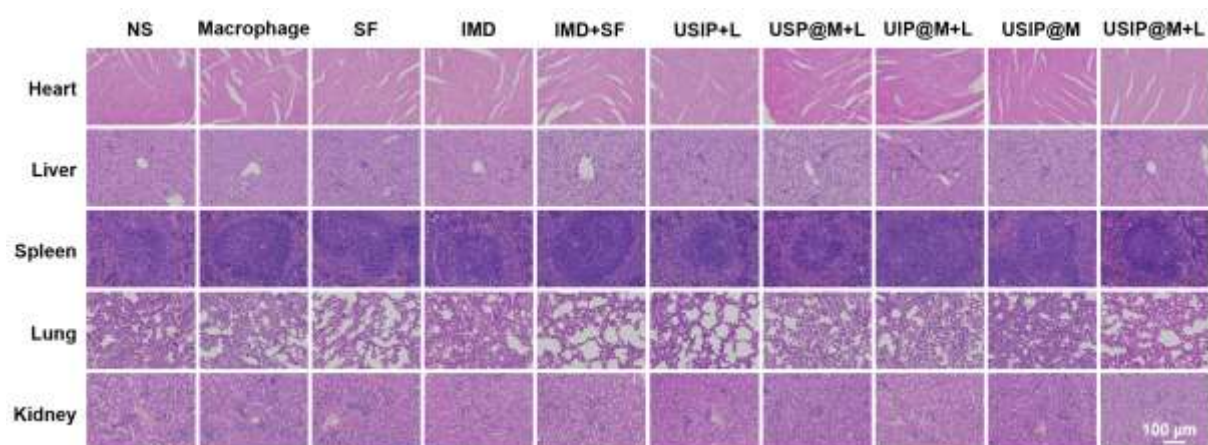
**Figure S4** Cytokine IL-10 analysis of USIP@M 24 h after irradiation with or without NIR light. The IMD group (RAW264.7 cells incubated with IMD) contained the same concentration of IMD as USIP@M and USIP@M+L group. Data are presented as mean $\pm$ SD ( $n=3$ ).  $**P < 0.01$ .



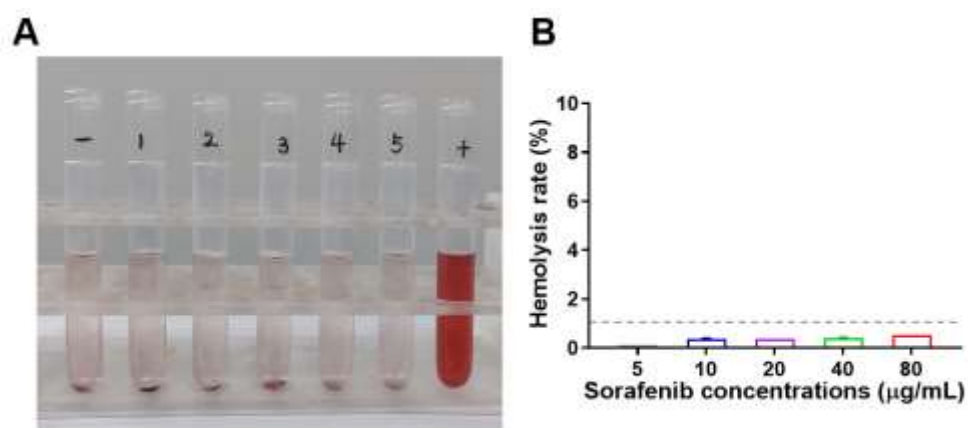
**Figure S5** (A) Phenotype analysis of adoptively transferred macrophage after injection with macrophage, USIP@M and USIP@M+L groups. (B) Phenotype analysis of endogenous TAM related to A. Data are presented as mean $\pm$ SD ( $n=3$ ). \*\*\* $P < 0.001$ .



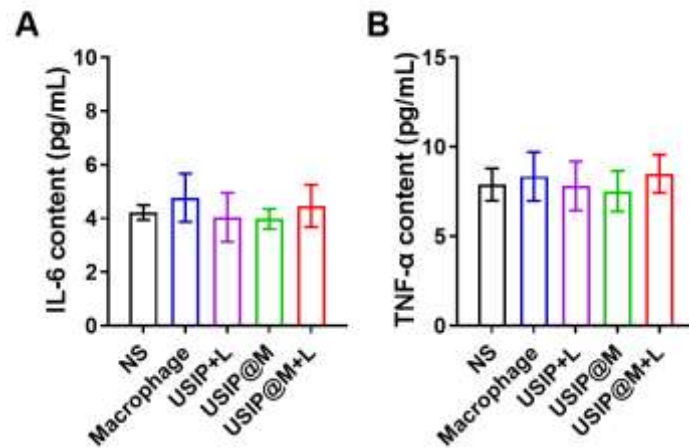
**Figure S6** Quantitative results for evaluating migration capacity of macrophage and USIP@M *in vitro*. Data are presented as mean $\pm$ SD ( $n=3$ ). \*\*\* $P < 0.001$ .



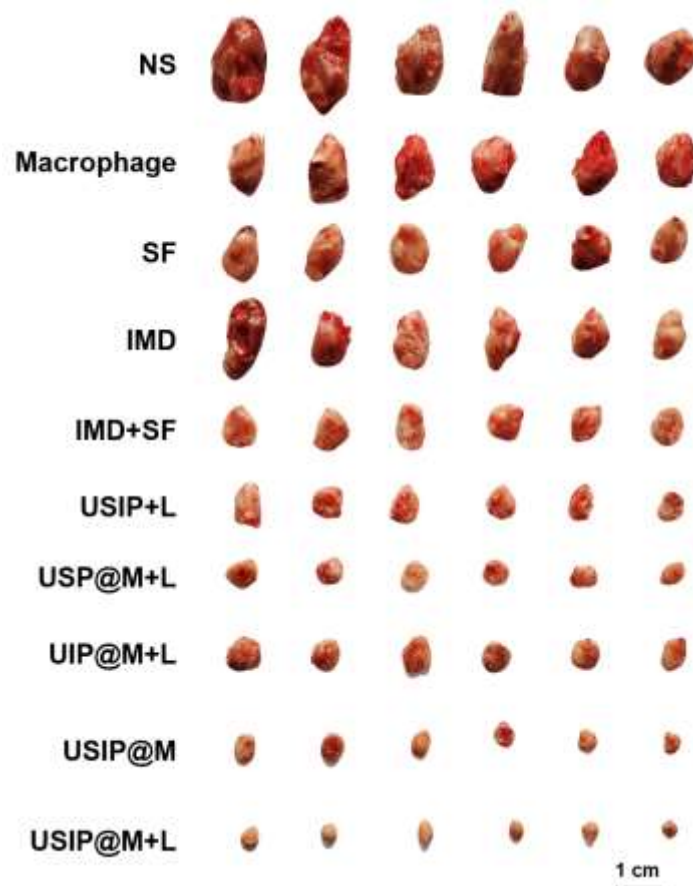
**Figure S7** H&E stained images of main organs (heart, liver, spleen, lung and kidney) after *i.v.* administration with different formulations, scale bar = 100  $\mu\text{m}$ .



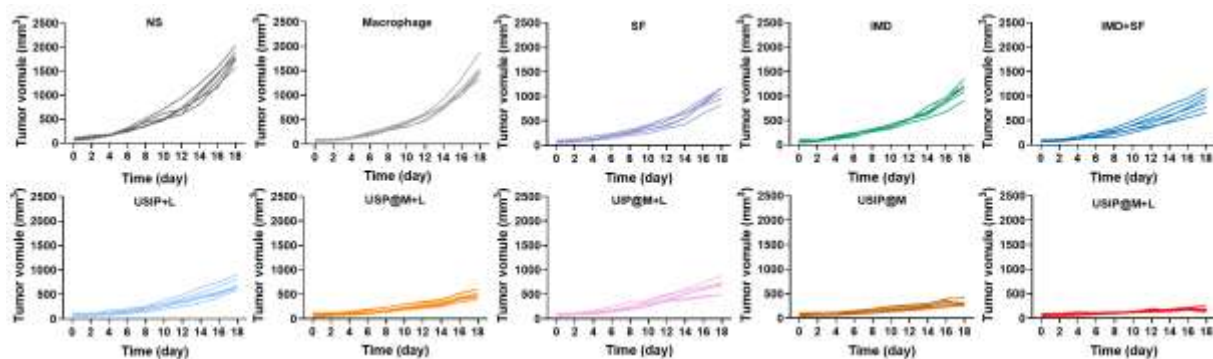
**Figure S8** (A) The hemolysis image of USIP@M. (B) The hemolysis rate with a series of doses of USIP@M at the concentration of SF in 5, 10, 20, 40 and 80  $\mu\text{g/mL}$ . Data are presented as mean $\pm$ SD ( $n=3$ ).



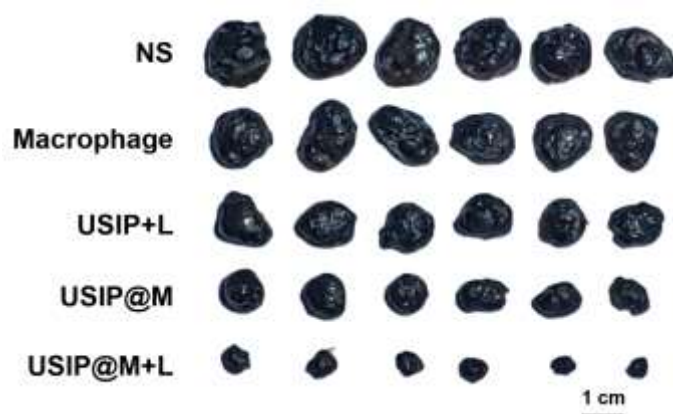
**Figure S9** (A) Serum cytokine concentrations of IL-6 assessed at 48 h after the first injection. (B) Serum cytokine concentrations of TNF- $\alpha$  assessed at 48 h after the first injection. Data are presented as mean $\pm$ SD ( $n=3$ ).



**Figure S10** Tumor photographs of H22 tumor-bearing mice receiving the indicated treatments related to Fig. 4 ( $n = 6$ ), scale bar = 1 cm.

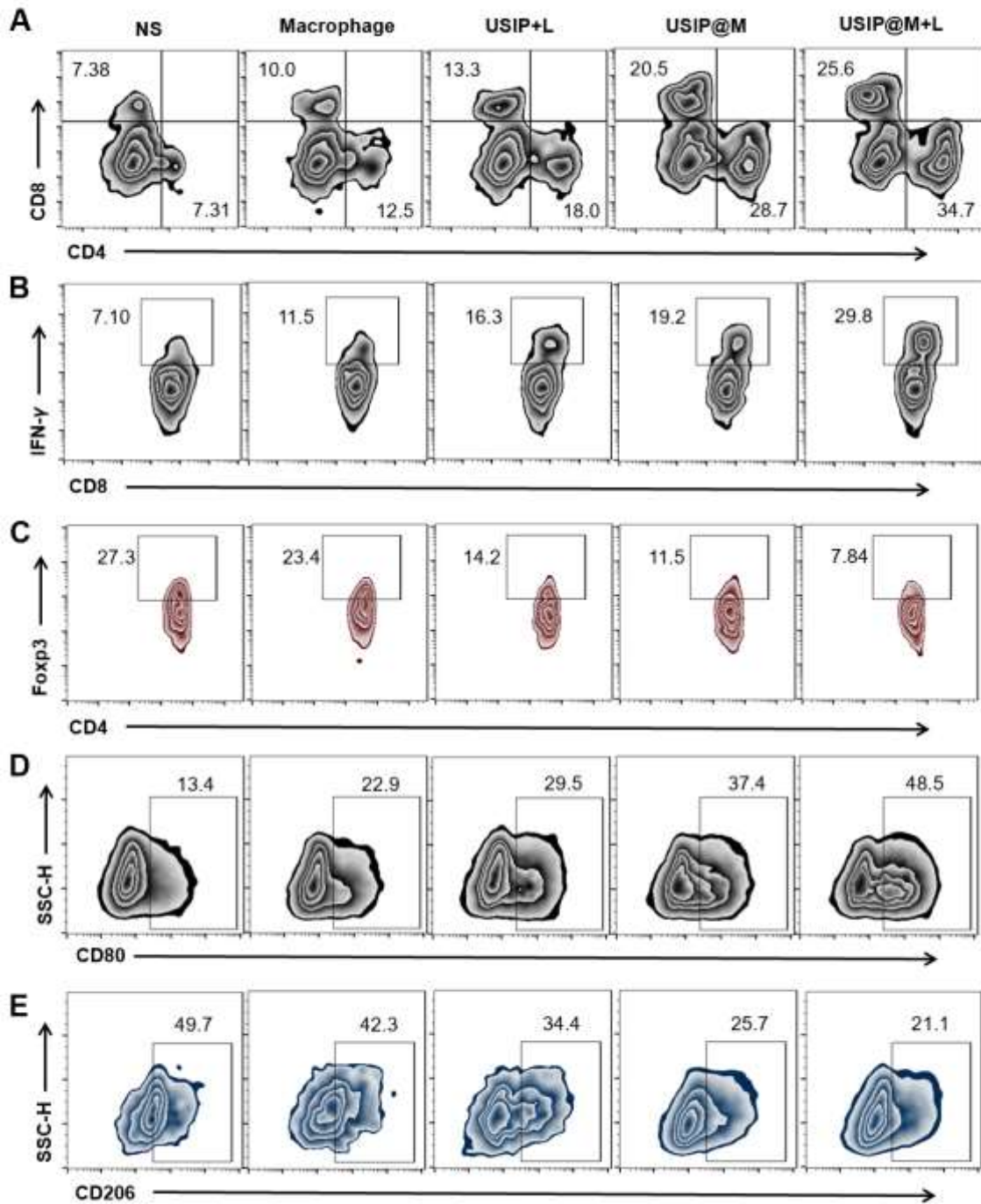


**Figure S11** Individual tumor growth curves of mice after *i.v.* administration with different formulations ( $n = 6$ ).



**Figure S12** Tumor photographs of B16F10 tumor-bearing mice receiving the indicated treatments related to Fig. 6 ( $n = 6$ ), scale bar = 1 cm.





**Figure S13** Representative flow cytometry plots of intratumoural (A) CD4<sup>+</sup> and CD8<sup>+</sup> T cells, (B) CD8<sup>+</sup>IFN- $\gamma$ <sup>+</sup> T cells, (C) Treg in CD4<sup>+</sup> T cells, (D) M1-type macrophages and (E) M2-type macrophages related to Fig 6F-L.



**Table S1.** Characterization of exosomes from different treatment groups

Group	Intensity size (nm)	Zeta potential (mV)
Macrophage	106.93 ± 3.39	-13.13 ± 1.07
USIP@M	117.87 ± 6.36	-13.00 ± 1.14
USIP@M+L	112.80 ± 3.35	-13.37 ± 1.42

**Table S2.** IC<sub>50</sub> in different treatment groups related to Fig. 2M

Group	Free SF	USIP	USIP+L	Released medium from USIP@M+L
IC <sub>50</sub> (µg/mL)	5.56 ± 2.22***	20.20 ± 8.01***	3.77 ± 0.47***	0.76 ± 0.34

\**P* indicate statistical significance compared to the released medium from USIP@M+L group.

Data are presented as mean±SD (*n* = 3).\*\*\**P* < 0.001.