

Sono-photodynamic Therapy on Cancer and Bacteria

Literature Seminar

2023/9/14

Keiichi Kawabata (M2)

Contents

- Introduction
- Sono-photodynamic Therapy on Cancer
- Sono-photodynamic Therapy against Bacteria
- Summary

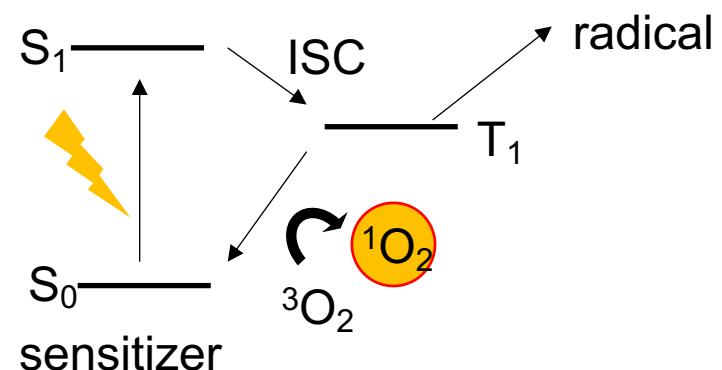
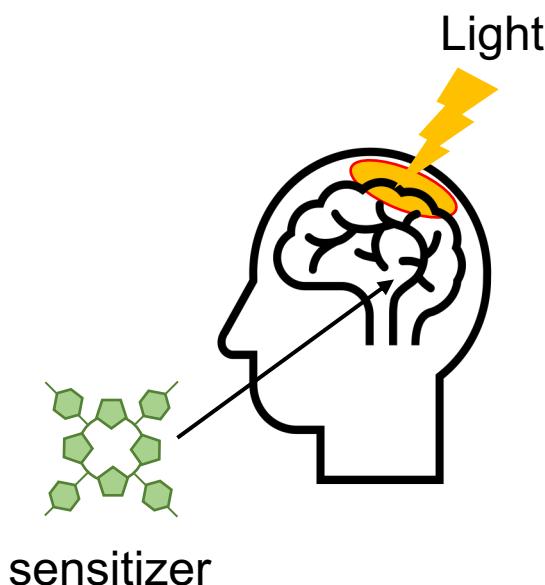
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Photodynamic Therapy

PDT

- ✓ High selectivity
- ✓ Low risk of side effects
- ✗ Low permeability



Sonodynamic Therapy

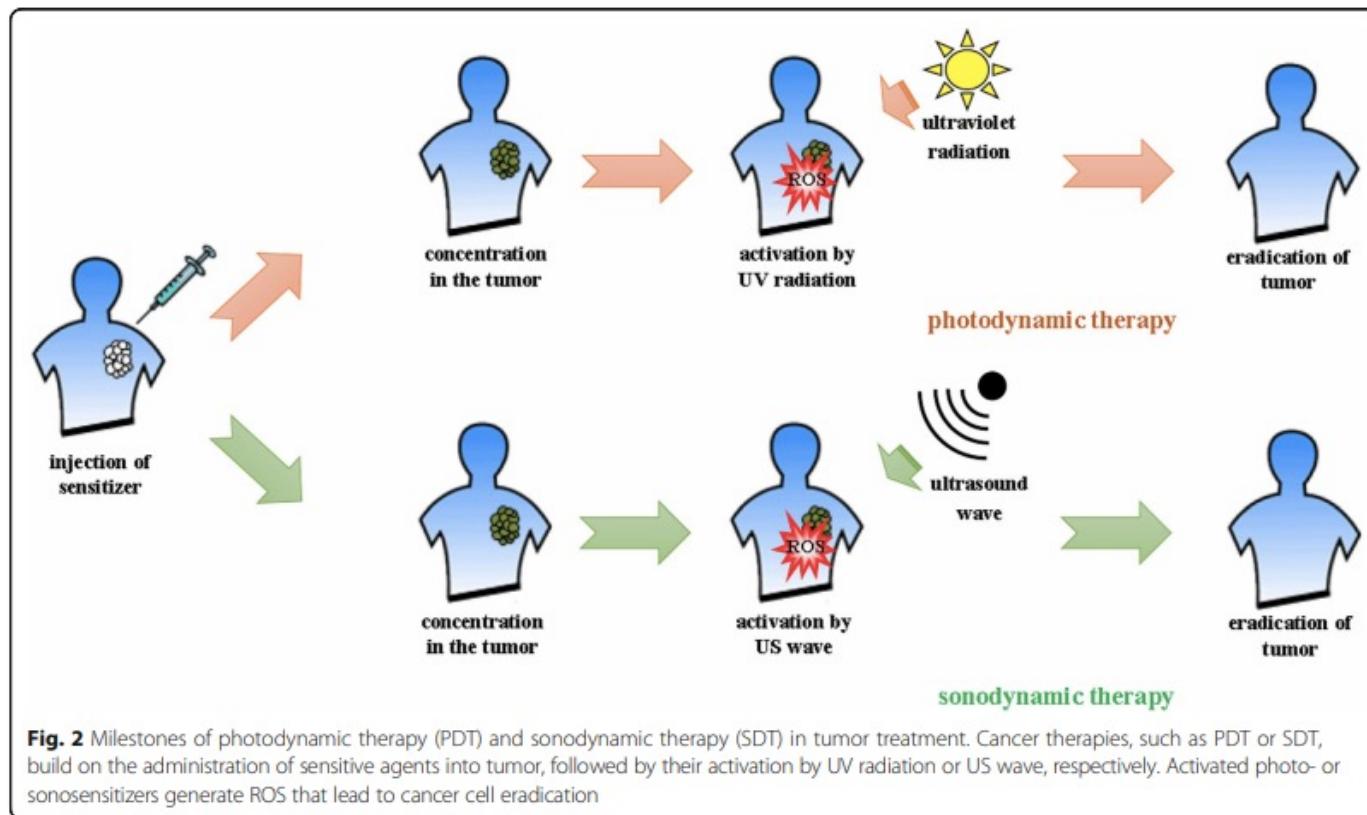


Fig. 2 Milestones of photodynamic therapy (PDT) and sonodynamic therapy (SDT) in tumor treatment. Cancer therapies, such as PDT or SDT, build on the administration of sensitive agents into tumor, followed by their activation by UV radiation or US wave, respectively. Activated photo- or sonosensitizers generate ROS that lead to cancer cell eradication

SDT

Bogdan, et al., *Nanoscale Res. Lett.*, 2017, 12, 225

- ✓ Non-invasive
- ✓ High permeability

Cavitation

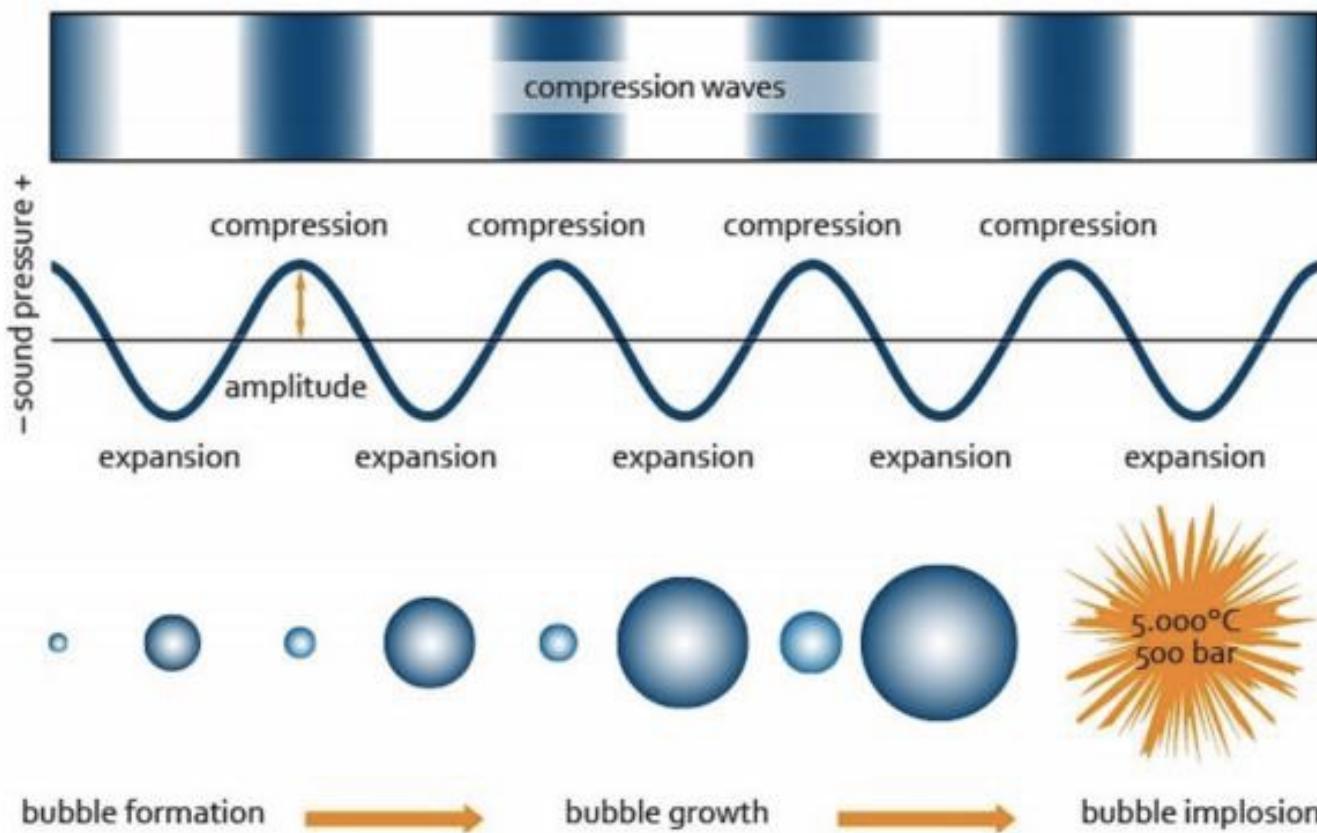
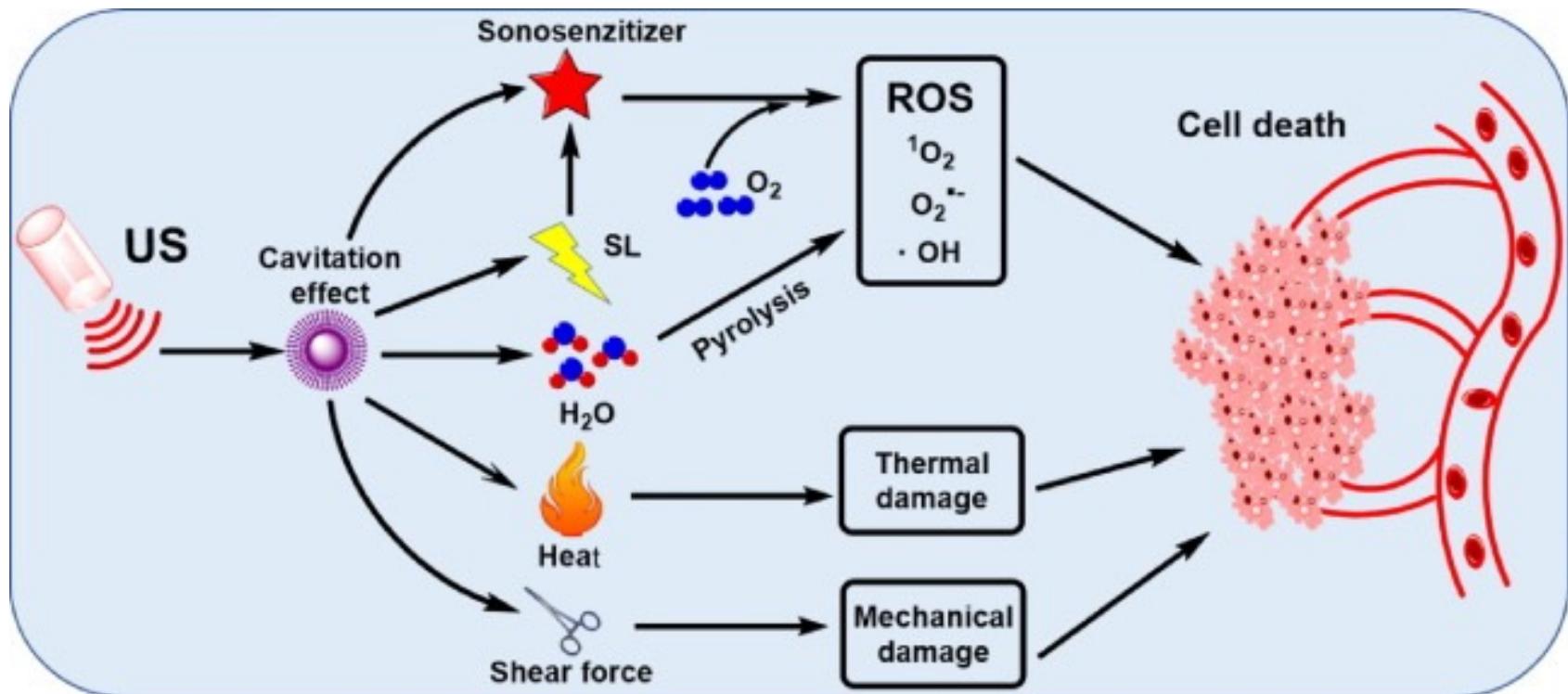


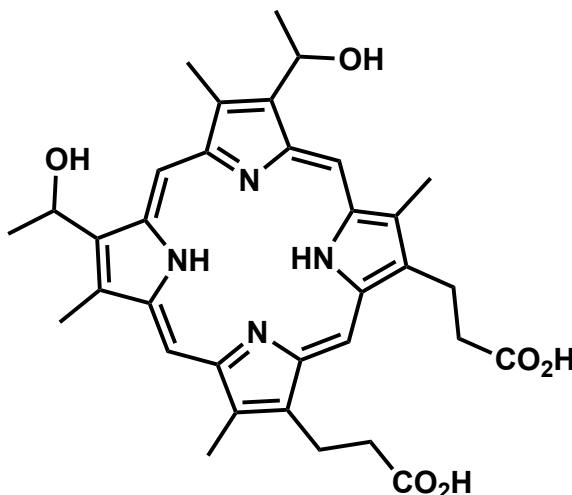
Figure 1. Principle of ultrasound cavitation [16]. The initiated bubbles grow due to evaporation and finally reach critical size (resonant) when it grows quickly and collapse violently.

Mechanism of SDT

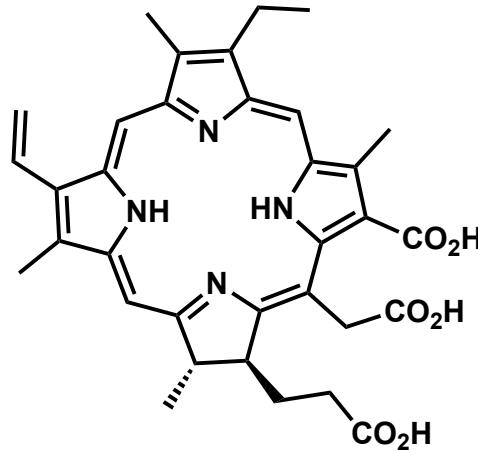


Xing, X. et al., *Coordination Chemistry Reviews*, 2021, 445, 214087

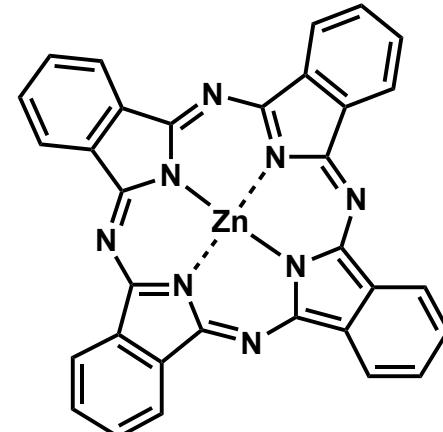
Sensitizers



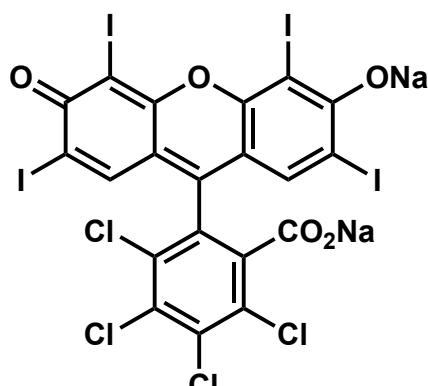
Hematoporphyrin



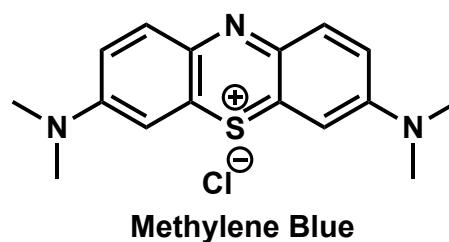
Chlorin e6



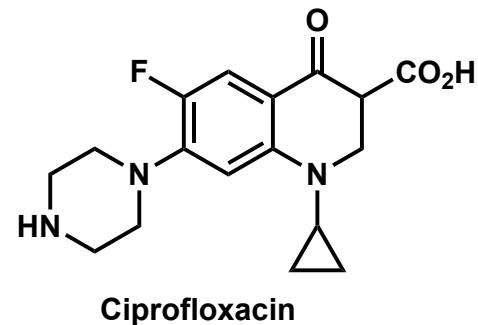
Zinc Phthalocyanine



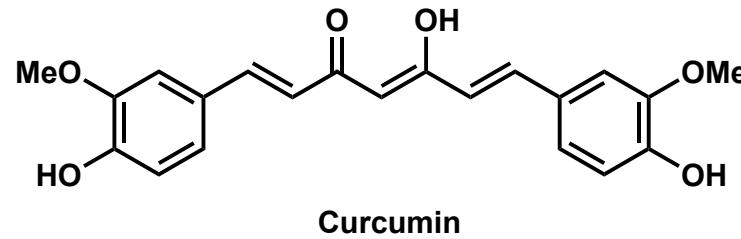
Rose Bengal



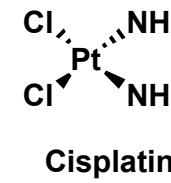
Methylene Blue



Ciprofloxacin



Curcumin

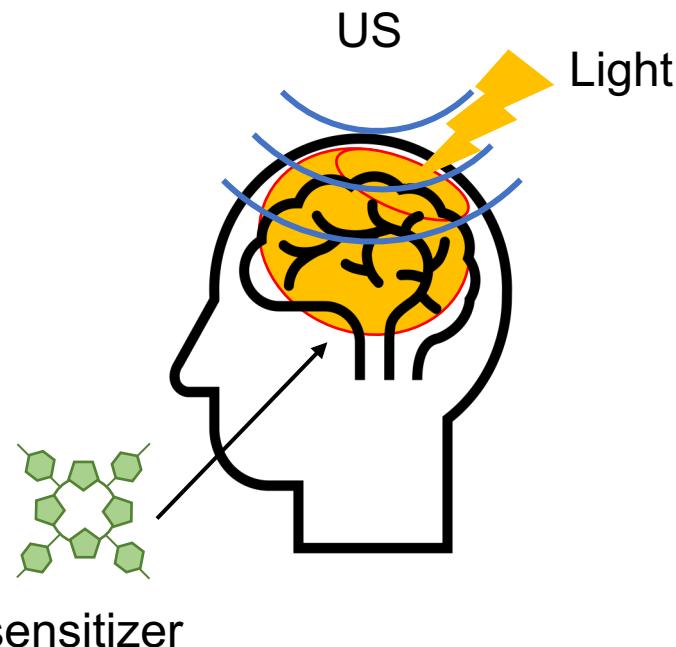


Cisplatin 8

Sono-photodynamic Therapy

SPDT

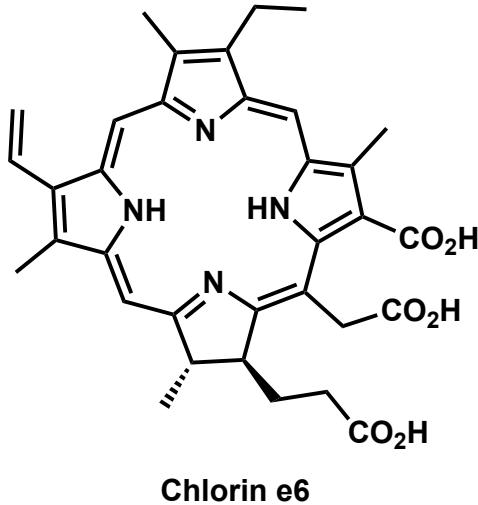
- ✓ High selectivity
- ✓ Low risk of side effects
- ✓ Low permeability
- ✓ High efficiency



Contents

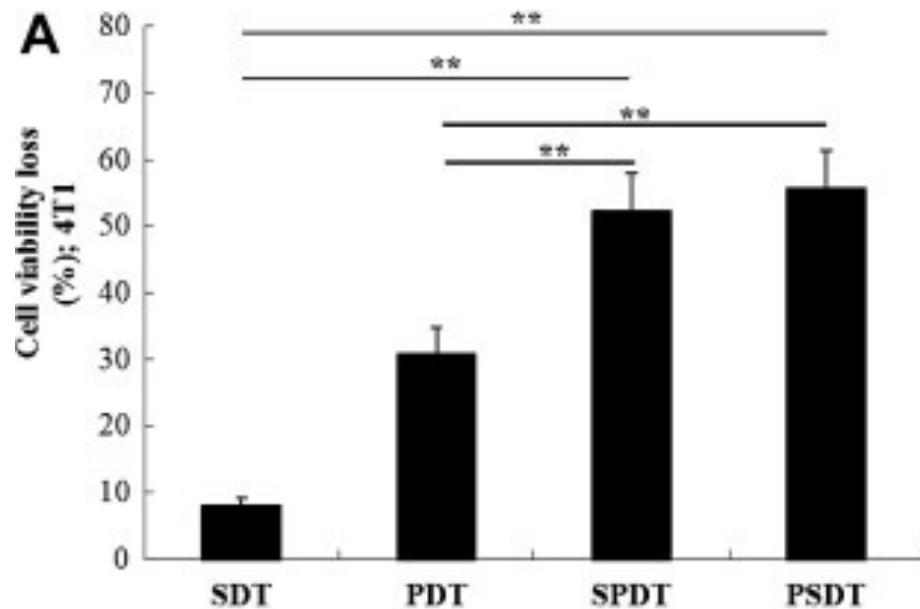
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- Sono-photodynamic Therapy against Bacteria
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Cytotoxicity of Ce6

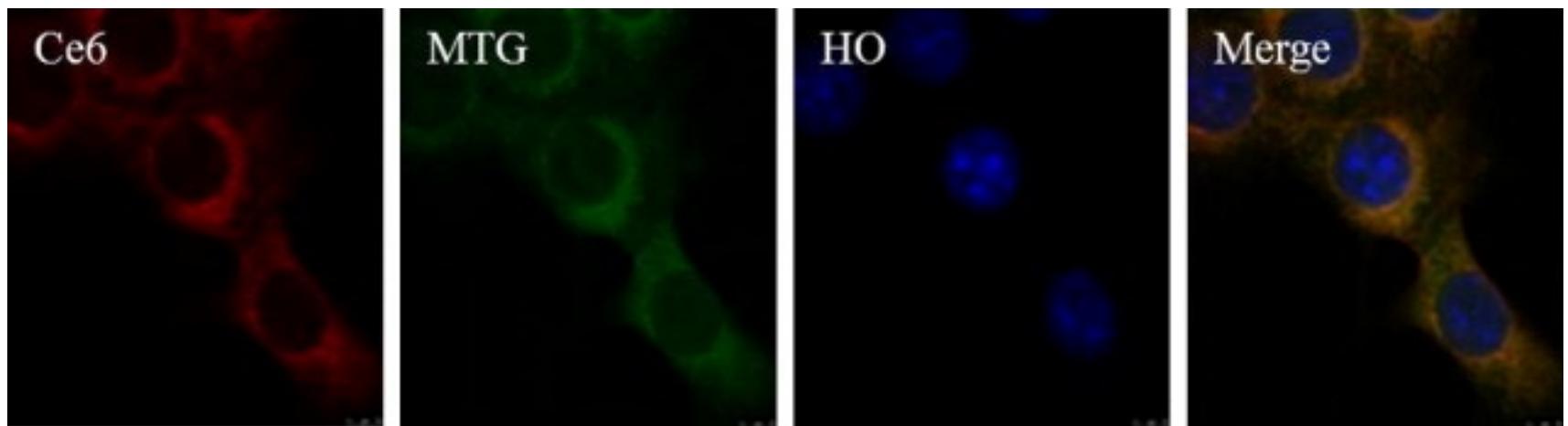


Light: 650 nm, 1.2 J/cm²
US: 0.36 W/cm²

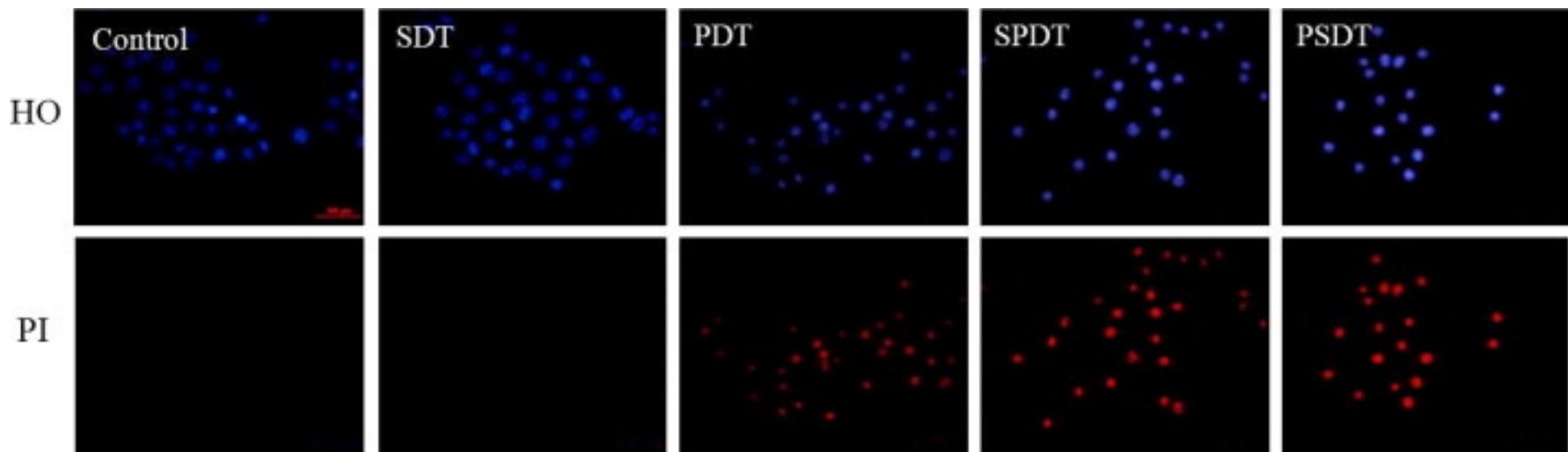
PDT: light only
SDT: US only
SPDT: light → US
PSDT: US → light



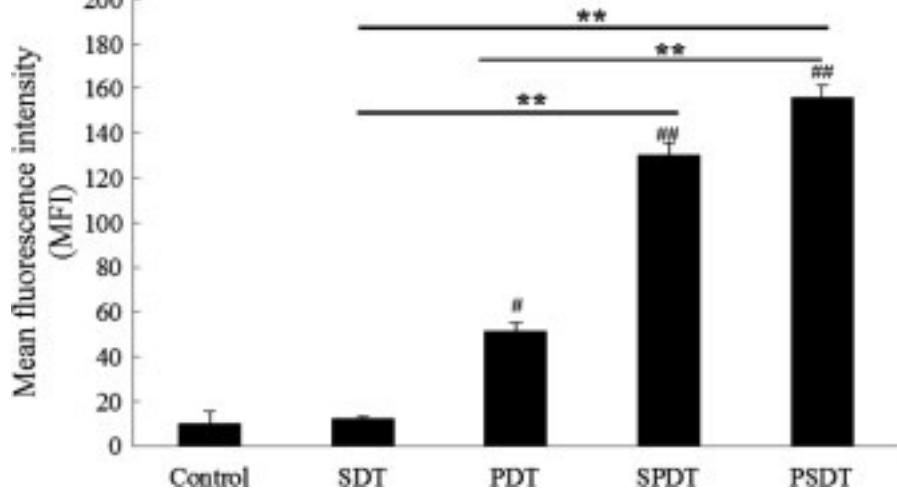
Localization



Morphological Analysis

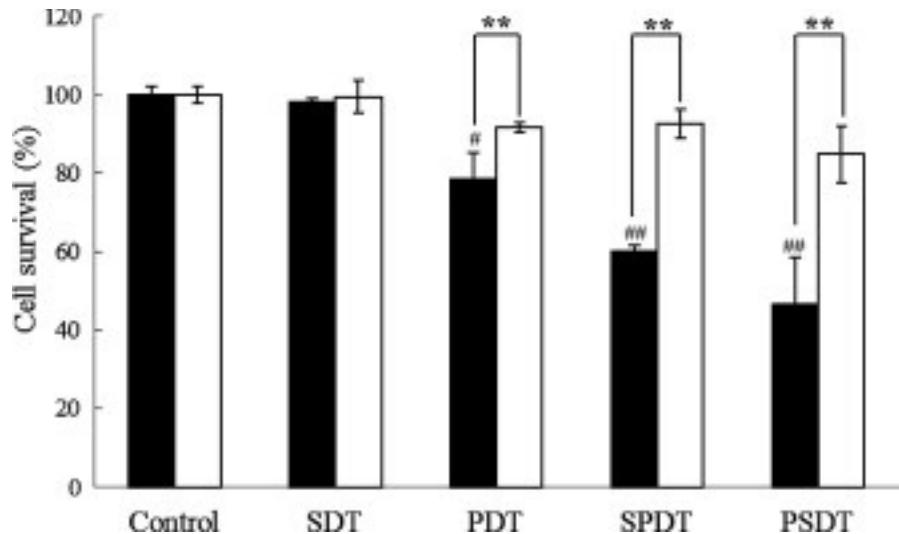


ROS Detection

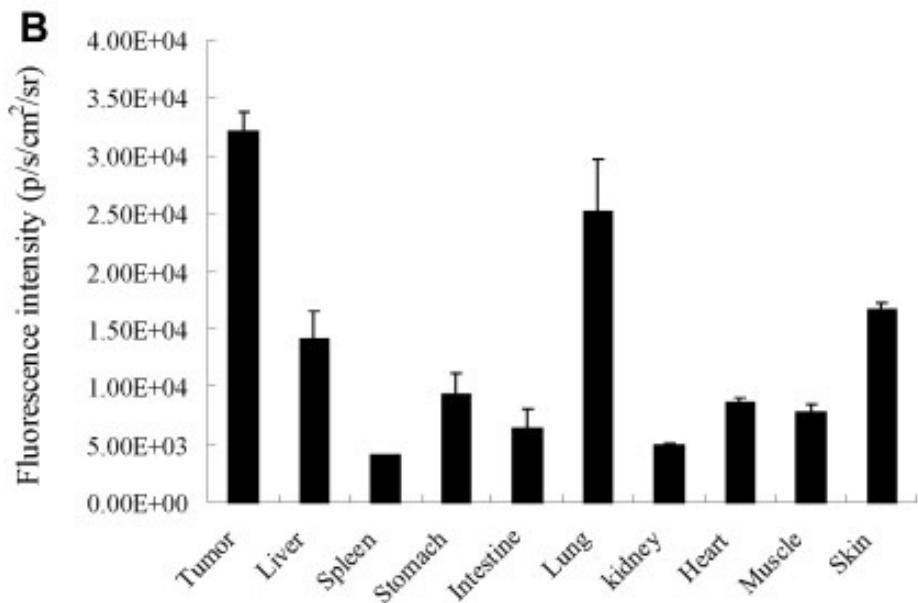
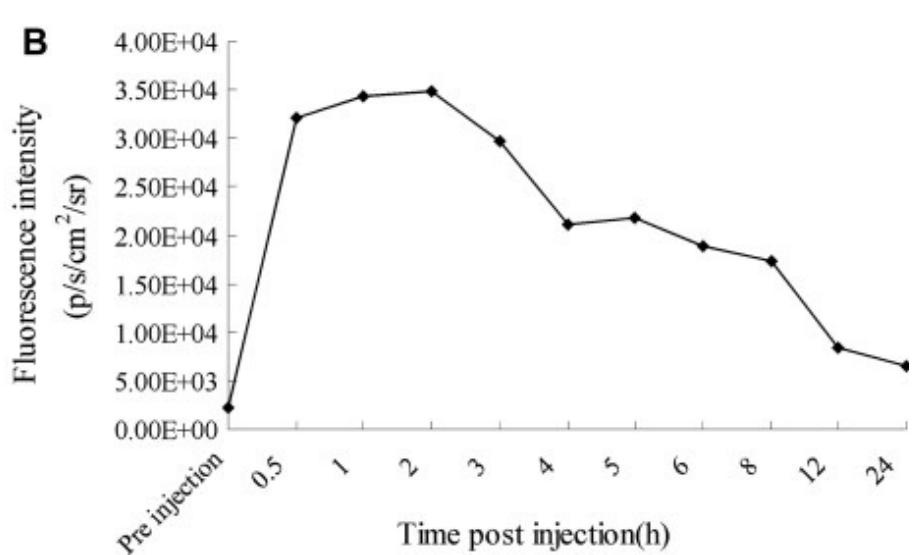


Rescue experiment

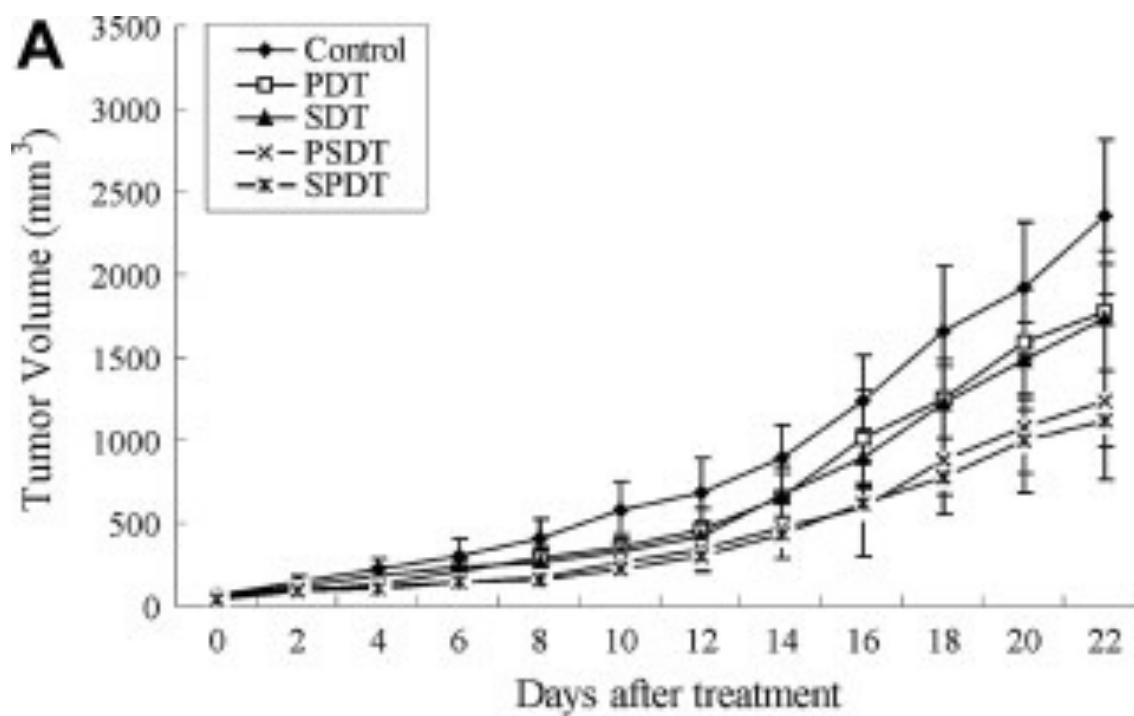
Black: NAC-
White: NAC+



Distribution of Ce6

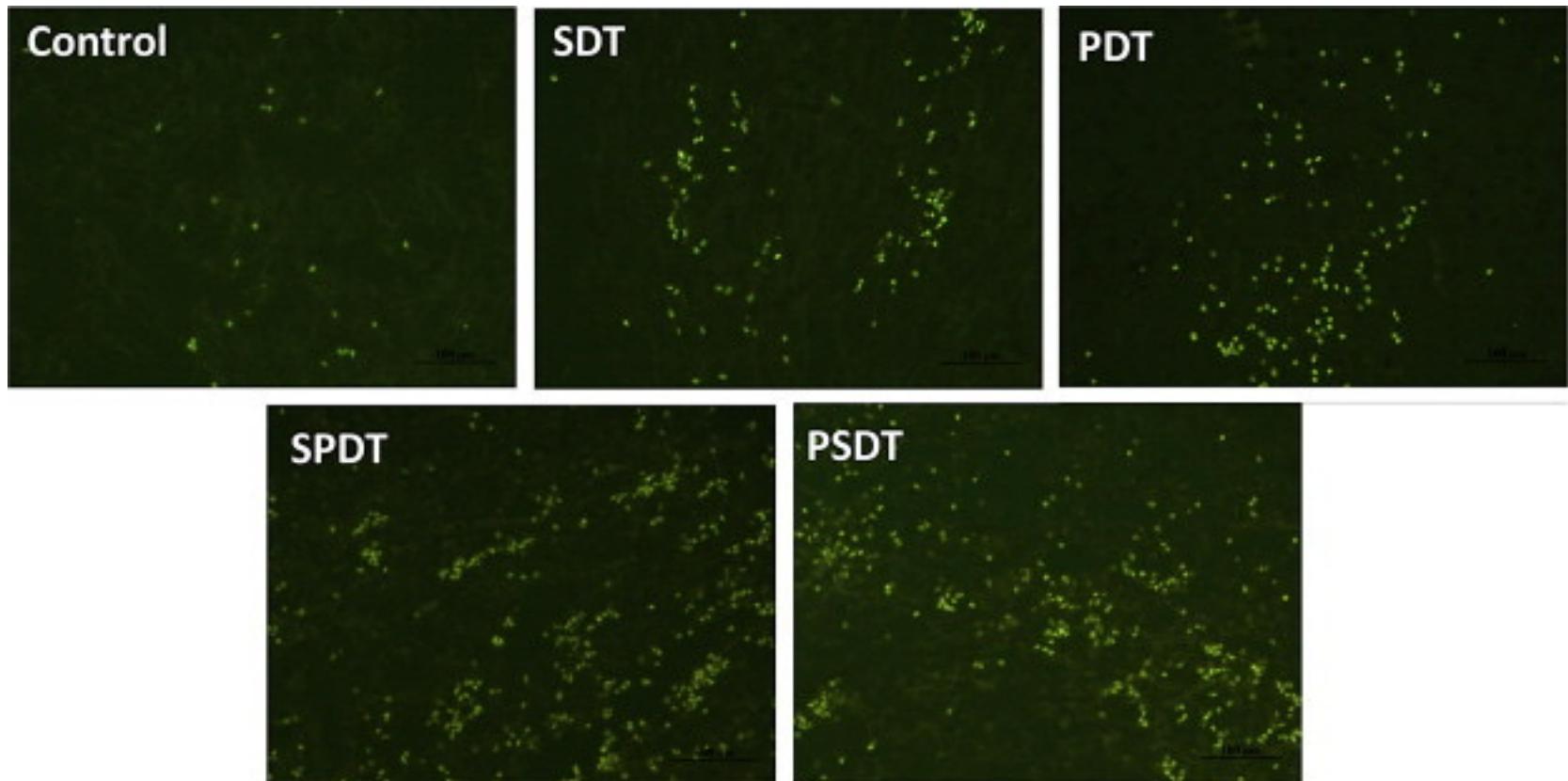


Inhibition of Tumor Growth



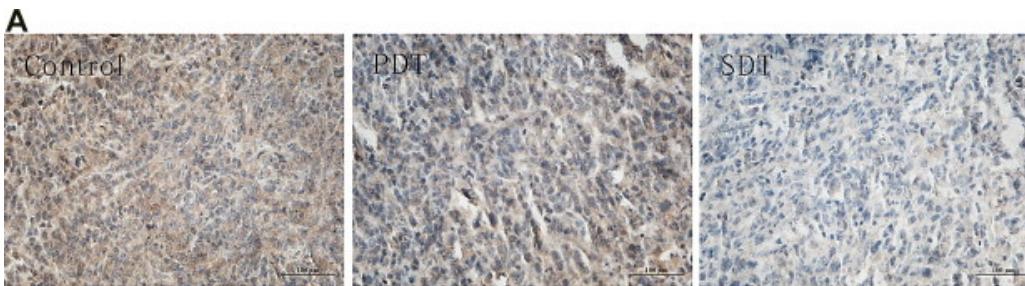
Apoptosis In Vivo

TUNEL assay

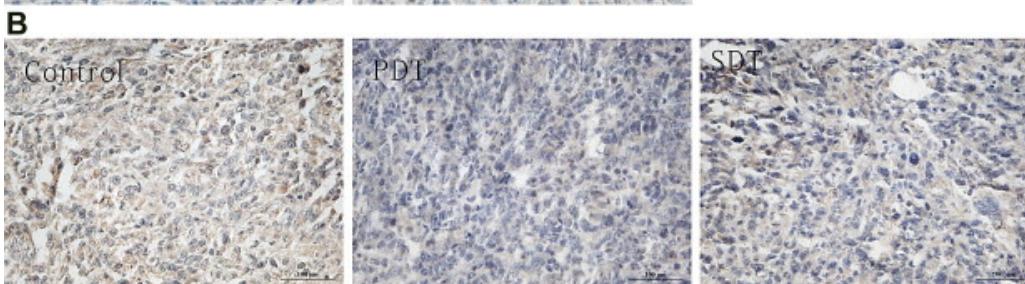


Decrease of VEGF & MMP-9

IHC assay



VEGF:
血管新生促進遺伝子



MMP-9:
がん転移促進遺伝子

Short Summary

- SPDT is a promising cancer therapy combining PDT and SDT.
- SPDT is more effective than PDT or SDT alone.
- ROS damage → mitochondria → apoptosis
- Decrease in expression level of VEGF and MMP-9

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Antibacterial Strategies

- Antibiotics
 - ✗ Drug resistance

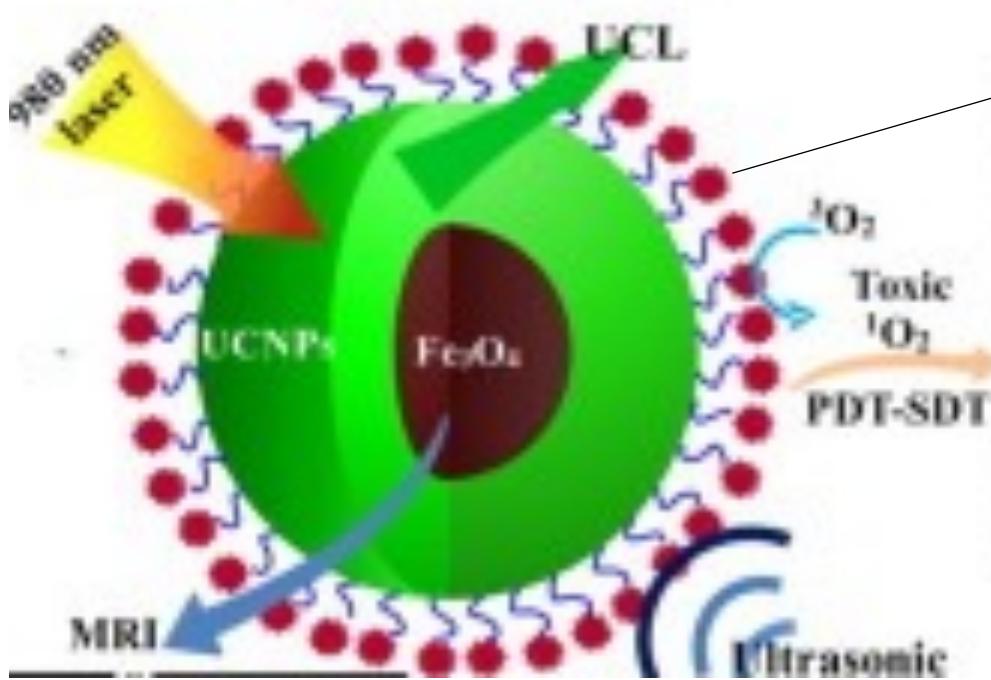


- PDT
 - ✓ High selectivity
 - ✓ Low risk of side effects
 - ✓ Low potential of drug resistance

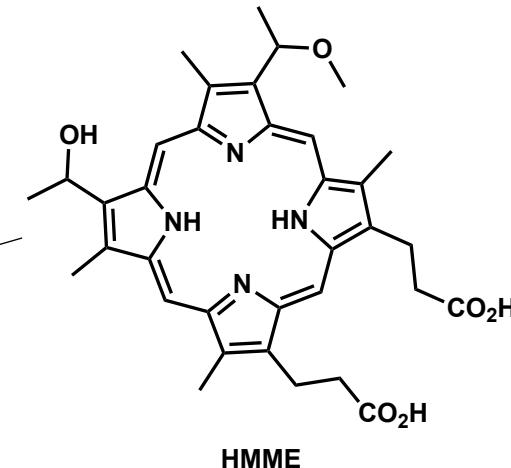


- SDT/SPDT
 - ✓ Permeability

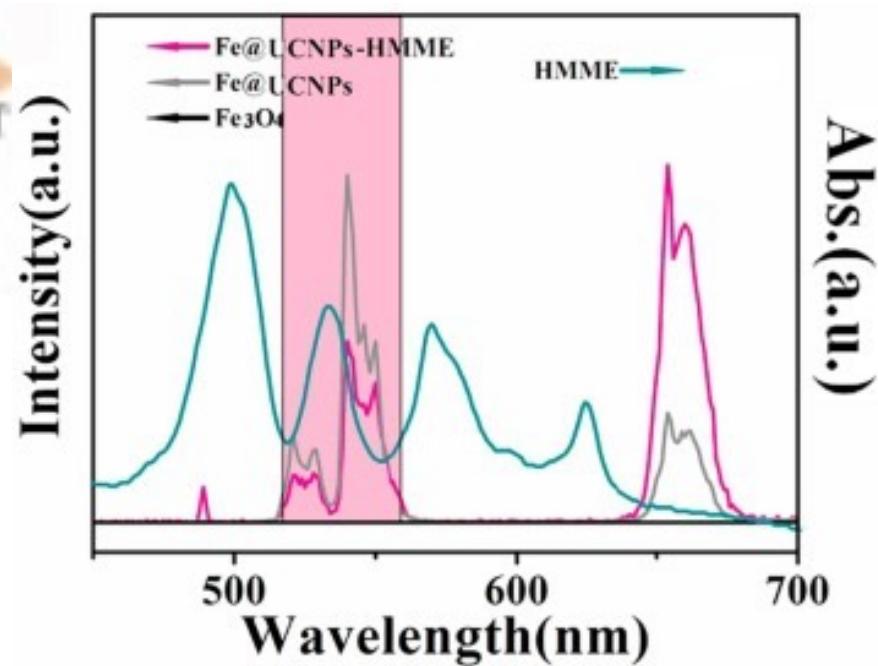
Nanoparticle



Fe@UCNP-HMME

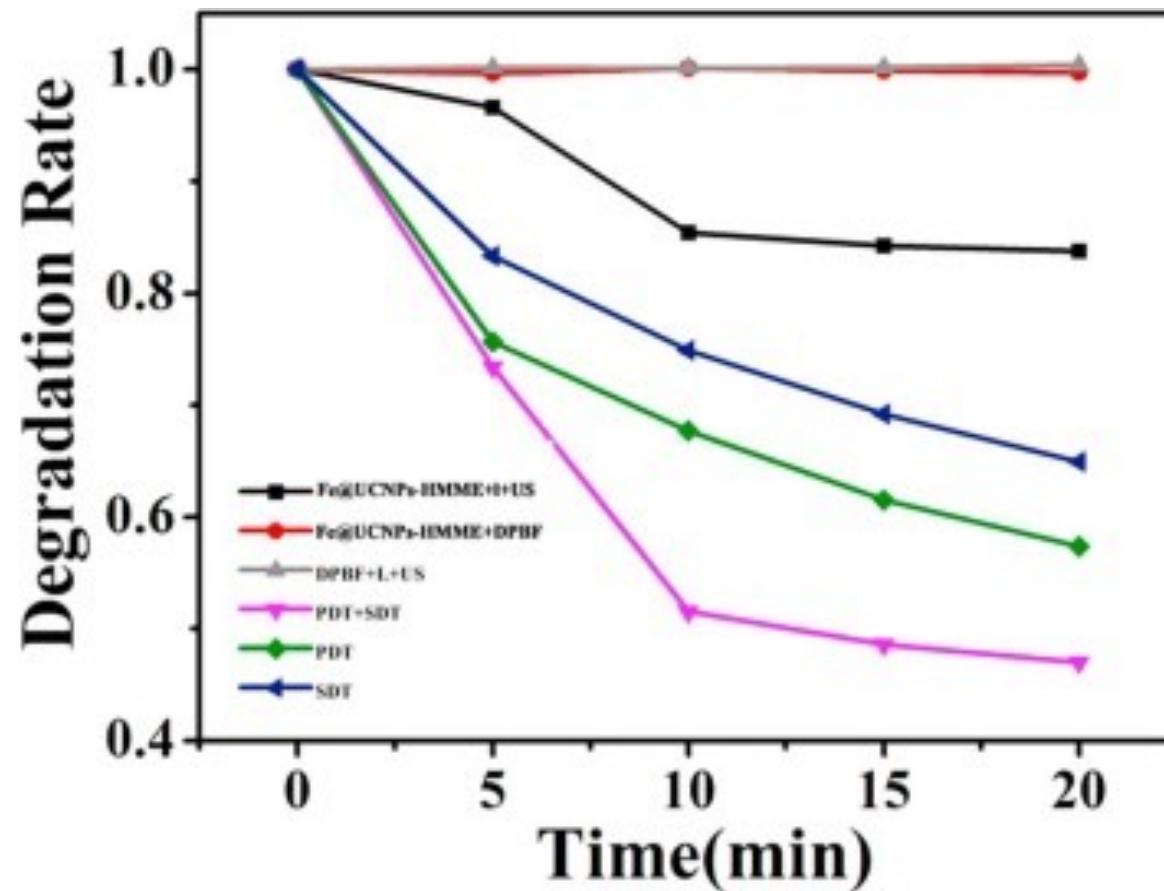


HMME

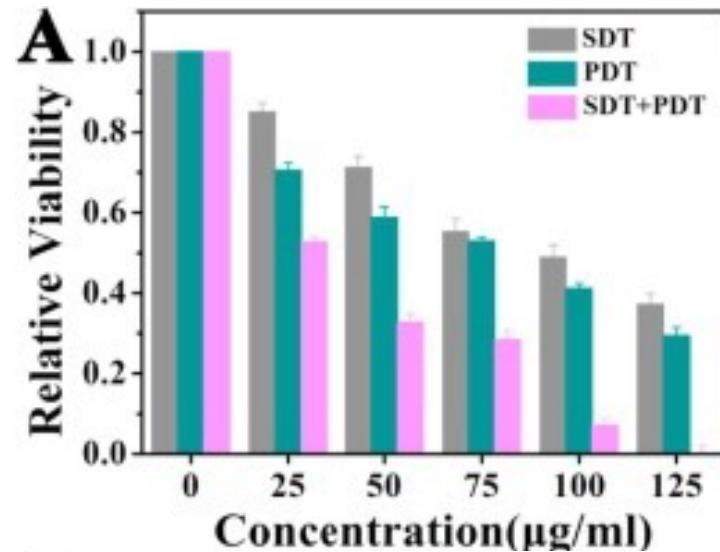


$^1\text{O}_2$ Generation

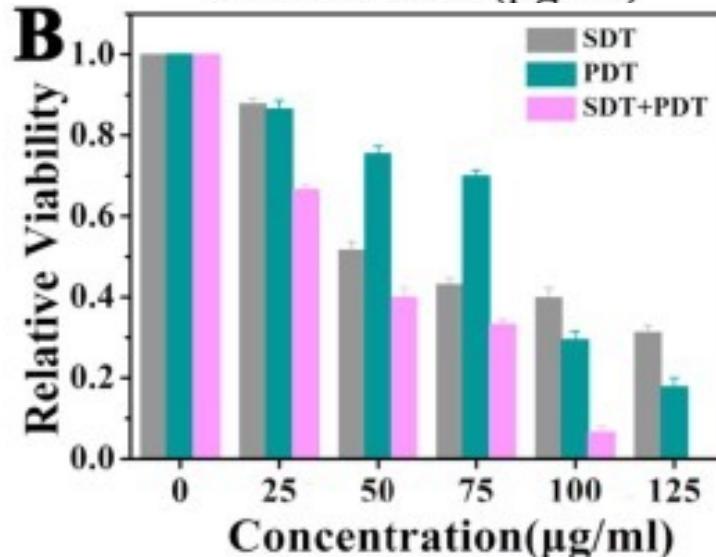
Light: 980 nm, 1 W/cm²
US: 2 W/cm²



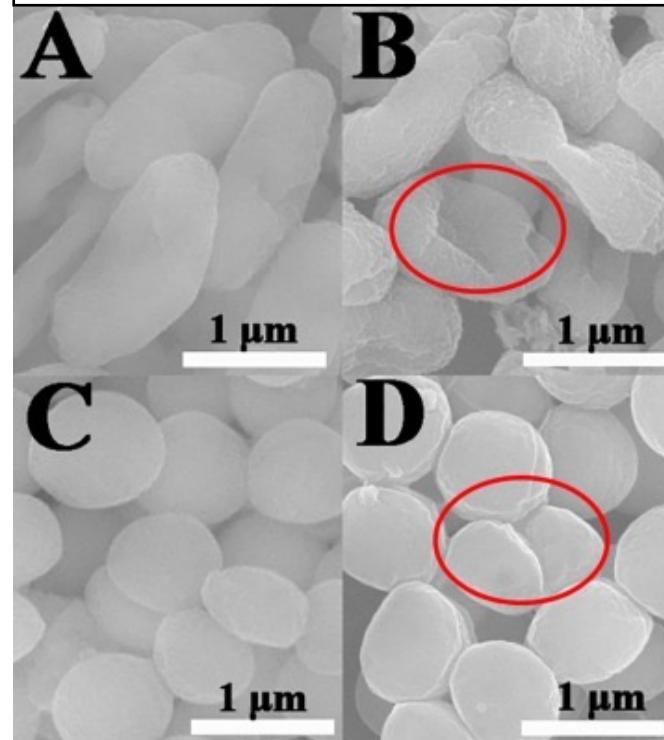
Bacterial Viability



A: ESBL-producing *E. coli*
B: MRSA



A, B: ESBL-producing *E. coli*
C, D: MRSA



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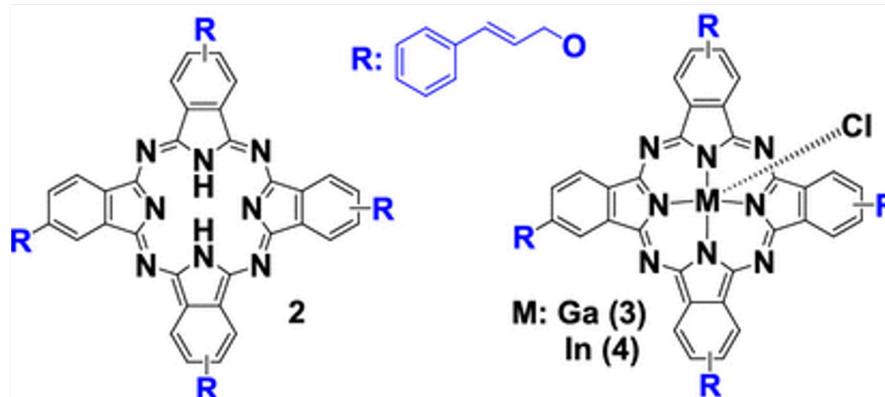
Summary

- SPDT is applied to cancer and bacteria treatment.
- SPDT is more effective than PDT and SDT.
- Mechanism of SPDT remains unclear.

APPENDIX

$^1\text{O}_2$ Quantum Yield

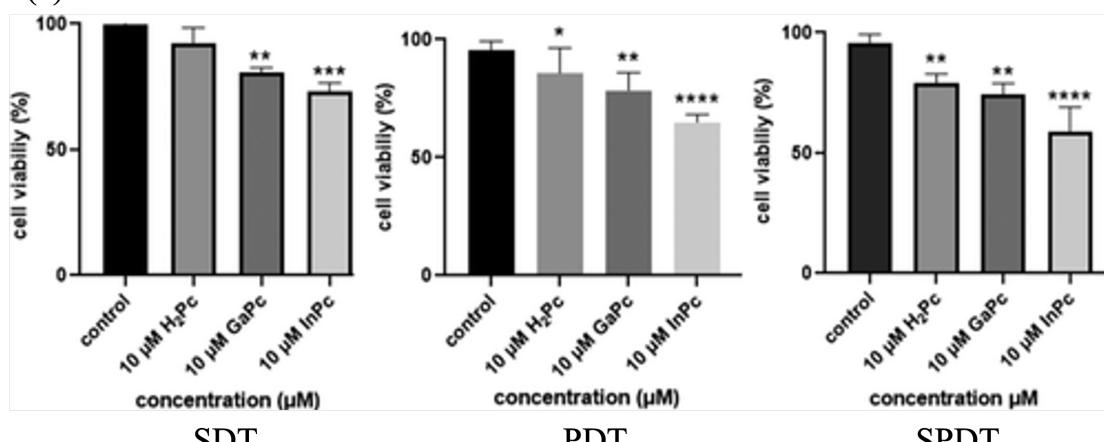
(a)



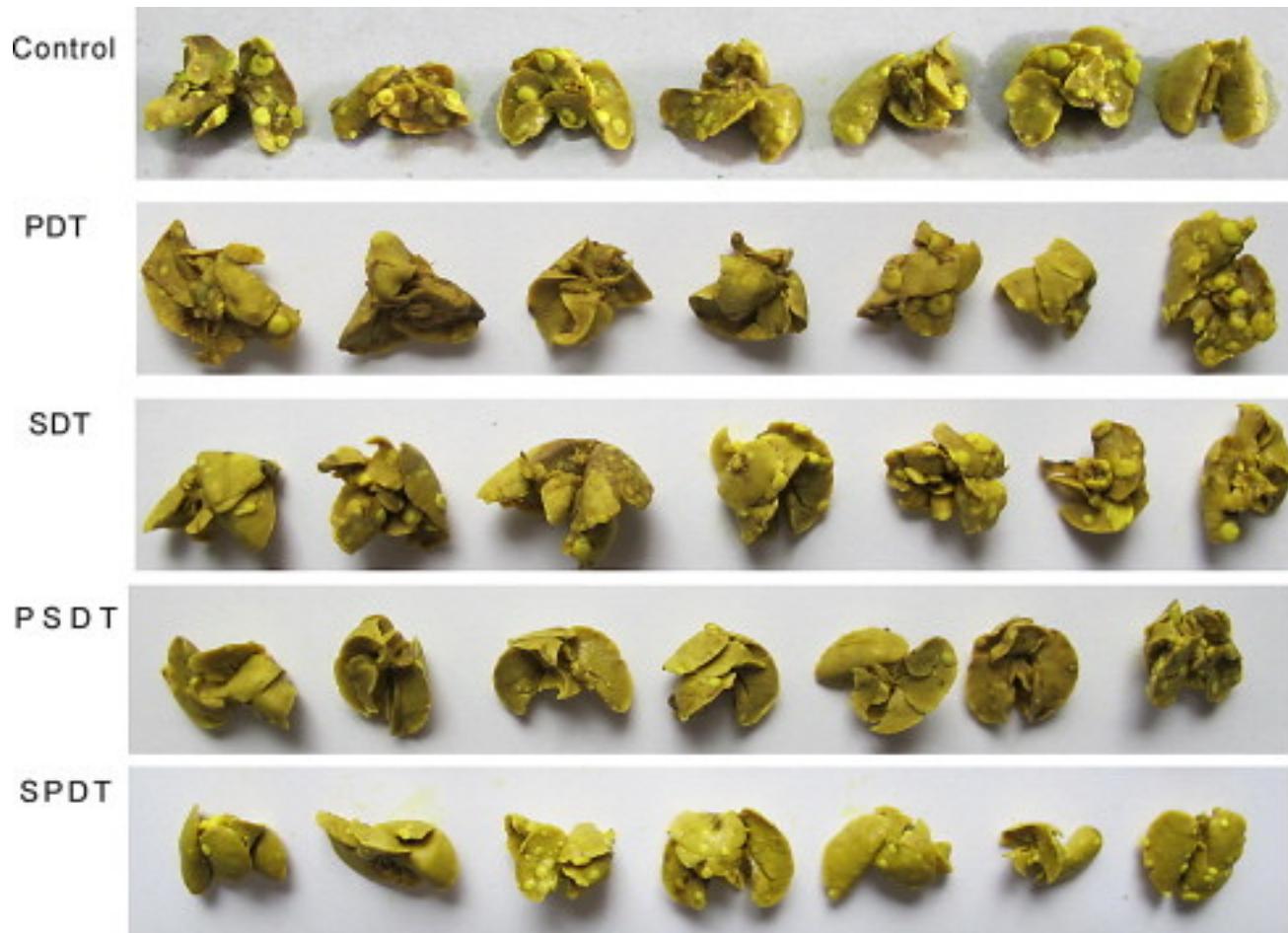
(b)

Pcs	$\Phi_F (10^{-2})$	$\Phi_{\Delta}(\text{PDT})$	$\Phi_{\Delta}(\text{SPDT})$
H ₂ Pc (2)	8.60	0.22	0.55
GaPc (3)	0.13	0.61	0.85
InPc (4)	0.27	0.78	0.96

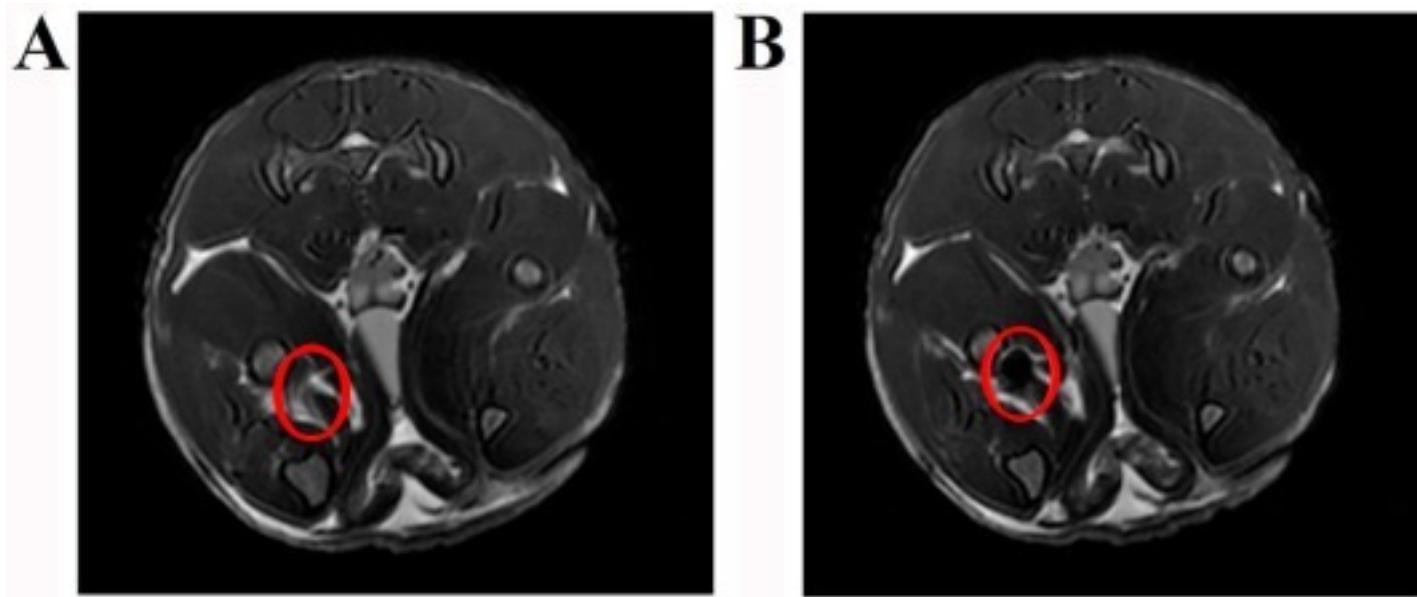
(c)



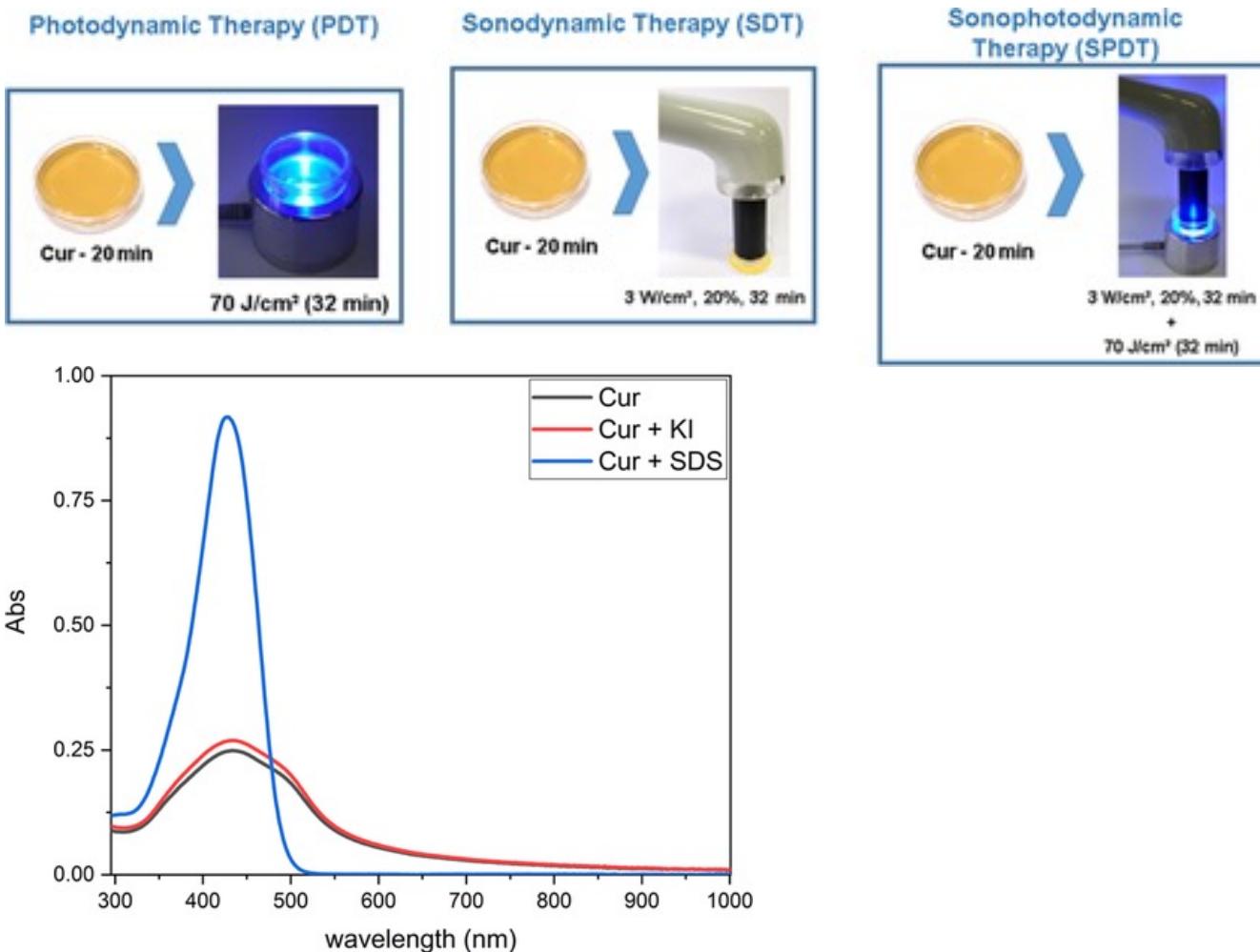
Metastasis to Lung



Imaging



Biofilm



Biofilm

