# Literature Report

## Reporter: Wang Guangying Date: 2020-12-3

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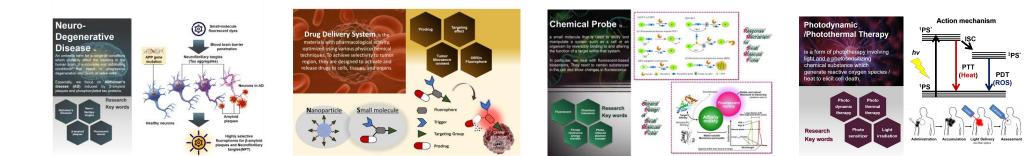
#### **RESEARCH ARTICLE**

### An Ethacrynic Acid-Brominated BODIPY Photosensitizer (EA-BPS) Construct Enhances the Lethality of Reactive Oxygen Species in Hypoxic Tumor-Targeted Photodynamic Therapy

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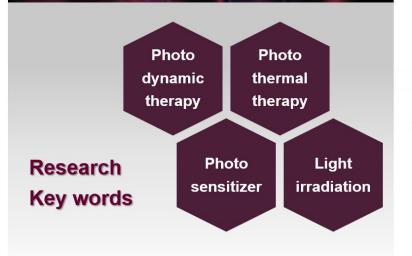


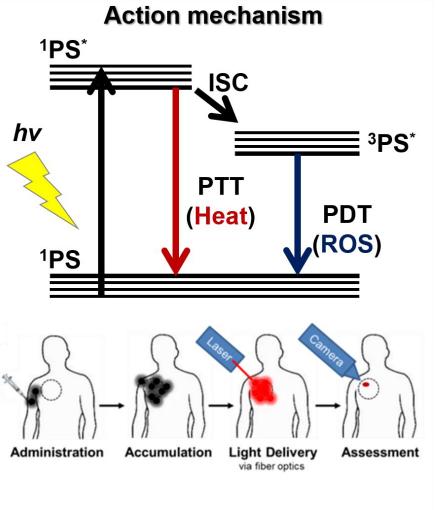
*Jong Seung Kim* Korea University Neurodegenerative disease, Drug delivery system, Chemical probe, Photodynamic / thermal therapy.

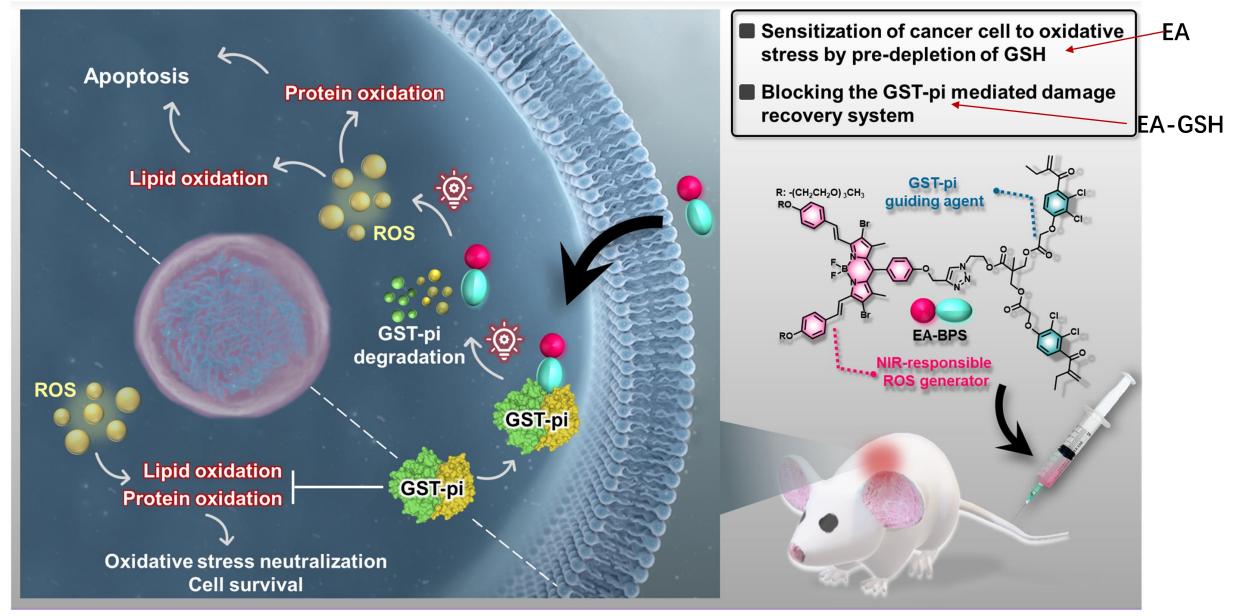


#### Photodynamic /Photothermal Therapy

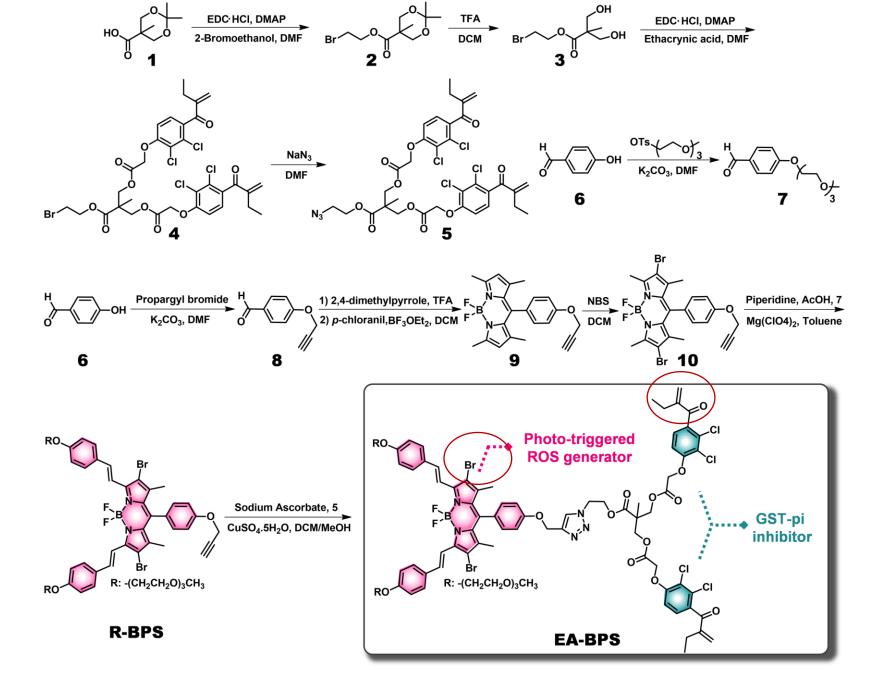
is a form of phototherapy involving light and a photosensitizing chemical substance which generate reactive oxygen species / heat to elicit cell death.







1.EA-BP与PDT发挥协同作用; 2.EA-BP降低了GST-pi活性,并增强了脂质过氧化作用; 3.EA-BPS在缺氧条件下还可以改善基于PDT的细胞毒性。



Scheme 1. Synthesis of R-BPS and EA-BPS.

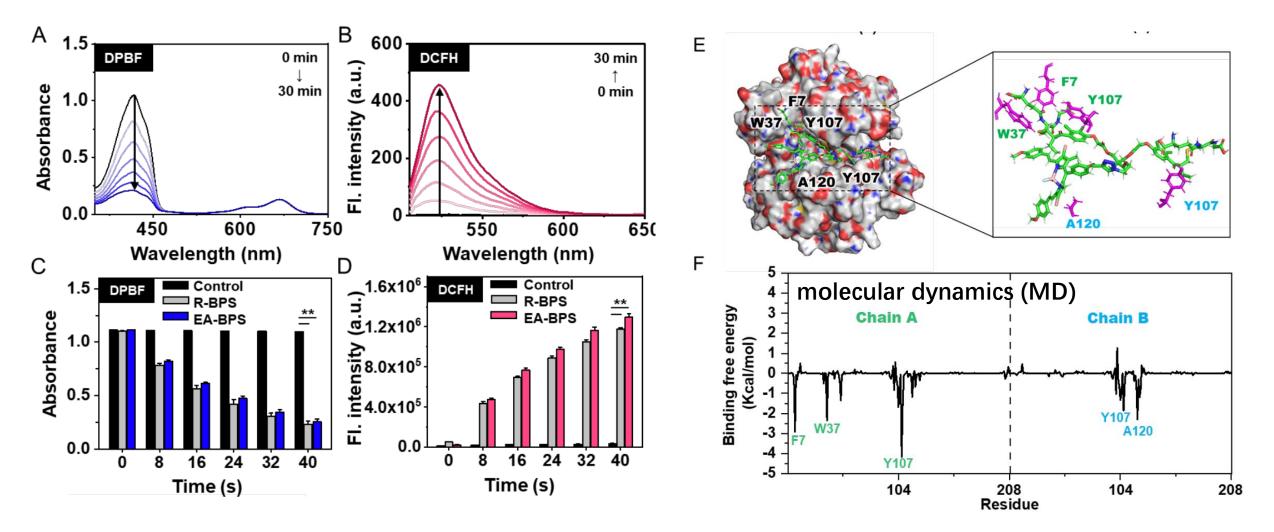


Figure 2. Photodynamic and GST-pi binding properties of EA-BPS.

Fig a : a mixture of EA-BPS and DPBF

Y107 residues are present in the active site

- DPBF : chemical probe for singlet oxygen (Type 2 ROS)
- DCFH ROS sensor : primarily for Type 1 ROS

Conclusion: R-BPS and EA-BPS are capable of promoting ROS production

- Fig 3A : In MDA-MB-231 cell line, EA-BPS was taken up more effectively than R-BPS and time dependent .(lipophilic and interactions)
- Fig3b:NAC, an effective ROS quencher; ROS probe (DCFH-DA, green).
- Fig3c: ascertain which of these mechanisms Singlet Oxygen Sensor Green (SOSG)
- Fig3d: a Type I mechanism, dihydroethidiu (DHE) , detection of O2<sup>-</sup> –  $\widehat{}$
- Fig3e: HPF, detection of hydroxyl radical E and peroxynitrite anions
- Fig3f: monitored the levels of GSH expression using a thiol-tracker

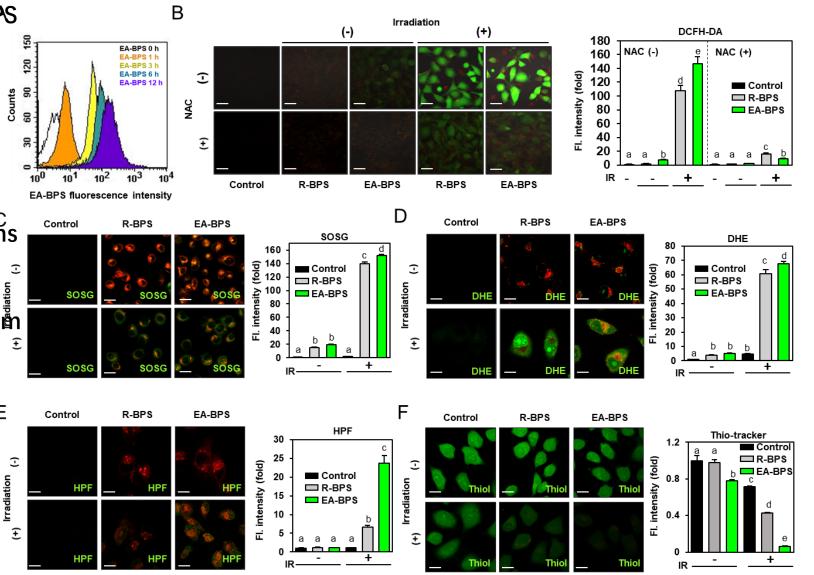
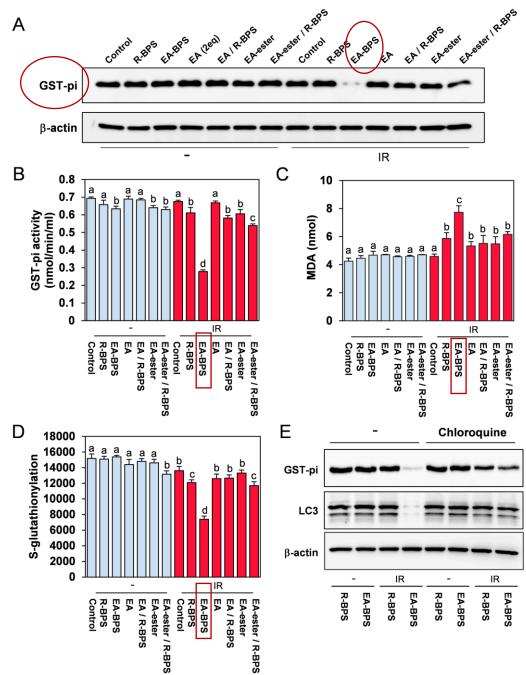
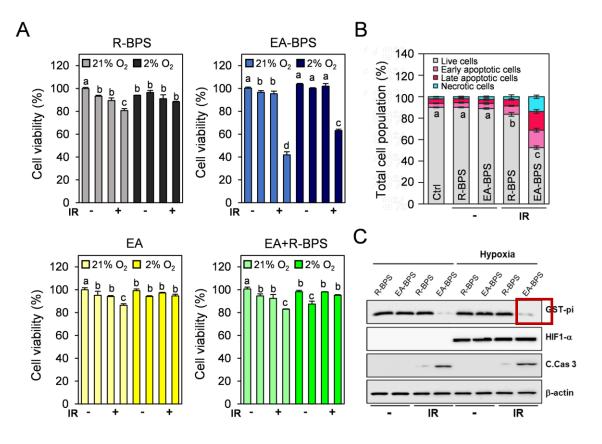


Figure 3. PDT sensitizer potential of EA-BPS.





*Figure 5.* Hypoxia induced apoptotic cell death by **EA-BPS**.

Fig4c: MDA, one of the final products of lipid peroxidation Fig4d: GST-pi can promote the Sglutathionylation of damaged proteins

Fig4e:(CQ) (a lysosomal inhibitor), LC3 (an autophagy marker) protein

*Figure 4.* Effect of **EA-BPS** with and without photoirradiation on lysosomal degradation-dependent way

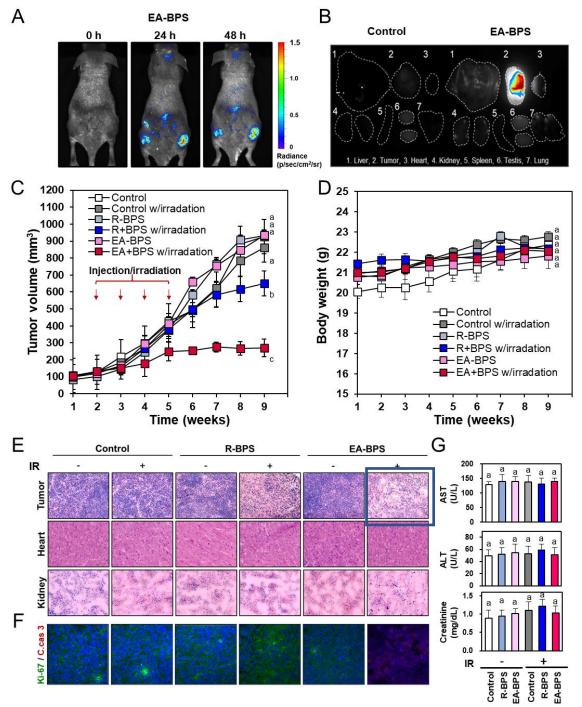


Fig 6c: at the tumor regions showed a statistically significant reduction in tumor growth and volumes

Fig 6e: characteristic apoptotic cells

Fig 6f: caspase-3 (red) , Ki-67 expression levels (green) :cell proliferation

Fig6g AST(天门冬氨酸7转氨酶),ALT(丙氨酸氨基转移 酶)和肌酐活性

*Figure 6.* In vivo photodynamic effects and tumor regression seen inMDA-MB-231 xenograft mouse models.