

Thanks for reviewing our manuscript entitled “Effect of photodynamic therapy using (17R,18R)-2-(1-Hexyloxyethyl)-2-devinyl chlorine E6 trisodium salt on pancreatic cancer in vitro and in vivo”. (ID: 42372). We thank the reviewers for the fair and professional evaluations of our manuscript. We are convinced that the constructive comments of reviewers have led to a significant improvement of our manuscript. We hope those corrections would meet with your approvals. The main corrections in the paper and the responds to the reviewer’s comments are as following:

To reviewer (code: 03656606):

1. Minor language polishing

Response: According to your suggestion, we have got the manuscript checked by a native speaker and uploaded the editorial certificate. The manuscript was revised extensively in regard to comprehension and readability. We hope those corrections would meet with your approvals.

To reviewer (code: 02544565):

1. One thing, it is recommended to add the explanation about the spirulina. (Ex, spirulina is being taken as a supplement recently but has a light sensitivity as a side effects. Then YLG-1 was extracted as a causative agent and then recognized as a photosensitizer.)

Response: We consider that your advice is pretty good. According to your suggestion, We have added the explanation about the spirulina in the paragraph 3 of introduction part (Marked in red) and the related reference in reference part [11-14] .

Spirulina is now being considered as a supplement but has a light sensitivity as a side effect[11]. It has been used as the raw material extracted for many photosensitizers including Methyl 3-[1'-(m-iodobenzyloxy) ethyl] pyropheophorbide-a 2, selenium-containing phycocyanin and chlorin e6 (Ce6) [12-14].

Reference

- 11 Zeinalian R, Farhangi MA, Shariat A, Saghafi-Asl M. The effects of *Spirulina Platensis* on anthropometric indices, appetite, lipid profile and serum vascular endothelial growth factor (VEGF) in obese individuals: a randomized double blinded placebo controlled trial. *BMC Complement Altern Med* 2017; 17: 225 [PMID: 28431534 DOI:10.1186/s12906-017-1670-y]
- 12 Jeon YM, Lee HS, Jeong D, Oh HK, Ra KH, Lee MY. Antimicrobial photodynamic therapy using chlorin e6 with halogen light for acne bacteria-induced inflammation. *Life Sci* 2015; 124: 56-63 [PMID: 25623849 DOI:10.1016/j.lfs.2014.12.029]
- 13 Liu Z, Fu X, Huang W, Li C, Wang X, Huang B. Photodynamic effect and mechanism study of selenium-enriched phycocyanin from *Spirulina platensis* against liver tumours. *J Photochem Photobiol B* 2018; 180:89-97 [PMID: 29413706 DOI:10.1016/j.jphotobiol.2017.12.020]
- 14 Srivatsan A, Pera P, Joshi P, Wang Y, Missert JR, Tracy EC, Tabaczynski WA, Yao R, Sajjad M, Baumann H, Pandey RK. Effect of chirality on cellular uptake, imaging and photodynamic therapy of photosensitizers derived from chlorophyll-a. *Bioorg Med Chem* 2015; 23: 3603-3617 [PMID: 25936263 DOI:10.1016/j.bmc.2015.04.006]