Author’s response to reviews

Title: β adrenergic receptor/cAMP/PKA signaling contributes to the intracellular Ca2+ release by tentacle extract from the jellyfish Cyanea capillata

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Author’s response to reviews:

Dear editor,

Thank you for your letter and for the reviewer’s generous comments concerning our manuscript entitled “β adrenergic receptor/cAMP/PKA signaling contributes to the intracellular Ca2+ release by tentacle extract from the jellyfish Cyanea capillata” (PHAT-D-17-00055). Those comments are all valuable and helpful for revising and improving our paper. We have studied the comments carefully and have made corrections which we hope meet with approval. The point-by-point responses are as follows:

Responses to Reviewer 1:

1. Pg2 L17:"… suffer with a severe pain… "… revise to " suffer from a severe pain "

Response: Thank you for your careful review. We have revised the “suffer with a severe pain” to “suffer from a severe pain”.

2. Pg2 L39: " … and just few …." … revise to "…. just a few …." 

Response: We have revised the “… and just few ….” to “… just a few ….”.

3. Pg3 L4: "…. information on the general mechanism of cardiotoxicity..." remove general.
Response: We have removed the “general” in the “…. information on the general mechanism of cardiotoxicity...”.

4. Pg4 L15: "… at -70 °C till use..." …revise to "… at -70 °C until use...".

Response: We have revised the "… at -70 °C till use..." to "… at -70 °C until use...".

5. Pg5 L57: "…. 5-bromo-2-deoxyuridine (Brdu). " revise to "…. 5-bromo-2-deoxyuridine (BrdU). "

Response: We have revised the "...5-bromo-2-deoxyuridine (Brdu)" to " ...5-bromo-2-deoxyuridine (BrdU). "

6. Results section (Pg9): Remove all references within the results including their data. They should be moved to appropriated sites in the discussion section.

Response: Thank you for your professional advice. We have removed all references within the results section.

7. Pg12 L32: "… Intravenous injection with 10 µg/kg nematocyst…” revise to "….. Intravenous injection of 10 µg/kg nematocyst …"

Response: We have revised the "… Intravenous injection with 10 µg/kg nematocyst…” to "….. Intravenous injection of 10 µg/kg nematocyst …".

8. Pg12 L59: "….. that might attributed …" …. Revise to " …. that might be attributed …"

Response: We have revised the "…… that might attributed …" to " .... that might be attributed ...

Responses to Reviewer 2:

1. The acronyms/abbreviations should be written in full at first appearance in the text and put into brackets in parenthesis.

Response: Thank you very much for your kind advice. We have checked the text carefully and written the abbreviations in full at first appearance in our revised manuscript, such as: cyclic adenosine monophosphate (cAMP), protein kinase A (PKA) and phosphate buffer saline (PBS).

2. The name "cyanea capillata" should be italicized though out the article and the rest of scientific words.

Response: We have re-checked and adjusted all the “cyanea capillata” in italic type though out the article in our new manuscript.
3. The authors should be clear whether they used mouse or rat in their methods.

Response: It’s my duty not to express it clearly. In fact, we used both mouse and rat for different experiments in our research. We used adult mice to isolate the ventricular myocytes for confocal determination of intracellular calcium, and all the rest of the experiments in our methods were performed using rats, which we have described in the “Material and methods” and “Results” of the revised manuscript.

4. Page no 9, line no 8-27; The authors should avoid using citations/references at the result section.

Response: Thanks for your professional advice. According to your suggestion, we have removed the references at the result section.

5. At page 13-14, line no: 59-1, the sentence have grammatical errors.

Response: Thanks for your kind advice. I am sorry for the grammatical errors, and we have modified this sentence as below:

“Although the extracellular Ca2+ entry can be prevented by the L-type Ca2+ blockers, the mechanism of extracellular Ca2+ entry is now dominated by the existence of pore-forming complex in jellyfish venom that allows a non-specific translocation of Ca2+ across the cell membrane.”

6. At page no 12, line no:49; The word "does" should be replace with "dose".

Response: I am very sorry for the spelling mistake, and we had replaced the word “does” with “dose” in the revised manuscript.

7. The figures legends are not well labelled and also the graphs on page 26, 27 and 28 are congested/not clear.

Response: Thanks for your kind advice. We had re-labelled the figure legends and modified the graphs in our revised manuscript.