

Dear Editors,

Thank you very much for considering publication of our manuscript. The reviewers have made great comments on our manuscript. After discussion with all the co-authors, we revised this manuscript seriously and carefully according to reviews' opinions, and highlight the changes to our manuscript by using the track changes mode in MS Word. The answers to each reviewer are as follows:

1. Respond to Reviewer 1' comments:

The article is interesting and brings innovations. Minor language polishing.

Answer to question 1: Thank you for your comments, we have turned to professional English language editing company. The relevant changes have been marked in blue in the text.

2. Respond to Reviewer 2' comments:

Excellent review apart from little language corrections. Garcia-Tsao is a she not he.

Answer to question 2: Thanks a lot for this correction to our manuscript. We also recognized this mistake and corrected them.

3. Respond to Reviewer 3' comments:

Article of great scientific relevance, serving as a guide for research questions to be worked on worldwide. Excellent study design. Congratulations to the authors.

Answer to question 3: We are grateful for your compliments on this article.

4. Respond to editorial office's comments:

Please provide decomposable Figures (in which all components are movable and editable), organize them into a single PowerPoint file. If the picture is 'original', the author needs to add the following copyright information to the bottom right-hand side of the picture in PowerPoint (PPT): Copyright ©The Author(s) 2022.

Answer to question 3: We have provided decomposable Figures and organized them into a PowerPoint file. All pictures in this article are original, and copyright information has been added to the bottom right-hand side of the PowerPoint (PPT) picture.

In addition to correct this manuscript according to the reviewers' advice, we also checked the manuscript time and time again to avoid grammatical or spelling errors throughout the article. Hope our corrected manuscript can be published in your journal soon. That will be our great honor.

Thank you very much! Best wishes!

yours sincerely,  
Xiaowei Tang