Dear Prof. Ma:

Thank you very much for your letter and advice. We have revised the manuscript, and would like to re-submit it for your consideration. We have addressed the comments raised by the reviewers in the revised manuscript. Point by point responses to the reviewers' comments are listed as following:

Replies to Reviewer 1

some concerns

References need to be supplemented with articles in recent years.

Answer: We have updated the relative references with great influence in recent years.

A supplementary scale bar is required in Figure 3.

Answer: We added the supplementary scale bar in Figure 3

·Please explain the full name of the abbreviation when it first appears in the text.

Answer: We have attached the full name before each abbreviation when it first appears in this manuscript

Please explain the clinical impact of the current results.

Answer: The clinical impact of the current results have been added and explained in last but one paragraph of the "Discussion".

Replies to Reviewer 2

Specific Comments

8. References —cited little recent literature, Please add some relevant research articles published in the past three years.

Answer: We have updated the relative references with great influence in recent years.

We hope that the revised version of the manuscript is now acceptable for publication in your journal. I look forward to hearing from you soon.

With best wishes,

Yours sincerely,

Wen-bin Yu