

Dear editor,

Thank you very much for your email and a chance to revise our manuscript (manuscript ID: 82684). We now resubmit the revision of our manuscript. We appreciate editor and reviewers very much for their positive and constructive comments and suggestions on our manuscript. We tried our best to improve the manuscript and made some changes in the manuscript.

Thank you again for your time and kind help.

Yours sincerely,

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Our point-by-point responses to editor and reviewers' comments are as the following.

Editor Comments:

I recommend the manuscript to be published in the World Journal of Gastrointestinal Oncology.

1. Before final acceptance, when revising the manuscript, the author must supplement and improve the highlights of the latest cutting-edge research results, thereby further improving the content of the manuscript.

Response: Thank you for your advice. We have made the appropriate modifications.

2. To this end, authors are advised to apply a new tool, the Reference Citation Analysis (RCA). RCA is an artificial intelligence technology-based

open multidisciplinary citation analysis database. In it, upon obtaining search results from the keywords entered by the author, "Impact Index Per Article" under "Ranked by" should be selected to find the latest highlight articles, which can then be used to further improve an article under preparation/peer-review/revision. Please visit our RCA database for more information at: <https://www.referencecitationanalysis.com/>.

Response: Greatly appreciate your suggestion. We used the RCA database, which is a useful database.

Reviewer Comments:

Reviewer 1:

1) Dear Authors, this is a well-written manuscript presenting interesting clinical and experimental findings supporting the immune-oncology-microbiome (OPM) axis in the pathogenesis of gastrointestinal malignancies. These findings may find valuable implications in the clinical praxis and deserve to be published. Best Regards

Response: Thank you for your review.

Reviewer 2:

1) The manuscript hypothesized that: Macrophages may be a key component in linking the microbiome and immunity, which has been reflected in the variety of tumors mentioned above. The microbiome may influence tumor immune responses through TLRs on macrophages. ILCs also play a vital role in the host microbiome and, together with T cells, regulate the IOM axis. The IOM axis provides a new direction for the treatment of gastrointestinal malignancies.

Does the author self-cite, omit, incorrectly cite and/or over-cite references?

Yes; a latest reference is omitted eg: PMID: PMC7940198 DOI: 10.3389/fimmu.2021.622064

Response: Greatly appreciate your suggestion. We have added this in the references.