



PEER-REVIEW REPORT

Name of journal: *World Journal of Gastrointestinal Oncology*

Manuscript NO: 81757

Title: Intratumour microbiome of pancreatic cancer

Provenance and peer review: Invited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 05719281

Position: Peer Reviewer

Academic degree: Doctor, PhD

Professional title: Academic Research, Doctor

Reviewer's Country/Territory: Japan

Author's Country/Territory: China

Manuscript submission date: 2022-12-02

Reviewer chosen by: AI Technique

Reviewer accepted review: 2022-12-09 08:16

Reviewer performed review: 2022-12-22 06:52

Review time: 12 Days and 22 Hours

| | |
|---------------------------|---|
| Scientific quality | <input type="checkbox"/> Grade A: Excellent <input checked="" type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish |
| Language quality | <input checked="" type="checkbox"/> Grade A: Priority publishing <input type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection |
| Conclusion | <input type="checkbox"/> Accept (High priority) <input checked="" type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection |
| Re-review | <input type="checkbox"/> Yes <input checked="" type="checkbox"/> No |
| Peer-reviewer | Peer-Review: <input checked="" type="checkbox"/> Anonymous <input type="checkbox"/> Onymous |



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| statements | Conflicts-of-Interest: [] Yes [Y] No |
|------------|--|

SPECIFIC COMMENTS TO AUTHORS

Manuscript by Shi-Wei Guan, Quan Lin, Hai-Bo Yu. describes the current status of the intratumor microbiome in pancreatic cancer. This review article summarizes the progression, diagnosis, treatment, chemotherapy resistance, and immune response to bacteria in pancreatic cancer. This review article is interesting and well-organized. However, we found the following points to be of concern. <Minor comments The association with viruses needs to be added. For example, some reports support or deny the association between HBV and HCV and pancreatic cancer carcinogenicity. These reports should also be summarized. A review of bacterial-viral interactions for the development and carcinogenesis of virus-associated cancers is also needed.



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Reviewer's code: 05601558

Position: Peer Reviewer

Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: Italy

Author's Country/Territory: China

Manuscript submission date: 2022-12-02

Reviewer chosen by: AI Technique

Reviewer accepted review: 2022-12-30 06:13

Reviewer performed review: 2022-12-31 06:05

Review time: 23 Hours

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| Scientific quality | <input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input checked="" type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish |
| Novelty of this manuscript | <input type="checkbox"/> Grade A: Excellent <input checked="" type="checkbox"/> Grade B: Good <input type="checkbox"/> Grade C: Fair <input type="checkbox"/> Grade D: No novelty |
| Creativity or innovation of this manuscript | <input type="checkbox"/> Grade A: Excellent <input checked="" type="checkbox"/> Grade B: Good <input type="checkbox"/> Grade C: Fair <input type="checkbox"/> Grade D: No creativity or innovation |



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|---|--|
| Scientific significance of the conclusion in this manuscript | <input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Good <input checked="" type="checkbox"/> Grade C: Fair <input type="checkbox"/> Grade D: No scientific significance |
| Language quality | <input type="checkbox"/> Grade A: Priority publishing <input checked="" type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection |
| Conclusion | <input type="checkbox"/> Accept (High priority) <input type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input checked="" type="checkbox"/> Major revision <input type="checkbox"/> Rejection |
| Re-review | <input checked="" type="checkbox"/> Yes <input type="checkbox"/> No |
| Peer-reviewer statements | Peer-Review: <input checked="" type="checkbox"/> Anonymous <input type="checkbox"/> Onymous |
| | Conflicts-of-Interest: <input type="checkbox"/> Yes <input checked="" type="checkbox"/> No |

SPECIFIC COMMENTS TO AUTHORS

The authors uncovered the intratumour microbiome of pancreatic cancer. Points to be addressed: 1) The rationale of why the authors came up with this research is scanty and is related to a lack of novelty: please highlight what this manuscript might add. 2) What is the information that is not exactly available that motivated the authors to come up with this information. What are the current caveats and how do the authors highlight the current research in answering them? If not they need to address in background and in future directions . 3)State of the art figures are required: scale bar should be provided in high resolution. 4)The authors could provide a little more consideration of genomic directed stratifications in clinical trial design and enrolments. 5)The underlying message here is that more precision and individualized approaches need to be tested in well-designed clinical trials – a challenge, but I would be interested in their perspective of how this might be done. If beyond the scope of the manuscript, this should be highlighted as a limitation 6) The authors need to highlight what new information the review is providing to enhance the research in progress 7) This reviewer personally misses some insights in the introduction/discussion regarding



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genetic alterations and the potential relationship between those and the microbiome/immune landscape of PDAC. Indeed, especially the K-Ras mutation, carry the heaviest burden in the progression of pancreatic precursor lesions into pancreatic ductal adenocarcinoma (PDAC). The tumor microenvironment is one of the challenges that hinder the therapeutic approaches from functioning sufficiently and leads to the immune evasion of pancreatic malignant cells. Mastering the mechanisms of these two hallmarks of PDAC can help us in dealing with the obstacles in the way of treatment. In this review, we have analyzed the signaling pathways involved in PDAC development and the immune system's role in pancreatic cancer and immune checkpoint inhibition as next-generation therapeutic strategy. The direct targeting of the involved signaling molecules and the immune checkpoint molecules, along with a combination with conventional therapies, have reached the most promising results in pancreatic cancer treatment (please refer to PMID: 33918146 and expand)



RE-REVIEW REPORT OF REVISED MANUSCRIPT

Name of journal: *World Journal of Gastrointestinal Oncology*

Manuscript NO: 81757

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Provenance and peer review: Invited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 05601558

Position: Peer Reviewer

Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: Italy

Author's Country/Territory: China

Manuscript submission date: 2022-12-02

Reviewer chosen by: Jia-Ru Fan

Reviewer accepted review: 2023-02-23 03:27

Reviewer performed review: 2023-02-23 03:30

Review time: 1 Hour

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|---------------------------------|--|
| Scientific quality | <input checked="" type="checkbox"/> Grade A: Excellent [] Grade B: Very good [] Grade C: Good [] Grade D: Fair [] Grade E: Do not publish |
| Language quality | <input checked="" type="checkbox"/> Grade A: Priority publishing [] Grade B: Minor language polishing [] Grade C: A great deal of language polishing [] Grade D: Rejection |
| Conclusion | <input checked="" type="checkbox"/> Accept (High priority) [] Accept (General priority) [] Minor revision [] Major revision [] Rejection |
| Peer-reviewer statements | Peer-Review: <input checked="" type="checkbox"/> Anonymous [] Onymous Conflicts-of-Interest: [] Yes <input checked="" type="checkbox"/> No |



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SPECIFIC COMMENTS TO AUTHORS

The authors have clarified several of the questions I raised in my previous review. Most of the major problems have been addressed by this revision.