

PEER-REVIEW REPORT

Name of journal: *World Journal of Gastrointestinal Oncology*

Manuscript NO: 80245

Title: Challenges and Exploration for Immunotherapy Targeting Cold Colorectal Cancer

Provenance and peer review: Invited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 03982286

Position: Peer Reviewer

Academic degree: N/A

Professional title: N/A

Reviewer's Country/Territory: Japan

Author's Country/Territory: China

Manuscript submission date: 2022-09-20

Reviewer chosen by: AI Technique

Reviewer accepted review: 2022-09-23 13:27

Reviewer performed review: 2022-09-23 13:34

Review time: 1 Hour

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 Grade E: Do not publish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 type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input checked="" 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

statements

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

SPECIFIC COMMENTS TO AUTHORS

The authors show the Challenges and Exploration for Immunotherapy Targeting Cold Colorectal Cancer. The investigation is interesting, however, I have some concerns to be discussed. -How about the anti-CTA therapy using anti--MAGE-A4 or NY-ESO for colon cancer? -Is inflammation involved in the pathogenesis of colon cancer? Please discuss referring the following article. Characterization of PD-1/PD-L1 immune checkpoint expression in soft tissue sarcomas. European journal of histochemistry : EJH, 65(3), 3203. <https://doi.org/10.4081/ejh.2021.3203> -Do you think the colon cancer "hot tumor"? -How do you think the combined therapy with anti-PD-1/PD-L1 and chemotherapy for colon cancer?

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Manuscript NO: 80245

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Provenance and peer review: Invited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 05753782

Position: Peer Reviewer

Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: Malaysia

Author's Country/Territory: China

Manuscript submission date: 2022-09-20

Reviewer chosen by: AI Technique

Reviewer accepted review: 2022-09-21 02:29

Reviewer performed review: 2022-10-11 00:37

Review time: 19 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 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
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statementsConflicts-of-Interest: [] Yes [**Y**] No**SPECIFIC COMMENTS TO AUTHORS**

Comments to the authors · Authors' discussions are mainly on primary articles. Reviewer wants to suggest to include systematic review (SR) and meta-analysis (MA) articles on immunotherapy and other therapeutic regimes in colorectal cancer as MA are evidence based results. · In the discussion on response of immunotherapy, it is suggestive to include the efficacies and responses related to the status of immunohistochemical expression of immune check point proteins such as PD-1, PD-L1, CTLA4 etc. · For the reader's understanding, it is suggestive to explain some of the processes in the flow chart or diagrams such as cold tumour formation. · References need to be cleaned up. For example, reference 57 and 58 overlapped. Please check all references.