

PEER-REVIEW REPORT

Name of journal: *World Journal of Gastroenterology*

Manuscript NO: 83404

Title: Metronomic Capecitabine Inhibits Liver Transplant Rejection in Rats by Triggering Recipients' T Cell Ferroptosis

Provenance and peer review: Unsolicited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 03742333

Position: Editorial Board

Academic degree: FACS, MD, PhD

Professional title: Doctor, Full Professor, Professor, Surgeon

Reviewer's Country/Territory: Brazil

Author's Country/Territory: China

Manuscript submission date: 2023-01-21

Reviewer chosen by: AI Technique

Reviewer accepted review: 2023-01-24 18:36

Reviewer performed review: 2023-01-25 15:15

Review time: 20 Hours

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 Grade E: Do not publish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

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 type="checkbox"/> Major revision <input checked="" 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

I have read with great interest the manuscript entitled “Metronomic Capecitabine Inhibits Liver Transplant Rejection in Rats by Triggering Recipients’ T Cell Ferroptosis”, submitted to the World Journal of Gastroenterology. The manuscript is well written, and all the experiments were conducted in accordance with the highest standards. Nevertheless, I have a few comments: **MAJOR** The major concern in the study is related to the study design. More specifically, how it translates or reproduces clinical practice; this is because, apparently, there was no initial immunosuppression to the transplanted rats from the control group until 7 days. This is not a routine practice of any centre. Therefore, I wonder if the effects seen were just a consequence and what would be this effect on the administration of the gold standard (tacrolimus). **MINOR** The Conclusion section in the abstract and the main text must be amended. It must state that all the findings were verified experimentally in a rat model. The same applies to the core tip. The discussion must emphasise how feasible the utilisation of the medication is after liver transplantation, considering the adverse effects. In addition, do the authors propose its use for the recurrence of HCC after liver transplantation? Or in cases of HCC



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transplants?

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Peer-review model: Single blind

Reviewer's code: 06468328

Position: Peer Reviewer

Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: Korea

Author's Country/Territory: China

Manuscript submission date: 2023-01-21

Reviewer chosen by: AI Technique

Reviewer accepted review: 2023-01-28 10:56

Reviewer performed review: 2023-02-06 02:06

Review time: 8 Days and 15 Hours

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 type="checkbox"/> Grade B: Good <input checked="" type="checkbox"/> Grade C: Fair <input type="checkbox"/> Grade D: No creativity or innovation

Scientific significance of the conclusion in 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 scientific significance
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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 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

This study is well-written, concise, and coherently organized, and identifies the role of metronomic CAP in patients with LT. The study flow and results are clear. 1. The authors confirmed the effect of CAP on CD3 T cells. Among T cells, which of CD4+ or CD8+ cells is affected by CAP. 2. It is recommended to change the expression from '5-Fu' to '5-FU'.

RE-REVIEW REPORT OF REVISED MANUSCRIPT

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Reviewer's Country/Territory: Brazil

Author's Country/Territory: China

Manuscript submission date: 2023-01-21

Reviewer chosen by: Jia-Ru Fan

Reviewer accepted review: 2023-04-01 21:01

Reviewer performed review: 2023-04-01 21:27

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 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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statements

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

SPECIFIC COMMENTS TO AUTHORS

The authors clarified the major concerns satisfactorily and amended the manuscript in accordance.