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PEER-REVIEW REPORT

Name of journal: *World Journal of Gastrointestinal Oncology*

Manuscript NO: 85408

Title: Physcion Increases the Sensitivity of Hepatocellular Carcinoma to Sorafenib through microRNA-370/PIM1 axis-regulated glycolysis

Provenance and peer review: Unsolicited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 03805472
Position: Peer Reviewer
Academic degree: MD, PhD

Professional title: Doctor

Reviewer's Country/Territory: Italy

Author's Country/Territory: China

Manuscript submission date: 2023-04-26

Reviewer chosen by: Geng-Long Liu

Reviewer accepted review: 2023-05-22 07:10

Reviewer performed review: 2023-05-31 01:11

Review time: 8 Days and 18 Hours

Scientific quality	[ ] Grade A: Excellent [Y] Grade B: Very good [ ] Grade C: Good [ ] Grade D: Fair [ ] Grade E: Do not publish
Novelty of this manuscript	[ ] Grade A: Excellent [Y] Grade B: Good [ ] Grade C: Fair [ ] Grade D: No novelty
Creativity or innovation of this manuscript	[ ] Grade A: Excellent [Y] Grade B: Good [ ] Grade C: Fair [ ] Grade D: No creativity or innovation

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Scientific significance of the conclusion in this manuscript	[ ] Grade A: Excellent [Y] Grade B: Good [ ] Grade C: Fair [ ] Grade D: No scientific significance
Language quality	[ ] Grade A: Priority publishing [Y] Grade B: Minor language polishing [ ] Grade C: A great deal of language polishing [ ] Grade D: Rejection
Conclusion	[ ] Accept (High priority) [ ] Accept (General priority) [Y] Minor revision [ ] Major revision [ ] Rejection
Re-review	[ ] Yes [Y] No
Peer-reviewer statements	Peer-Review: [Y] Anonymous [ ] Onymous
	Conflicts-of-Interest: [ ] Yes [Y] No

SPECIFIC COMMENTS TO AUTHORS

Aerobic glycolysis is a well-recognized hallmark of cancer cells, and it is critical for cancer initiation and development. It is worthy of note that a recent study reported that sorafenib impaired OXPHOS and promoted glycolysis in HCC. The physcion acts as a suppressor of metastasis and plays a pivotal role in chemosensitization. However, the effect of physcion during sorafenib resistance has not been studied yet. In this study, the authors designed the research to investigate the effect of physcion on the sensitivity of HCC cells to sorafenib. The research is well performed. The methods are described in detail, and the results are very interesting. Minor comments: 1. The statistical methods should be described in the method section. 2. How about the limit of this research? Please make a discussion about it. 3. The quality of the images should be improved. 4. Some minor language polishing should be corrected.

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Provenance and peer review: Unsolicited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 06143444
Position: Peer Reviewer
Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: Austria

Author's Country/Territory: China

Manuscript submission date: 2023-04-26

Reviewer chosen by: Geng-Long Liu

Reviewer accepted review: 2023-05-24 00:46

Reviewer performed review: 2023-05-31 01:40

Review time: 7 Days

Scientific quality	[ ] Grade A: Excellent [ ] Grade B: Very good [Y] Grade C: Good [ ] Grade D: Fair [ ] Grade E: Do not publish
Novelty of this manuscript	[ ] Grade A: Excellent [Y] Grade B: Good [ ] Grade C: Fair [ ] Grade D: No novelty
Creativity or innovation of this manuscript	[ ] Grade A: Excellent [Y] Grade B: Good [ ] Grade C: Fair [ ] Grade D: No creativity or innovation

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Conclusion	[ ] Accept (High priority) [Y] Accept (General priority) [ ] Minor revision [ ] Major revision [ ] Rejection
Re-review	[ ] Yes [Y] No
Peer-reviewer statements	Peer-Review: [Y] Anonymous [ ] Onymous
	Conflicts-of-Interest: [ ] Yes [Y] No

SPECIFIC COMMENTS TO AUTHORS

This is an interesting study about the physcion in hepatocellular carcinoma. The study is well designed and the findings are interesting. Discussion is good, and references are updated. The authors should make a carefully editing to the manuscript, and update the figures.

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Peer-review model: Single blind

Reviewer's code: 03633805
Position: Peer Reviewer
Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: Croatia

Author's Country/Territory: China

Manuscript submission date: 2023-04-26

Reviewer chosen by: Geng-Long Liu

Reviewer accepted review: 2023-06-02 04:44

Reviewer performed review: 2023-06-14 12:25

Review time: 12 Days and 7 Hours

Scientific quality	[ ] Grade A: Excellent [Y] Grade B: Very good [ ] Grade C: Good [ ] Grade D: Fair [ ] Grade E: Do not publish
Novelty of this manuscript	[ ] Grade A: Excellent [Y] Grade B: Good [ ] Grade C: Fair [ ] Grade D: No novelty
Creativity or innovation of this manuscript	[ ] Grade A: Excellent [Y] Grade B: Good [ ] Grade C: Fair [ ] Grade D: No creativity or innovation

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Re-review	[Y] Yes [ ] No
Peer-reviewer statements	Peer-Review: [Y] Anonymous [ ] Onymous
	Conflicts-of-Interest: [ ] Yes [Y] No

SPECIFIC COMMENTS TO AUTHORS

The paper is concise, logically written, and organized. The hypothesis and corresponding conclusion are clear, followed by a detailed and well-depicted scientific methodology. In order to further improve scientific value the Discussion/Conclusion section could also tackle the following questions: 1.Taking into account myriads of other anti-cancer mechanisms of action for Physcion in other cancer cell types , as well as established promotion of chemosensitivity, what would be the author's opinion about the potential effectiveness of Physcion in other targeted therapies resistances 2. Could numerous established anticancer mechanisms for Physcion (antineoplastic, anti-inflammatory, antiangiogenic, and antiproliferative) suggest that PIM1-regulated glycolysis is not the only mechanism underlying sorafenib sensitivity