

PEER-REVIEW REPORT

Name of journal: *World Journal of Stem Cells*

Manuscript NO: 82221

Title: Delineating the glioblastoma stemness by genes involved in cytoskeletal rearrangements and metabolic alterations

Provenance and peer review: Invited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 05200667

Position: Editorial Board

Academic degree: BSc, MPhil, PhD

Professional title: Adjunct Professor, Professor, Research Scientist, Senior Scientist, Vice Editor-in-Chief

Reviewer's Country/Territory: United States

Author's Country/Territory: Poland

Manuscript submission date: 2022-12-10

Reviewer chosen by: Dong-Mei Wang

Reviewer accepted review: 2023-01-16 22:22

Reviewer performed review: 2023-01-17 00:12

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 Grade E: Do not publish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
Conclusion	<input type="checkbox"/> Accept (High priority) <input type="checkbox"/> Accept (General priority) <input checked="" 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	Peer-Review: [<input type="checkbox"/>] Anonymous [<input checked="" type="checkbox"/>] Onymous
statements	Conflicts-of-Interest: [<input type="checkbox"/>] Yes [<input checked="" type="checkbox"/>] No

SPECIFIC COMMENTS TO AUTHORS

Comment: (MS #82221) The specific argument for the contribution of the link between metabolism and cytoskeleton to GBM stemness is of great interest. However, the current manuscript version did not fully capture the forceful logic with tightened clarity rather than generic narratives. Specific comments: 1) Abstract: "Previously, we proved that interplay between metabolism and cytoskeleton exists in GBM." A schematic diagram to summarize this interface could power up their argument. 2) Figure 1: Figure 1 Impact of described genes on biological processes related to stem cells. It is hard to follow with such complex black dashed lines or solid black lines over the places. A side-by-side table should be used to enhance clarity. 3) Page 4: "Adding tumor-treating electric fields (TTFields) to maintenance TMZ chemotherapy was found to prolong progression-free and overall survival but is currently limited due to the lack of a method to predict or quantify the efficacy of TTFields [5]." This statement contradicted: did "prolong progression-free and overall survival" quantify the efficacy of TTFields? 4) P4: "non-stem glioblastoma cells are less invasive than GBM stem cells (GSCs) [17], " How less is less? How did they determine "less" - do all GBM cells invade surrounding tissues? E.g., GSCs or non-GSCs came with enhanced MMP-family production. 5) P5: If only three genes PLEK2, RRM2, GCSH as shown in Ref #22, of metabolic alterations and cytoskeletal rearrangements, please focus on them to expand instead of generic statements. The list of either group should be provided. 6) The discussion lacks the grip of integration for all the genes in cross-talk networks. E.g., How could they integrate the glioblastoma biomarkers [213] with their specific argument for the



**Baishideng
Publishing
Group**

7041 Koll Center Parkway, Suite
160, Pleasanton, CA 94566, USA
Telephone: +1-925-399-1568
E-mail: bpgoffice@wjgnet.com
<https://www.wjgnet.com>

contribution of the link between metabolism and cytoskeleton to GBM stemness? The authors narrate many independent studies on various tumor types but do not tighten up and draw the line back to their focus. E.g., “to emphasize the role of described genes specifically in stem cells, setting aside the rest of the information provided for each gene (Figure 1). At first glance, the most frequently regulated processes are proliferation and chemoresistance, followed by differentiation, tumor growth, invasion, and apoptosis.” Note that these functions were not entirely gravitated toward their specific argument for the contribution of the link between metabolism and cytoskeleton to GBM stemness.

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Position: Peer Reviewer

Academic degree: MD, PhD

Professional title: Attending Doctor, Neurosurgeon, Postdoc, Research Assistant

Reviewer's Country/Territory: China

Author's Country/Territory: Poland

Manuscript submission date: 2022-12-10

Reviewer chosen by: Dong-Mei Wang

Reviewer accepted review: 2023-01-17 11:54

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Review time: 1 Hour

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
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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 review is comprehensive and timely. One minor poits need to be revised.
Demonstration of figure is chaotic, need to be redraw.