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PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 47507

Title: Pharmacogenetics of the systemic treatment in advanced hepatocellular carcinoma

Reviewer's code: 02985646

Reviewer's country: United States

Science editor: Jia-Ping Yan

Reviewer accepted review: 2019-03-18 13:44

Reviewer performed review: 2019-04-02 17:34

Review time: 15 Days and 3 Hours

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
[] Grade A: Excellent	[] Grade A: Priority publishing	[] Accept	Peer-Review:
[] Grade B: Very good	[Y] Grade B: Minor language	(High priority)	[Y] Anonymous
[] Grade C: Good	polishing	[] Accept	[] Onymous
[Y] Grade D: Fair	[] Grade C: A great deal of	(General priority)	Peer-reviewer's expertise on the
[] Grade E: Do not	language polishing	[] Minor revision	topic of the manuscript:
publish	[] Grade D: Rejection	[Y] Major revision	[Y] Advanced
		[] Rejection	[] General
			[] No expertise
			Conflicts-of-Interest:
			[] Yes
			[Y] No

SPECIFIC COMMENTS TO AUTHORS

I appreciate the thoroughness and completeness of the review paper. However, there are many areas for improvement. Overall, the review paper is very long, unfocused, and does not provide a higher-level synthesis of the data (i.e. much of the manuscript simply restates conclusions from other studies without providing the authors



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interpretation of what the results really mean, how do they impact current pharmacogenomics research, will it impact clinical practice, what are the next steps needed in research to validate findings, etc.). Further, the authors review many different trials, but did not include any numbers or statistics for most of the trials (see attachment). There is a lot of information that is not pertinent to the aim or title of the manuscript (see attachment).

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PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 47507

Title: Pharmacogenetics of the systemic treatment in advanced hepatocellular carcinoma

Reviewer's code: 03074879

Reviewer's country: China

Science editor: Jia-Ping Yan

Reviewer accepted review: 2019-04-17 08:14

Reviewer performed review: 2019-04-17 09:05

Review time: 1 Hour

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
[] Grade A: Excellent	[] Grade A: Priority publishing	[] Accept	Peer-Review:
[Y] Grade B: Very good	[Y] Grade B: Minor language	(High priority)	[] Anonymous
[] Grade C: Good	polishing	[Y] Accept	[Y] Onymous
[] Grade D: Fair	[] Grade C: A great deal of	(General priority)	Peer-reviewer's expertise on the
[] Grade E: Do not	language polishing	[] Minor revision	topic of the manuscript:
publish	[] Grade D: Rejection	[] Major revision	[] Advanced
		[] Rejection	[Y] General
			[] No expertise
			Conflicts-of-Interest:
			[] Yes
			[Y] No

SPECIFIC COMMENTS TO AUTHORS

The paper had carefully read by us. This paper summarized and critically discussed the pharmacogenetic literature evidence, with particular attention to sorafenib and regorafenib, which have been used longer than the others in HCC treatment and introduced the the previous research progress of approved drugs. It is a well written



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manuscript with a large quantity of data and detailed description.

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PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 47507

Title: Pharmacogenetics of the systemic treatment in advanced hepatocellular carcinoma

Reviewer's code: 02444752

Reviewer's country: China

Science editor: Jia-Ping Yan

Reviewer accepted review: 2019-04-17 08:55

Reviewer performed review: 2019-04-19 07:34

Review time: 1 Day and 22 Hours

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
[Y] Grade A: Excellent	[Y] Grade A: Priority publishing	[Y] Accept	Peer-Review:
[] Grade B: Very good	[] Grade B: Minor language	(High priority)	[Y] Anonymous
[] Grade C: Good	polishing	[] Accept	[] Onymous
[] Grade D: Fair	[] Grade C: A great deal of	(General priority)	Peer-reviewer's expertise on the
[] Grade E: Do not	language polishing	[] Minor revision	topic of the manuscript:
publish	[] Grade D: Rejection	[] Major revision	[] Advanced
		[] Rejection	[Y] General
			[] No expertise
			Conflicts-of-Interest:
			[] Yes
			[Y] No

SPECIFIC COMMENTS TO AUTHORS

This review is by far the most comprehensive introduction to the topic of pharmacogenetics in the treatment of advanced liver cancer. I suggest that priority be given to the publication of this review article.



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PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 47507

Title: Pharmacogenetics of the systemic treatment in advanced hepatocellular carcinoma

Reviewer's code: 02861333

Reviewer's country: China

Science editor: Jia-Ping Yan

Reviewer accepted review: 2019-04-17 06:53

Reviewer performed review: 2019-04-20 02:04

Review time: 2 Days and 19 Hours

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
[Y] Grade A: Excellent	[Y] Grade A: Priority publishing	[] Accept	Peer-Review:
[] Grade B: Very good	[] Grade B: Minor language	(High priority)	[Y] Anonymous
[] Grade C: Good	polishing	[Y] Accept	[] Onymous
[] Grade D: Fair	[] Grade C: A great deal of	(General priority)	Peer-reviewer's expertise on the
[] Grade E: Do not	language polishing	[] Minor revision	topic of the manuscript:
publish	[] Grade D: Rejection	[] Major revision	[Y] Advanced
		[] Rejection	[] General
			[] No expertise
			Conflicts-of-Interest:
			[] Yes
			[Y] No

SPECIFIC COMMENTS TO AUTHORS

This review study summarized and discussed the pharmacogenetic literature evidence for targeted agents in the systemic treatment of advanced HCC. No comments.



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PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 47507

Title: Pharmacogenetics of the systemic treatment in advanced hepatocellular carcinoma

Reviewer's code: 03408355

Reviewer's country: China

Science editor: Jia-Ping Yan

Reviewer accepted review: 2019-04-20 04:21

Reviewer performed review: 2019-04-20 06:51

Review time: 2 Hours

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
[] Grade A: Excellent	[] Grade A: Priority publishing	[] Accept	Peer-Review:
[] Grade B: Very good	[Y] Grade B: Minor language	(High priority)	[Y] Anonymous
[Y] Grade C: Good	polishing	[] Accept	[] Onymous
[] Grade D: Fair	[] Grade C: A great deal of	(General priority)	Peer-reviewer's expertise on the
[] Grade E: Do not	language polishing	[Y] Minor revision	topic of the manuscript:
publish	[] Grade D: Rejection	[] Major revision	[Y] Advanced
		[] Rejection	[] General
			[] No expertise
			Conflicts-of-Interest:
			[] Yes
			[Y] No

SPECIFIC COMMENTS TO AUTHORS

1. The abstract was a bit longer. Please shorten it. 2. There were up to 10 keywords, which was too many.



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PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 47507

Title: Pharmacogenetics of the systemic treatment in advanced hepatocellular carcinoma

Reviewer's code: 03546970

Reviewer's country: China

Science editor: Jia-Ping Yan

Reviewer accepted review: 2019-04-22 11:09

Reviewer performed review: 2019-04-23 09:52

Review time: 22 Hours

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
[] Grade A: Excellent	[] Grade A: Priority publishing	[] Accept	Peer-Review:
[Y] Grade B: Very good	[Y] Grade B: Minor language	(High priority)	[Y] Anonymous
[] Grade C: Good	polishing	[Y] Accept	[] Onymous
[] Grade D: Fair	[] Grade C: A great deal of	(General priority)	Peer-reviewer's expertise on the
[] Grade E: Do not	language polishing	[] Minor revision	topic of the manuscript:
publish	[] Grade D: Rejection	[] Major revision	[] Advanced
		[] Rejection	[Y] General
			[] No expertise
			Conflicts-of-Interest:
			[] Yes
			[Y] No

SPECIFIC COMMENTS TO AUTHORS

In the present manuscript, Dr. Elena De Mattia et al. reviewed and discussed the pharmacogenetic literature evidence of target therapy drugs. The identification of biomarkers for therapeutic effect prediction is helpful for the choice of clinical treatment. The present manuscript could be accepted on World Journal of Gastroenterology.



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Comments are as follows: 1. The abstract should be more concise. 2. Tivantinib is c-Met inhibitor, rather than mesenchymal-epithelial transcription factor inhibitor. 3. There were some results described in the article without references, such as chemoembolization clinical trials. 4. Wrong reference cited in reference 54 and 55, please revise it.

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