



PEER-REVIEW REPORT

Name of journal: World Journal of Hepatology

Manuscript NO: 41893

Title: Hepatic Encephalopathy: lessons from preclinical studies.

Reviewer's code: 03529813

Reviewer's country: Portugal

Science editor: Ying Dou

Date sent for review: 2018-08-29

Date reviewed: 2018-09-03

Review time: 5 Days

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

SPECIFIC COMMENTS TO AUTHORS

Dear authors, first of all, congratulations for the manuscript. Although the text is well written, the discussion of each model of hepatic encephalopathy is very short and you should think to improve it before publication and also to explain better the pros and cons of each model. Also in the text it is stated that "Ideally, models of HE associated to



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portosystemic shunting (type B) and due to chronic liver disease (type C) must exhibit liver cirrhosis...". This paragraph may lead to misunderstandings since type B encephalopathy does not occur only in cirrhotic patients, in fact most of the case have no chronic liver disease, so may you explain why the models for type B should exhibit liver cirrhosis?

INITIAL REVIEW OF THE MANUSCRIPT

Google Search:

- The same title
- Duplicate publication
- Plagiarism
- No

BPG Search:

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- Plagiarism
- No



PEER-REVIEW REPORT

Name of journal: World Journal of Hepatology

Manuscript NO: 41893

Title: Hepatic Encephalopathy: lessons from preclinical studies.

Reviewer’s code: 03350345

Reviewer’s country: France

Science editor: Ying Dou

Date sent for review: 2018-08-29

Date reviewed: 2018-09-17

Review time: 19 Days

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	<input type="checkbox"/> Accept	Peer-Review:
<input type="checkbox"/> Grade B: Very good	<input checked="" type="checkbox"/> Grade B: Minor language	(High priority)	<input checked="" type="checkbox"/> Anonymous
<input checked="" type="checkbox"/> Grade C: Good	polishing	<input type="checkbox"/> Accept	<input type="checkbox"/> Onymous
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade C: A great deal of	(General priority)	Peer-reviewer’s expertise on the
<input type="checkbox"/> Grade E: Do not	language polishing	<input type="checkbox"/> Minor revision	topic of the manuscript:
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		<input type="checkbox"/> Rejection	<input type="checkbox"/> General
			<input type="checkbox"/> No expertise
			Conflicts-of-Interest:
			<input type="checkbox"/> Yes
			<input checked="" type="checkbox"/> No

SPECIFIC COMMENTS TO AUTHORS

Dr Lima et al present a review article on Hepatic Encephalopathy (HE) animal models. This article give a rapid overview on the available models in the study of HE pathophysiology. Some shortcomings are however present: - it would be important for the reader to state for each model whether it is associated with hyperammonemia and



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systemic inflammation (cf the pathophysiology section). This should be added in the text and the tables - in Introduction, the references 2 and 9 are very general and do not clearly state for what they are appealed - in session 3.1, it would be interesting to talk to the reader about the possible local production of ammonia in enterocytes - in section 3.1, just before section 3.2, mention of some work about lowering ammonia agents could be tied (Rockey et al, Hepato 2014, Weiss et al Fund Clin Pharmacol 2017, STOP-HE study by Ocera pharmaceuticals) - section 3.2, CXCL1 is probably CXCL-1 - the carbon tetrachloride (CCL4) model should be discussed since it has been used by some authors. - what about the model of just giving hyperammonemia diet to Wistar rats? some teams use this model - Tables should be reformatted: clinical signs should be separated from biological findings - a figure would be helpful

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PEER-REVIEW REPORT

Name of journal: World Journal of Hepatology

Manuscript NO: 41893

Title: Hepatic Encephalopathy: lessons from preclinical studies.

Reviewer's code: 00030603

Reviewer's country: Greece

Science editor: Ying Dou

Date sent for review: 2018-09-26

Date reviewed: 2018-10-08

Review time: 12 Days

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	<input type="checkbox"/> Accept	Peer-Review:
<input type="checkbox"/> Grade B: Very good	<input checked="" type="checkbox"/> Grade B: Minor language	(High priority)	<input checked="" type="checkbox"/> Anonymous
<input checked="" type="checkbox"/> Grade C: Good	polishing	<input checked="" type="checkbox"/> Accept	<input type="checkbox"/> Onymous
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publish	<input type="checkbox"/> Grade D: Rejection	<input type="checkbox"/> Major revision	<input type="checkbox"/> Advanced
		<input type="checkbox"/> Rejection	<input checked="" type="checkbox"/> General
			<input type="checkbox"/> No expertise
			Conflicts-of-Interest:
			<input type="checkbox"/> Yes
			<input checked="" type="checkbox"/> No

SPECIFIC COMMENTS TO AUTHORS

Nice review about preclinical studies in hepatic encephalopathy.

INITIAL REVIEW OF THE MANUSCRIPT

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