

ESPS PEER-REVIEW REPORT

Name of journal: World Journal of Hepatology

ESPS manuscript NO: 20571

Title: Role of interleukin-1 and its antagonism of hepatic stellate cell proliferation and liver fibrosis in the Abcb4-/- mouse model

Reviewer's code: 00006459

Reviewer's country: Australia

Science editor: Shui Qiu

Date sent for review: 2015-06-11 22:31

Date reviewed: 2015-08-30 09:06

CLASSIFICATION	LANGUAGE EVALUATION	SCIENTIFIC MISCONDUCT	CONCLUSION
<input type="checkbox"/> Grade A: Excellent	<input checked="" type="checkbox"/> Grade A: Priority publishing	Google Search:	<input type="checkbox"/> Accept
<input checked="" type="checkbox"/> Grade B: Very good	<input type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> The same title	<input type="checkbox"/> High priority for publication
<input type="checkbox"/> Grade C: Good	<input type="checkbox"/> Grade C: A great deal of language polishing	<input type="checkbox"/> Duplicate publication	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade D: Rejected	<input checked="" type="checkbox"/> No	<input checked="" type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor		BPG Search:	<input type="checkbox"/> Major revision
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input checked="" type="checkbox"/> No	

COMMENTS TO AUTHORS

This paper shows that the IL1 signaling antagonist Anakinra can influence mouse HSC in vitro but not fibrosis in the Abcb4 mouse. Additional novel data is that female mice in this model are more affected than male mice. As PSC is more common in human females than males, perhaps this model will be useful in gaining additional understanding of the human situation. Needed improvements:

1. Fig 2: Please also analyse this data to compare Anakinra treatment versus control: is the Anakinra treatment lowering proliferation compared to control? If so, is there a background level of IL1 signaling ?
2. In the methods section please add a statement on whether ethics approval was obtained and the rules adhered to.
3. Please correct the spelling of hydroxyproline and phenotype in abstract and figure labels [fig 1]

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Name of journal: World Journal of Hepatology

ESPS manuscript NO: 20571

Title: Role of interleukin-1 and its antagonism of hepatic stellate cell proliferation and liver fibrosis in the Abcb4-/- mouse model

Reviewer's code: 00160393

Reviewer's country: China

Science editor: Shui Qiu

Date sent for review: 2015-06-11 22:31

Date reviewed: 2015-09-02 15:41

CLASSIFICATION	LANGUAGE EVALUATION	SCIENTIFIC MISCONDUCT	CONCLUSION
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	Google Search:	<input type="checkbox"/> Accept
<input type="checkbox"/> Grade B: Very good	<input type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> The same title	<input type="checkbox"/> High priority for publication
<input type="checkbox"/> Grade C: Good		<input type="checkbox"/> Duplicate publication	
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade C: A great deal of language polishing	<input type="checkbox"/> Plagiarism	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade E: Poor	<input type="checkbox"/> Grade D: Rejected	<input type="checkbox"/> No	<input type="checkbox"/> Minor revision
		BPG Search:	<input type="checkbox"/> Major revision
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input type="checkbox"/> No	

COMMENTS TO AUTHORS

In this manuscript, the authors showed that IL-1 β promoted the proliferation of murine HSC and this effect could be abolished by the IL-1-receptor antagonist Anakinra in vitro. In addition, Anakinra failed to improve liver-fibrosis in this preclinical PSC model. There are a few questions: 1) The conclusion that "IL-1-antagonism shows antifibrotic effects in vitro.... is not correct because antifibrotic effects should be tested in vivo. In addition, the author should also observe the effect of Anakinra on HSC activation. 2) Another experimental animal model (such as bile duct ligation) should be used in order to confirm antifibrotic effect of Anakinra. Why the author only chose PSC model? 3) The paper is not well written..

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Name of journal: World Journal of Hepatology

ESPS manuscript NO: 20571

Title: Role of interleukin-1 and its antagonism of hepatic stellate cell proliferation and liver fibrosis in the Abcb4-/- mouse model

Reviewer's code: 02861134

Reviewer's country: Bulgaria

Science editor: Shui Qiu

Date sent for review: 2015-06-11 22:31

Date reviewed: 2015-08-27 17:03

CLASSIFICATION	LANGUAGE EVALUATION	SCIENTIFIC MISCONDUCT	CONCLUSION
<input checked="" type="checkbox"/> Grade A: Excellent	<input checked="" type="checkbox"/> Grade A: Priority publishing	Google Search:	<input checked="" type="checkbox"/> Accept
<input type="checkbox"/> Grade B: Very good	<input type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> The same title	<input type="checkbox"/> High priority for publication
<input type="checkbox"/> Grade C: Good	<input type="checkbox"/> Grade C: A great deal of language polishing	<input type="checkbox"/> Duplicate publication	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade D: Rejected	<input checked="" type="checkbox"/> No	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor		BPG Search:	<input type="checkbox"/> Major revision
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input checked="" type="checkbox"/> No	

COMMENTS TO AUTHORS

The method applied in the study to analyze the research topic is praise-worthy. There is a good consistency of analysis throughout the whole paper. I would like to recommend this paper to publish with no significant modification.

ESPS PEER-REVIEW REPORT

Name of journal: World Journal of Hepatology

ESPS manuscript NO: 20571

Title: Role of interleukin-1 and its antagonism of hepatic stellate cell proliferation and liver fibrosis in the Abcb4^{-/-} mouse model

Reviewer's code: 01810523

Reviewer's country: United States

Science editor: Shui Qiu

Date sent for review: 2015-06-11 22:31

Date reviewed: 2015-08-24 06:37

CLASSIFICATION	LANGUAGE EVALUATION	SCIENTIFIC MISCONDUCT	CONCLUSION
<input checked="" type="checkbox"/> Grade A: Excellent	<input checked="" type="checkbox"/> Grade A: Priority publishing	Google Search:	<input checked="" type="checkbox"/> Accept
<input type="checkbox"/> Grade B: Very good	<input type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> The same title	<input type="checkbox"/> High priority for publication
<input type="checkbox"/> Grade C: Good	<input type="checkbox"/> Grade C: A great deal of language polishing	<input type="checkbox"/> Duplicate publication	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade D: Rejected	<input checked="" type="checkbox"/> No	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor		BPG Search:	<input type="checkbox"/> Major revision
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input checked="" type="checkbox"/> No	

COMMENTS TO AUTHORS

Activation of HSC is considered a major pathological event in development of liver fibrosis. In this study, Reiter et al. evaluated the role of HSC IL-1 pathway in the pathogenesis of liver fibrosis. The authors provided strong evidence demonstrating the pro-fibrogenic effect of IL-1 pathway, and more importantly the distinct anti-fibrogenic effect of Anakinra in vitro vs. in vivo using the Abcb4 deficient mouse model. These results suggest that multiple pathways need to be considered for therapeutic intervention of different types of liver fibrosis.