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PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 38213

Title: Multiomics Biomarkers for the Prediction of Nonalcoholic Fatty Liver Disease Severity

Reviewer's code: 00199807

Reviewer's country: Turkey

Science editor: Ya-Juan Ma

Date sent for review: 2018-02-01

Date reviewed: 2018-02-06

Review time: 5 Days

CLASSIFICATION	LANGUAGE EVALUATION	SCIENTIFIC MISCONDUCT	CONCLUSION
<input type="checkbox"/> Grade A: Excellent	<input checked="" type="checkbox"/> Grade A: Priority publishing	Google Search:	<input checked="" 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 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

Dear Editor, I reviewed the manuscript titled "Multiomics Biomarkers for the Prediction of Nonalcoholic Fatty Liver Disease Severity". I think this paper can be accepted in this current form. Sincerely yours.



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PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 38213

Title: Multiomics Biomarkers for the Prediction of Nonalcoholic Fatty Liver Disease Severity

Reviewer's code: 02861277

Reviewer's country: Italy

Science editor: Ya-Juan Ma

Date sent for review: 2018-02-01

Date reviewed: 2018-02-09

Review time: 8 Days

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 checked="" type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> The same title	<input type="checkbox"/> High priority for publication
<input checked="" 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"/> Plagiarism	<input checked="" type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor		<input checked="" type="checkbox"/> No	<input type="checkbox"/> Major revision
		BPG Search:	
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input checked="" type="checkbox"/> No	

COMMENTS TO AUTHORS

In the manuscript, entitled "Multiomics Biomarkers for the Prediction of Nonalcoholic Fatty Liver Disease Severity" the authors provided an exhaustive overview concerning biomarkers and their integrative analysis in predicting NASH severity. In my opinion, they focused on a very promising clinical tool that in the era of personalized medicine might have a huge scientific relevance. Comments I think that it should be always kept in mind that NAFLD/NASH is an inflammatory disease involving both innate and adaptive immune responses and for these latter their interplay with the oxidative stress. Indeed some potential therapeutic options are directed against these pathogenetic mechanisms such as Vitamin E and Cenicriviroc (CVC) etc. Considering the relevance of these molecular targets, we cannot exclude in the near future to use additional biomarkers immunity-related such as circulating levels of cyto/chemokines, antibodies



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etc. in predicting NASH progression toward advanced phases. Taking into account these premises, I would suggest for a compelling analysis to give some hints (in the introduction section) to the role of innate and adaptive immunity in NASH (see for instance: Therapeutic Inhibition of Inflammatory Monocyte Recruitment Reduces Steatohepatitis and Liver Fibrosis. *Hepatology*. 2017 Sep 21. doi: 10.1002/hep.29544 and Is there a role for adaptive immunity in nonalcoholic steatohepatitis? *World J Hepatol*. 2015 Jul 8; 7(13):1725-9.). I did not succeed to visualize the Figure 3



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PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 38213

Title: Multiomics Biomarkers for the Prediction of Nonalcoholic Fatty Liver Disease Severity

Reviewer's code: 01560058

Reviewer's country: Japan

Science editor: Ya-Juan Ma

Date sent for review: 2018-02-01

Date reviewed: 2018-02-10

Review time: 9 Days

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"/> 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 type="checkbox"/> Plagiarism	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor		<input type="checkbox"/> No	<input type="checkbox"/> Major revision
		BPG Search:	
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input type="checkbox"/> No	

COMMENTS TO AUTHORS

In this review article, the authors summarized novel findings obtained from various omics studies on the pathogenesis of nonalcoholic fatty liver disease (NAFLD). The Reviewer considers that this article allow readers to gain new insights into the molecular pathogenesis of NAFLD.



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PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 38213

Title: Multiomics Biomarkers for the Prediction of Nonalcoholic Fatty Liver Disease Severity

Reviewer's code: 02942902

Reviewer's country: Japan

Science editor: Ya-Juan Ma

Date sent for review: 2018-02-01

Date reviewed: 2018-02-10

Review time: 9 Days

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 type="checkbox"/> Plagiarism	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor		<input type="checkbox"/> No	<input type="checkbox"/> Major revision
		BPG Search:	
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input type="checkbox"/> No	

COMMENTS TO AUTHORS

The authors summarized the OMICS-based findings regarding NAFLD/NASH. I consider this paper is interesting and informative. Comments: 1) They mentioned that "Rigorous steps that must include validation and replication are mandatory before utilizing OMICs biomarkers in diagnostics to identify patients at risk of advanced disease, including liver cancer." I would like to know whether they consider that the risk of disease severity (fibrosis) was equal to that of HCC development. (Is it unnecessary to take into account some additional conditions in relation to the development of NASH-related HCC?) 2) Perhaps due to the capacity of my PC, I could not visualize the Figure 3. Kindly scale down the file size.



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PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 38213

Title: Multiomics Biomarkers for the Prediction of Nonalcoholic Fatty Liver Disease Severity

Reviewer's code: 02942798

Reviewer's country: Slovakia

Science editor: Ya-Juan Ma

Date sent for review: 2018-02-01

Date reviewed: 2018-02-11

Review time: 10 Days

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"/> 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 type="checkbox"/> Plagiarism	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor		<input type="checkbox"/> No	<input type="checkbox"/> Major revision
		BPG Search:	
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input type="checkbox"/> No	

COMMENTS TO AUTHORS

Dear sir, thank you to select me for reviewing manuscript: Pirola JC, Sookoian S. Multiomics Biomarkers for the Prediction of Nonalcoholic Fatty Liver Disease Severity. Paper is well written. Only 2 technical changes are needed: 1) I can not open Figure 3 2) Please edit the references as recommended by World Journal of Gastroenterology. My final decision is acceptance for publication in World Journal of Gastroenterology.