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ESPS PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

ESPS manuscript NO: 16527

Title: Cardiovascular risk across the histological spectrum and the clinical manifestations of non-alcoholic fatty liver disease: an update

Reviewer's code: 01806327

Reviewer's country: Spain

Science editor: Ya-Juan Ma

Date sent for review: 2015-01-22 09:52

Date reviewed: 2015-01-30 16:52

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

This is a comprehensive and very well written Review on the relationships between NAFLD and cardiovascular risk. I recommend publication as is.



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Name of journal: World Journal of Gastroenterology

ESPS manuscript NO: 16527

Title: Cardiovascular risk across the histological spectrum and the clinical manifestations of non-alcoholic fatty liver disease: an update

Reviewer's code: 02462668

Reviewer's country: Czech Republic

Science editor: Ya-Juan Ma

Date sent for review: 2015-01-22 09:52

Date reviewed: 2015-02-24 22:41

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 checked="" 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 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 review is concisely written and contains recent clinical data. I would recommend only minor changes in the manuscript. 1. I would recommend changing the Short title (should correspond more with the main title). 2. On the page 5/37, there is no citation for fibrosis stage (Is it according to NAFLD activity score?). 3. There are two different terms used for the same enzyme - gamma-glutamyltransferase (GGT) and gamma-glutamyl transpeptidase (GTT), I would recommend unifying it. Also, abbreviations oGTT and OGTT should be unified and HDL-C should be explained in the text. 4. Adding a table summarizing the clinical trials mentioned in the manuscript could increase readability.



ESPS PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology
ESPS manuscript NO: 16527
Title: Cardiovascular risk across the histological spectrum and the clinical manifestations of non-alcoholic fatty liver disease: an update
Reviewer's code: 00034635
Reviewer's country: Spain
Science editor: Ya-Juan Ma
Date sent for review: 2015-01-22 09:52
Date reviewed: 2015-02-08 18:08

Table with 4 columns: CLASSIFICATION, LANGUAGE EVALUATION, SCIENTIFIC MISCONDUCT, CONCLUSION. It contains checkboxes for various review criteria like 'Grade A: Excellent', 'Priority publishing', 'Google Search', etc.

COMMENTS TO AUTHORS

This is a comprehensive well-structured review of CV risk in NAFLD. NAFLD is a recognized independent risk factor for cardiovascular disease and mortality. While this review provide a modern review on the association between NAFLD and CV risk, the treatment option, mainly focused on statins is less evidence-based, and thus far, there is no available drugs with long-term safety and efficacy to treat NAFLD. I would like authors to address the following concerns about the manuscript. 1. Do they suggest to routinely measure arterial stiffness in all subjects with NAFLD? If so, would they recommend statins for patients with significant AS?. Please, discuss current guidelines recommendation on statin use. 2. Readers may deduce that NASH result from simple steatosis through a time progression, however the multiple parallel hit theory on NASH pathogenesis suggests that NASH may occur directly in many individuals without previous bland steatosis- 3. It is mentioned that there are available drugs for NAFLD and references 54-56 (from the same group) are cited to support this comment. Actually, there is no approved drug for NASH and well-designed clinical trials on statin use are lacking. In a well-designed controlled trial, vitamin E provided benefit



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in NASH, however, this long-term use of this compound may increase the risk for hemorrhagic stroke and for prostate cancer. In addition, long-term use vitamin E decrease survival. Therefore, it should be clearly stated that evidence for efficacy of statins in NAFLD is poor and large good quality clinical trials should be undertaken until these drugs can be recommended. Meanwhile, their use should be consistently aligned with the current guidelines. Minor: A list of Acronyms explanation will ease readability.



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ESPS PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

ESPS manuscript NO: 16527

Title: Cardiovascular risk across the histological spectrum and the clinical manifestations of non-alcoholic fatty liver disease: an update

Reviewer's code: 00182901

Reviewer's country: South Korea

Science editor: Ya-Juan Ma

Date sent for review: 2015-01-22 09:52

Date reviewed: 2015-02-09 16:19

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

This review paper is written nicely and concisely with extensive up-to-date information and the reviewer hardly finds a point to be added. The reviewer only recommend that the author would provide some tables that summarize the clinical trials they introduced in the text, or figures that represent schematic pathophysiologic mechanisms or interrelationship of each risk factors in order to help readers understand.



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ESPS PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

ESPS manuscript NO: 16527

Title: Cardiovascular risk across the histological spectrum and the clinical manifestations of non-alcoholic fatty liver disease: an update

Reviewer's code: 00053485

Reviewer's country: Japan

Science editor: Ya-Juan Ma

Date sent for review: 2015-01-22 09:52

Date reviewed: 2015-01-25 10:32

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 type="checkbox"/> Grade C: Good	<input type="checkbox"/> Grade C: A great deal of language polishing	<input type="checkbox"/> Duplicate publication	<input checked="" type="checkbox"/> Rejection
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<input checked="" 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 this report, Vasiliou et al. reviewed beneficial effects of lifestyle measures and statin treatment on cardiovascular disease in patients with NAFLD/NASH. Although the authors summarized a number of papers, difference from previous reviews is unclear. Major comments 1. There are a lot of review papers are recently published regarding cardiovascular risk in patients with non-alcoholic fatty liver disease (Eur J Gastroenterol Hepatol. 2015 in press, J Tehran Heart Cent. 2014 Jan 12;9(1):1-8., Arterioscler Thromb Vasc Biol. 2014 Jun;34(6):1155-61.). In addition, a similar review has been published in World J Gastroenterol. (World J Gastroenterol. 2014 Feb 21;20(7):1724-45.). The authors have to cite these previous reviews and emphasize the difference from previous reviews. 2. Addition of table and figures will be able to make this paper easier to understand.