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PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology
Manuscript NO: 35349
Title: Nonalcoholic fatty liver disease: Evolving paradigms
Reviewer's code: 02890067
Reviewer's country: Croatia
Science editor: Ze-Mao Gong
Date sent for review: 2017-07-18
Date reviewed: 2017-07-19

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"/> 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 type="checkbox"/> No	

COMMENTS TO AUTHORS

The manuscript is good and suitable for publication



PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology
Manuscript NO: 35349
Title: Nonalcoholic fatty liver disease: Evolving paradigms
Reviewer’s code: 02926997
Reviewer’s country: Iran
Science editor: Ze-Mao Gong
Date sent for review: 2017-07-18
Date reviewed: 2017-07-22

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 Associate Editor, Thank you for sending this review. This review focus on the updates on epidemiology, pathogenesis, diagnosis and treatment of NAFLD. The manuscript is well written.



PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology
Manuscript NO: 35349
Title: Nonalcoholic fatty liver disease: Evolving paradigms
Reviewer’s code: 03647461
Reviewer’s country: United States
Science editor: Ze-Mao Gong
Date sent for review: 2017-07-18
Date reviewed: 2017-07-29

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 checked="" 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"/> Plagiarism	<input 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

Please adjust the grammatical and spelling errors pointed out in the manuscript. Also, define terminologies as they are encountered for the first time, hence the appropriate usage of their acronyms can follow. For example, the same way you correctly defined and abbreviated NAFLD in the abstract, you should also do the same for MetS the first time it is mentioned in the abstract. Furthermore, the first-time IR was mentioned in its acronym form on page 3, it was not spelled out; but it was later spelled out on page 5. It should be the other way around. It is not clear when you cite epidemiological figures whether you are referring to US or world populations. For example, you stated: "In the general population, the prevalence of NAFLD has been reported to widely range from 6.3% to 51% related to the different population/ethnicity evaluated as well as to the diagnostic methods utilized to assess the deposition of liver fat content [43]". Please be specific! Also, please revise the statement "At variance with what was found in the general population, women with T2D are exposed to the same risk of NAFLD as men,



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indicating that T2D abrogates that protection from developing NAFLD which is usually provided to ladies by their hormonal profile and/or set of chromosomes [7, 14].? Please revise this phrase." I have modified this sentence to state what I think you meant. if not, please revise as it does not convey any meaningful message as it currently stands. On page 9, the assertion that "Steatosis, namely a minimal threshold of 5% hepatocytes containing fat droplets." - is the 5% by weight of the liver? There also seems to be a discrepancy between the above 5% hepatocytes mentioned in the manuscript and the sentence on page 10 that states "The main advantages of ultrasonography are its safety, low cost, wide availability, the overall scanning of abdominal organs and the accuracy for the diagnosis of steatosis affecting >10% of hepatocytes". This statement implies that ultrasound cannot be used for determining steatosis in less than 10% of the affected hepatocytes. Again, there is a discrepancy in relating the word "accuracy" between the statements "The main advantages of ultrasonography are its safety, low cost, wide availability, the overall scanning of abdominal organs and the accuracy for the diagnosis of steatosis affecting >10% of hepatocytes" and "The main limitations of ultrasonography are its inaccuracy in differentiating steatosis from fibrosis, the issues with morbid obese individuals, and its operator and machinery dependency". The standard abbreviation or acronym for colorectal cancer is "CRC" and not "CCR".



PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 35349

Title: Nonalcoholic fatty liver disease: Evolving paradigms

Reviewer's code: 03665102

Reviewer's country: Italy

Science editor: Ze-Mao Gong

Date sent for review: 2017-07-18

Date reviewed: 2017-08-01

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"/> 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 checked="" type="checkbox"/> No	<input checked="" 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 checked="" type="checkbox"/> No	

COMMENTS TO AUTHORS

This manuscript by Lonardo et al. is a comprehensive review on Nonalcoholic Fatty Liver Disease, thoroughly exploring all the issues concerning this emerging global health problem. The manuscript is well written and conceived. I only have few comments: - the acronyms should be written in extenso when the authors mention them for the first time - page 6, last paragraph: I suggest to better point out this topic on nuclear receptors to their role in the pathogenesis of NAFLD - page 10, second paragraph, line 5: I suggest to better describe the SAF score, because the the authors' comments are not easily understandable. A table may be added to compare the Kleiner's classification and the FLIP algorithm - page 10, fourth paragraph: in several studies and guidelines, US is considered an imaging technique with limited sentitivity in the detection of liver steatosis, since it does not reliably detect steatosis when <20-25%. The ability of US to detect liver steatosis when >10% is deducted from the results of a recent study by Ballestri et al, but these results should be confirmed. Please comment - after the



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diagnostic section, I suggest to rearrange the layout as follow: 1. NAFLD: clinico-laboratory features; 2. NAFLD as multisystem disease; 3. NAFLD and T2D; 4. NAFLD: a large spectrum of clinical associations; 5. principles of treatment - page 14, third paragraph: the topic on NOSA is not strictly linked to IR and hyperuricemia. If the authors prefer, they should address this issue in another paragraph. Furthermore, as laboratory features, at least liver function tests, lipid profile, iron parameters should be evaluated page 18, second paragraph: please specify that it's a phase 2 randomized study - table 1 should be removed - the manuscript should be revised for typo



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PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology
Manuscript NO: 35349
Title: Nonalcoholic fatty liver disease: Evolving paradigms
Reviewer's code: 03478530
Reviewer's country: Brazil
Science editor: Ze-Mao Gong
Date sent for review: 2017-07-18
Date reviewed: 2017-08-07

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
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		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input checked="" type="checkbox"/> No	

COMMENTS TO AUTHORS

The theme presented in the manuscript is relevant in scientific community, extremely interesting and current, and the authors described relevant aspects with fluidity. Some minor revisions should be making.