Name of journal: *World Journal of Gastroenterology*

Manuscript NO: 74800

Title: Metabolic Aspects of Hepatitis C Virus

Provenance and peer review: Invited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer’s code: 00159278

Position: Editorial Board

Academic degree: MD, PhD

Professional title: Professor

Reviewer’s Country/Territory: Romania

Author’s Country/Territory: Egypt

Manuscript submission date: 2022-01-06

Reviewer chosen by: AI Technique

Reviewer accepted review: 2022-01-07 07:46

Reviewer performed review: 2022-01-17 19:51

Review time: 10 Days and 12 Hours

<table>
<thead>
<tr>
<th>Scientific quality</th>
<th>[ ] Grade A: Excellent</th>
<th>[ ] Grade B: Very good</th>
<th>[ Y] Grade C: Good</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>[ ] Grade D: Fair</td>
<td>[ ] Grade E: Do not publish</td>
<td></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Language quality</th>
<th>[ ] Grade A: Priority publishing</th>
<th>[ Y] Grade B: Minor language polishing</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>[ ] Grade C: A great deal of language polishing</td>
<td>[ ] Grade D: Rejection</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Conclusion</th>
<th>[ ] Accept (High priority)</th>
<th>[ ] Accept (General priority)</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>[ ] Minor revision</td>
<td>[ Y] Major revision</td>
</tr>
</tbody>
</table>

| Re-review          | [ Y] Yes | [ ] No |

| Peer-reviewer      | Peer-Review: [ Y] Anonymous | [ ] Onymous |
SPECIFIC COMMENTS TO AUTHORS

I was asked to review the paper entitled: “Metabolic Aspects of Hepatitis C Virus”. It is an interesting paper, however I think there are some things that can be improved; Major comments: 1. Page 5, last paragraph: You say “In this context, a high prevalence of cirrhosis and non-SVR was observed among patients with diabetes and CHC”. It should be mentioned that the rate of SVR is indeed lower in patients with insulin resistance, but only in Interferon based treatment. It seems that SVR rates following DAA are not influenced by IR (PMID: 30024489) 2. Page 6, end of first paragraph you say: “Consequently, further evidence from long-term follow-up studies is still required to determine if successful eradication of HCV can help to ameliorate IR and improve glycemic control and clinical outcomes in patients with established DM2.28, 29”. Several studies have proven better control of DM and amelioration of IR following SVR and I think they should be discussed (DOI: 10.1080/14787210.2018.1505500; DOI: 10.1016/S1665-2681(19)31578-9; https://doi.org/10.1371/journal.pone.0209216) 3. Page 7, first paragraph, you say: “Studies reported that steatosis may improve and even disappear following successful antiviral treatment with interferon and ribavirin; however, data on direct acting antivirals are still limited.34, 35”. References you quoted refer to peg IFN treatment only, not DAA. Suggested references that should be discussed: DOI: 10.15761/JTS.1000194; doi: 10.1097/MEG.0000000000001106; https://doi.org/10.1038/s41598-021-93881-7; https://doi.org/10.1371/journal.pone.0209615 Minor comments: 1. Paragraph with Insulin resistance, you say: “IR is associated with HCV infection in up to 80% of cases”, I think that correct wording is “HCV infection is associated with IR in up to 80% of cases” 2. English polishing is needed to avoid awkward wording such as: -  page 5, first
paragraph: “the systemic burden of hepatitis C infection surpasses its liver disease burden because it is a major burden on the overall health of the human body, causing increased economic burden to patients and healthcare systems” - page 6, second paragraph: “Not only are IR and DM are more prevalent in the course of HCV infection...” - page 8, last paragraph: “Plausible explanations include the feasibility of adipose tissue to promote fatty substrates....” - page 9, last paragraph: “HCV infection can be an independent risk factor for increased carotid IMT49 and cerebrovascular deaths50, as reported in many types of study; .....” - page 11, second paragraph: “There is a high incidence rate of sarcopenia, of up to 70%, in patients with cirrhosis. Dasarathy S. Cause and management of muscle wasting in chronic liver diseas.”
Name of journal: World Journal of Gastroenterology
Manuscript NO: 74800
Title: Metabolic Aspects of Hepatitis C Virus
Provenance and peer review: Invited Manuscript; Externally peer reviewed
Peer-review model: Single blind
Reviewer’s code: 05769618
Position: Peer Reviewer
Academic degree: PhD
Professional title: Lecturer
Reviewer’s Country/Territory: Indonesia
Author’s Country/Territory: Egypt
Manuscript submission date: 2022-01-06
Reviewer chosen by: AI Technique
Reviewer accepted review: 2022-01-26 12:11
Reviewer performed review: 2022-02-03 08:21
Review time: 7 Days and 20 Hours

<table>
<thead>
<tr>
<th>Scientific quality</th>
<th>Grade A: Excellent</th>
<th>Grade B: Very good</th>
<th>Grade C: Good</th>
<th>Grade D: Fair</th>
<th>Grade E: Do not publish</th>
</tr>
</thead>
<tbody>
<tr>
<td>Language quality</td>
<td>Grade A: Priority publishing</td>
<td>Grade B: Minor language polishing</td>
<td>Grade C: A great deal of language polishing</td>
<td>Grade D: Rejection</td>
<td></td>
</tr>
<tr>
<td>Conclusion</td>
<td>Accept (High priority)</td>
<td>Accept (General priority)</td>
<td>Minor revision</td>
<td>Major revision</td>
<td>Rejection</td>
</tr>
<tr>
<td>Re-review</td>
<td>Yes</td>
<td>No</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Peer-reviewer</td>
<td>Peer-Review: Yes</td>
<td>Anonymous</td>
<td>Onymous</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>
SPECIFIC COMMENTS TO AUTHORS
• in Abstract section, author mentioned insulin resistance in Keywords section, however, reviewers do not identify the explanation of insulin resistance does not found in the abstract section, pleases clarified and revised • author should mention the full name and abbreviation of HCC in the second paragraph of abstract • please mention the VLDL by full name in the page number 8 • page number 9. Please add vitamin D by the role of Vitamin D • please mention more detail the acknowledgments • please add the DOI in the reference number 10, 17, 20, 21, 22, 27, 32, 38, 4153, • please make sure the graphical abstract is more visualize and more attractive. The current version is not well organized, please fixed, such as iron metabolism reproductive status and menopause and insulin resistance does not have figures.
PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 74800

Title: Metabolic Aspects of Hepatitis C Virus

Provenance and peer review: Invited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer’s code: 05562744

Position: Editorial Board

Academic degree: FACS, MD, PhD

Professional title: Professor, Senior Scientist

Reviewer’s Country/Territory: Turkey

Author’s Country/Territory: Egypt

Manuscript submission date: 2022-01-06

Reviewer chosen by: AI Technique

Reviewer accepted review: 2022-01-27 00:41

Reviewer performed review: 2022-02-03 10:07

Review time: 7 Days and 9 Hours

<table>
<thead>
<tr>
<th>Scientific quality</th>
<th>[ ] Grade A: Excellent</th>
<th>[ ] Grade B: Very good</th>
<th>[ ] Grade C: Good</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>[ ] Grade D: Fair</td>
<td>[ ] Grade E: Do not publish</td>
<td></td>
</tr>
<tr>
<td>Language quality</td>
<td>[ ] Grade A: Priority publishing</td>
<td>[ ] Grade B: Minor language polishing</td>
<td></td>
</tr>
<tr>
<td></td>
<td>[ ] Grade C: A great deal of language polishing</td>
<td>[ ] Grade D: Rejection</td>
<td></td>
</tr>
<tr>
<td>Conclusion</td>
<td>[ ] Accept (High priority)</td>
<td>[ ] Accept (General priority)</td>
<td></td>
</tr>
<tr>
<td></td>
<td>[ ] Minor revision</td>
<td>[ ] Major revision</td>
<td>[ ] Rejection</td>
</tr>
<tr>
<td>Re-review</td>
<td>[ ] Yes</td>
<td>[ ] No</td>
<td></td>
</tr>
<tr>
<td>Peer-reviewer</td>
<td>Peer-Review: [ ] Anonymous</td>
<td>[ ] Onymous</td>
<td></td>
</tr>
</tbody>
</table>
SPECIFIC COMMENTS TO AUTHORS
Title: well presented  Abstract: needs language revision  Introduction: generally well written but few grammatical errors are present  The present study is a minireview summarizing the metabolic effects of HCV infection. The quality of the article is very well but language is the main problem and needs revision