Response Letter

Dear Editor and Reviewers,

We thank both the editor and reviewers for their valuable comments and constructive suggestions and are grateful for the insightful comments and valuable improvements to our paper. We have substantially revised the manuscript and incorporated the suggestions from all reviewers to improve our manuscript. We believe that the manuscript has been substantially improved. The revised contents of the manuscript are highlighted with yellow color in the revised manuscript. Please see below, in blue, for a point-by-point response to the reviewers’ comments and concerns.

Reviewer(s)' Comments to Author:

Dear Authors, I have reviewed your manuscript titled "Function of macrophage-derived exosomes in chronic liver disease from etiology to treatment" and found it to be a well-structured and informative mini-review in the field of hepatology. Your comprehensive coverage of the topic is commendable, and I believe your work has the potential to contribute significantly to our understanding of the role of macrophage-derived exosomes in liver diseases. I would like to acknowledge the strengths of your manuscript, including its comprehensiveness, clarity of writing, and scientific rigor. You have provided a balanced perspective on the current state of knowledge in this area, and your incorporation of illustrative figures enhances the clarity of the content.

Response: We feel great thanks for your professional review work on our article. As you are concerned, there are several problems that need to be addressed. According to your nice suggestions, we have made extensive corrections to our previous draft, the detailed corrections are listed below.
1. **In-Text Citations:** Please ensure that in-text citations are specific and included where needed. This will help support your statements with appropriate references, adding credibility to your review.

Response: We very much appreciate your suggestion. We have reviewed and altered the citation of references in order to make it more impartial and worthwhile. In addition, we have modified the figures as new research was included and also updated the citation of references in the figure captions.

2. **Discussion on Clinical Applications:** While you briefly mention the therapeutic potential of macrophage-derived exosomes, consider expanding this section to include discussions on clinical trials, translational challenges, and the future outlook for clinical applications. This will provide readers with a more comprehensive view of the field's clinical relevance.

Response: Thank you for your considerate comments. As suggested, we develop a discussion on the promising clinical applications of macrophage-derived exosomes in the treatment of liver diseases in the conclusion part of the article. Although exosomes are potential biotherapeutic and drug delivery vectors with the advantages of low immunogenicity and evasion of macrophage phagocytosis, there are drawbacks such as poor stability and low delivery efficiency, and therefore there is still a gap in clinical trials of macrophage-derived exosomes for the treatment of CLD. However, it is encouraging that with the advancement of surface modification and nanotechnology for targeted delivery, the conversion of macrophage-derived exosomes from theoretical research to clinical application can become a reality. Therefore, in the future, the research on macrophyge-
derived exosomes should gradually shift to clinical applications to provide more significant value in solving the global liver disease problem.

3. **Incorporating Recent Developments:** Given the rapidly evolving nature of exosome research, consider including any recent key developments or emerging trends in the field to keep the review up-to-date.

   Response: We are grateful for your expert suggestion. We revised this article after looking through the most recent material. As of October 2023, we have revised seven publications, which have reference numbers 72, 73, 74, 81, 87, 91, and 98. Regarding the specifics of the citations, we have highlighted them in the text.

   I appreciate your efforts in addressing these suggestions in your revised manuscript. Overall, I recommend your work for publication, as it contributes significantly to our understanding of the role of macrophage-derived exosomes in liver disease etiology and therapy.

   Response: We really appreciate your approval of our work. We trust these revisions will address the concerns and that the manuscript is now suitable for publication in *World Journal of Hepatology*.
Company editor-in-chief Comments to Author:

I have reviewed the Peer-Review Report and the full text of the manuscript, all of which have met the basic publishing requirements of the World Journal of Hepatology, and the manuscript is conditionally accepted. I have sent the manuscript to the author(s) for its revision according to the Peer-Review Report, Editorial Office’s comments and the Criteria for Manuscript Revision by Authors.

Response: We feel great appreciation for your professional review work on our manuscript. As you are concerned, there are several problems that need to be addressed. According to your expert suggestions, we have made appropriate modifications to the previous draft, the detailed corrections are listed below.

The quality of the English language of the manuscript does not meet the requirements of the journal. Before final acceptance, the author(s) must provide the English Language Certificate issued by a professional English language editing company. Please visit the following website for the professional English language editing companies we recommend: https://www.wjgnet.com/bpg/gerinfo/240.

Response: Thanks for your expert comment. Based on your suggestions, we sent our revised manuscript to a professional English language editing company to polish the manuscript further and received a new language certificate.

Before final acceptance, when revising the manuscript, the author must supplement and improve the highlights of the latest cutting-edge research results, thereby further improving the content of the manuscript. To this end, authors are advised to apply a new tool, the Reference Citation Analysis (RCA).
RCA is an artificial intelligence technology-based open multidisciplinary citation analysis database. In it, upon obtaining search results from the keywords entered by the author, "Impact Index Per Article" under "Ranked by" should be selected to find the latest highlight articles, which can then be used to further improve an article under preparation/peer-review/revision. Please visit our RCA database for more information at: https://www.referencecitationanalysis.com/.

Response: We are grateful for your suggestion. With the help of RCA, we found the latest literature and improved the content of the article. As of October 2023, we have supplemented seven publications, which the reference numbers is 72, 73, 74, 81, 87, 91, and 98. Regarding the specifics of the citations, we have highlighted them in the text.

Uniform presentation should be used for figures showing the same or similar contents; for example, “Figure 1Pathological changes of atrophic gastritis after treatment. A: ...; B: ...; C: ...; D: ...; E: ...; F: ...; G: ...”. Please provide decomposable Figures (in which all components are movable and editable), organize them into a single PowerPoint file. Please check and confirm whether the figures are original (i.e. generated de novo by the author(s) for this paper). If the picture is ‘original’, the author needs to add the following copyright information to the bottom right-hand side of the picture in PowerPoint (PPT): Copyright ©The Author(s) 2023.

Response: We very much appreciate your suggestion. We have modified the figures so that all its components are movable and editable, and organized it into a PowerPoint file (87924-Image File-revision. pptx). After our careful confirmation, we guarantee that all these figures are original. We trust these revisions will address the concerns and that the manuscript is now suitable for publication in World Journal of Hepatology.
Name of journal: World Journal of Hepatology
Manuscript NO: 87924
Title: Function of macrophage-derived exosomes in chronic liver disease: From etiology to treatment
Journal Editor-in-Chief/Associate Editor/Editorial Board Member: Amany Sholkamy
Country/Territory: Egypt
Editorial Director: Jia-Ping Yan
Date accepted review: 2023-10-20 08:43
Date reviewed: 2023-10-20 09:49
Review time: 1 Hour

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<tr>
<th>SCIENTIFIC QUALITY</th>
<th>LANGUAGE QUALITY</th>
<th>CONCLUSION</th>
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<tr>
<td>[ Y] Grade B: Very good</td>
<td>[ ] Grade B: Minor language polishing</td>
<td>[ ] High priority for publication</td>
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<tr>
<td>[ ] Grade C: Good</td>
<td>[ ] Grade C: A great deal of language polishing</td>
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<td>[ ] Grade D: Fair</td>
<td>[ ] Grade D: Rejected</td>
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JOURNAL EDITORIAL BOARD COMMENTS TO AUTHORS
Exosomes are one of the trending topics in medicine and hepatology. Exosomes are among the proposed future targets in diagnosis and treatment of different diseases. Disorders like HCC, NAFLD and hepatic fibrosis (of whatever etiology) are the mostly targeted. Regarding the title; it is better to put the word "pathogenesis" instead of the word "etiology". Exosomes are playing in the pathogenesis of disease. The discussion of this subject in the present minireview was informative and comprehensive. The discussion covered the most important areas in this field. References are too many. the minireview is about 18 pages. References are 22pages. Figures are of informative and of good quality and properly lied down. So long it is not referenced or quoted from another source; I suppose it is original. Kindly state if not. The abbreviation HF” in medicine in general us commonly used for heart failure. So, kindly use "liver fibrosis at least in the subtitle. There should be a section of the abbreviations.
Reply: Thanks for your comment, we had revised the manuscript according to your revision.