

## PEER-REVIEW REPORT

**Name of journal:** *World Journal of Diabetes*

**Manuscript NO:** 80553

**Title:** Future applications of exosomes delivering resolvins and cytokines in facilitating diabetic foot ulcer healing

**Provenance and peer review:** Invited manuscript; Externally peer reviewed

**Peer-review model:** Single blind

**Reviewer's code:** 05438678

**Position:** Peer Reviewer

**Academic degree:** MD

**Professional title:** Doctor

**Reviewer's Country/Territory:** Italy

**Author's Country/Territory:** United States

**Manuscript submission date:** 2022-10-10

**Reviewer chosen by:** AI Technique

**Reviewer accepted review:** 2022-10-10 08:59

**Reviewer performed review:** 2022-10-14 07:52

**Review time:** 3 Days and 22 Hours

Scientific quality	<input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input checked="" type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
Judgment by peer reviewers	Does this manuscript meet the code of ethics standards? <input checked="" type="checkbox"/> Yes <input type="checkbox"/> No Does this manuscript have important novelty? <input checked="" type="checkbox"/> Yes <input type="checkbox"/> No Does this manuscript have important creativity or innovation?

	<input type="checkbox"/> [31] Yes <input type="checkbox"/> [30] No Does this manuscript use reliable research methods? <input type="checkbox"/> [41] Yes <input type="checkbox"/> [40] No Are the manuscript-accompanying data and figures authentic? <input type="checkbox"/> [51] Yes <input type="checkbox"/> [50] No Does this manuscript make scientifically significant conclusions? <input type="checkbox"/> [61] Yes <input type="checkbox"/> [60] No
<b>Language quality</b>	<input type="checkbox"/> Grade A: Priority publishing <input checked="" type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
<b>Conclusion</b>	<input type="checkbox"/> Accept (High priority) <input type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input checked="" type="checkbox"/> Major revision <input type="checkbox"/> Rejection
<b>Re-review</b>	<input checked="" type="checkbox"/> Yes <input type="checkbox"/> No
<b>Peer-reviewer statements</b>	Peer-Review: <input checked="" type="checkbox"/> Anonymous <input type="checkbox"/> Onymous Conflicts-of-Interest: <input type="checkbox"/> Yes <input checked="" type="checkbox"/> No

## SPECIFIC COMMENTS TO AUTHORS

MAJOR COMMENTS: -Section :Inflammation-resolving mediators   should be reduced. Just mention the beneficial effects of resolvins in attenuating inflammation in various inflammatory conditions but the table is not needed. -Section : Future perspectives needs to be more clearly written. The review is related to diabetic foot ulcer healing while the authors report the promising role in the treatment of other inflammatory diseases. Also in this paragraph the table is not necessary.

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**Peer-review model:** Single blind

**Reviewer's code:** 03122873

**Position:** Peer Reviewer

**Academic degree:** DSc, FRCS (Ed), MBBS

**Professional title:** Academic Research, Director, Doctor, Professor

**Reviewer's Country/Territory:** China

**Author's Country/Territory:** United States

**Manuscript submission date:** 2022-10-10

**Reviewer chosen by:** AI Technique

**Reviewer accepted review:** 2022-10-17 03:19

**Reviewer performed review:** 2022-11-01 11:10

**Review time:** 15 Days and 7 Hours

Scientific quality	<input type="checkbox"/> Grade A: Excellent <input checked="" type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
Judgment by peer reviewers	Does this manuscript meet the code of ethics standards? <input checked="" type="checkbox"/> Yes <input type="checkbox"/> No Does this manuscript have important novelty? <input checked="" type="checkbox"/> Yes <input type="checkbox"/> No Does this manuscript have important creativity or innovation?

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<b>Conclusion</b>	<input type="checkbox"/> Accept (High priority) <input checked="" type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection
<b>Re-review</b>	<input type="checkbox"/> Yes <input checked="" type="checkbox"/> No
<b>Peer-reviewer statements</b>	Peer-Review: <input checked="" type="checkbox"/> Anonymous <input type="checkbox"/> Onymous Conflicts-of-Interest: <input type="checkbox"/> Yes <input checked="" type="checkbox"/> No

## SPECIFIC COMMENTS TO AUTHORS

Specific Comments to Authors: It is a valuable review on one important aspect of wound healing in DM patients viz. control of inflammation. As authors rightly pointed out, the healing process is related also to angiogenesis and granulation formation, they could discuss more about these aspects apart from concentrating on exosomes alone.

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**Peer-review model:** Single blind

**Reviewer's code:** 00503405

**Position:** Editorial Board

**Academic degree:** MD, PhD

**Professional title:** Senior Lecturer, Senior Scientist

**Reviewer's Country/Territory:** Hungary

**Author's Country/Territory:** United States

**Manuscript submission date:** 2022-10-10

**Reviewer chosen by:** AI Technique

**Reviewer accepted review:** 2022-10-31 06:46

**Reviewer performed review:** 2022-11-03 18:18

**Review time:** 3 Days and 11 Hours

Scientific quality	<input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input checked="" type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
Judgment by peer reviewers	Does this manuscript meet the code of ethics standards? <input checked="" type="checkbox"/> Yes <input type="checkbox"/> No Does this manuscript have important novelty? <input checked="" type="checkbox"/> Yes <input type="checkbox"/> No Does this manuscript have important creativity or innovation?

	<p><input type="checkbox"/>[31] Yes   <input type="checkbox"/>[30] No</p> <p>Does this manuscript use reliable research methods?</p> <p><input type="checkbox"/>[41] Yes   <input type="checkbox"/>[40] No</p> <p>Are the manuscript-accompanying data and figures authentic?</p> <p><input type="checkbox"/>[51] Yes   <input type="checkbox"/>[50] No</p> <p>Does this manuscript make scientifically significant conclusions?</p> <p><input type="checkbox"/>[61] Yes   <input type="checkbox"/>[60] No</p>
<b>Language quality</b>	<p><input type="checkbox"/> Grade A: Priority publishing   <input checked="" type="checkbox"/> Grade B: Minor language polishing</p> <p><input type="checkbox"/> Grade C: A great deal of language polishing   <input type="checkbox"/> Grade D: Rejection</p>
<b>Conclusion</b>	<p><input type="checkbox"/> Accept (High priority)   <input type="checkbox"/> Accept (General priority)</p> <p><input type="checkbox"/> Minor revision   <input checked="" type="checkbox"/> Major revision   <input type="checkbox"/> Rejection</p>
<b>Re-review</b>	<p><input type="checkbox"/> Yes   <input checked="" type="checkbox"/> No</p>
<b>Peer-reviewer statements</b>	<p>Peer-Review: <input checked="" type="checkbox"/> Anonymous   <input type="checkbox"/> Onymous</p> <p>Conflicts-of-Interest: <input type="checkbox"/> Yes   <input checked="" type="checkbox"/> No</p>

## SPECIFIC COMMENTS TO AUTHORS

The authors summarized the therapeutic effects of the intake of resolvins and cytokins via exosomes in the diabetic foot. The article introduces the pathogenesis of the subtypes of diabetes mellitus in a modern and comprehensible manner, as well as the steps in the development of diabetic foot. Then, the molecular and immunological connections between diabetes and inflammation are properly discussed. In the next chapter, the resolvins involved in reducing inflammation and their immunobiological effects are illustrated. In this section, they also cover a number of diseases that are not the subject of the article. Table 1 is also like this. The presentation of the structure of exosomes and their therapeutic application in DFU is correct. However, Chapters 4.2 and 4.3 repeatedly deviate from the original objective of the article: in the summary of drug delivery and loading possibilities of exosomes, no mention is made of DM and



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DFU. In chapter 4.4, the pros and cons of the use of exosomes are also presented through aspects not related to the topic of the article. Chapters 4.5 and 4.6 are correct. The conclusion is also appropriate. I recommend a DFU-focused rewrite of the chapters indicated above. If there is no literature data on DFU, then the indicated chapters should be combined and shortened. It is also necessary to check the grammar of the English language and correct syntactical errors. A major revision is required.

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**Peer-review model:** Single blind

**Reviewer's code:** 03809896

**Position:** Peer Reviewer

**Academic degree:** PhD

**Professional title:** Assistant Professor

**Reviewer's Country/Territory:** Iran

**Author's Country/Territory:** United States

**Manuscript submission date:** 2022-10-10

**Reviewer chosen by:** AI Technique

**Reviewer accepted review:** 2022-10-31 12:39

**Reviewer performed review:** 2022-11-09 20:00

**Review time:** 9 Days and 7 Hours

Scientific quality	<input checked="" type="radio"/> Grade A: Excellent <input type="radio"/> Grade B: Very good <input type="radio"/> Grade C: Good <input type="radio"/> Grade D: Fair <input type="radio"/> Grade E: Do not publish
Judgment by peer reviewers	Does this manuscript meet the code of ethics standards? <input checked="" type="radio"/> Yes <input type="radio"/> No Does this manuscript have important novelty? <input checked="" type="radio"/> Yes <input type="radio"/> No Does this manuscript have important creativity or innovation?



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<b>Re-review</b>	<input checked="" type="checkbox"/> [Y] Yes <input type="checkbox"/> No
<b>Peer-reviewer statements</b>	Peer-Review: <input checked="" type="checkbox"/> [Y] Anonymous <input type="checkbox"/> Onymous Conflicts-of-Interest: <input type="checkbox"/> Yes <input checked="" type="checkbox"/> [Y] No

## SPECIFIC COMMENTS TO AUTHORS

On manuscript entitled "Future Applications of Exosomes Delivering Resolvins and Cytokines in Facilitating Diabetic Foot Ulcer Healing": In my opinion, the authors have considered all the details of their main topics and have done all the necessary discussions around it well. Moreover, I think that the manuscript is well organized and useful to many interested readers.

## RE-REVIEW REPORT OF REVISED MANUSCRIPT

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**Position:** Peer Reviewer

**Academic degree:** MD

**Professional title:** Doctor

**Reviewer's Country/Territory:** Italy

**Author's Country/Territory:** United States

**Manuscript submission date:** 2022-10-10

**Reviewer chosen by:** Jing-Jie Wang

**Reviewer accepted review:** 2022-11-25 14:01

**Reviewer performed review:** 2022-11-25 14:30

**Review time:** 1 Hour

<b>Scientific quality</b>	<input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input checked="" type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
<b>Language quality</b>	<input checked="" type="checkbox"/> Grade A: Priority publishing <input type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
<b>Conclusion</b>	<input type="checkbox"/> Accept (High priority) <input checked="" type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection
<b>Peer-reviewer</b>	Peer-Review: <input type="checkbox"/> Anonymous <input checked="" type="checkbox"/> Onymous



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statements

Conflicts-of-Interest: [ ] Yes [Y] No

#### **SPECIFIC COMMENTS TO AUTHORS**

I sincerely thank the authors that have responded to all comments and suggestions.