

Response to reviewers' comments

We would like to express our sincere gratitude to the reviewers for their valuable feedback and insightful comments on our manuscript. Your expertise and dedication to providing a thorough evaluation have significantly contributed to the improvement of our work. We appreciate the time and effort you have invested in reviewing our paper, and we have carefully considered your suggestions in order to address any concerns and enhance the overall quality and clarity of the manuscript. Your input has been instrumental in refining our research, and we believe that the final version of the paper will greatly benefit from your constructive criticism. Once again, thank you for your invaluable contribution to the advancement of our research and for helping us shape our manuscript into a more impactful publication.

Reviewer 1 comments	Response
1. As mentioned in abstract one of objection is to identify areas where further research is needed in order to fully understand the effects of adiponectin on DFUs and to establish its safety and efficacy as a treatment for DFUs in the clinical setting... but I could not find any research information or any other future direction research on areas where further research is needed its safety in the manuscript. Author has to cover these areas of future research as mentioned in abstract	Areas for future research have been added before conclusion.
2. As per literature evidence that Adiponectin have a great healing potential on DFU, However it could checked out and discussed about other ie route of administration either by oral or topical if topical route, what is the stability of Adiponectin on site /type of formulations etc..	A paragraph has been added under "Role of adiponectin in wound healing and metabolic conditions: insights from in vitro and in vivo studies" to summarize the findings from in vivo and in vitro studies on the route of administration of adiponectin and its effectiveness in various diseases.
3. The underlying mechanism of Adiponectin healing foot ulcer via different molecular signalling mechanism could have been explained well with more appropriate diagram /figures	Picture Modified to include specific signalling pathway to better the effect of adiponectin
4. What about evidence data about anti-oxidant and anti-bacterial studies? DFU is normally infected with bacterial biofilm.	Antibacterial and antioxidant actions of adiponectin have been elaborated with additional references.
5. In addition to the above comments, I recommend the authors to update more recent relevant publications on treatment of DFU in the references list.	Current & relevant references have been updated
Reviewer 2	
It is a very good review on the importance of adiponectin in the molecular assessment for Diabetic foot ulcer. The facts about its	Thank you

relationship with inflammation, fibrosis, angiogenesis and cellular activities are well described	
DFU is a complicated pathology related to DM, infection, regional and local blood supply etc. To deal with it, the multifactorial influences should be discussed together	A paragraph describing the multifactorial influence in the pathogenesis of DFU is added to enhance coherence of this literature.
Not to target only at one area, one marker. Authors could discuss, to be realistic, whether adiponectin is suitable only as a marker of pathological process.	We have addressed the role of adiponectin both as a marker for DFU as well as being a potential therapeutic agent. Figures 1, 2, 3 support the above fact.