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The Editor,
World Journal of Diabetes
Dear Sir,

Response to reviewers

We are grateful for the comments and contributions of the reviewers. All changes required of us have been implemented and highlighted in red. We are also grateful to the reviewers for the suggested articles; they were of immense benefit in improving the quality of this manuscript.

Reviewer 1

It will be nice if the authors can put a figure outlining the relationship between circadian rhythm changes and its relationship to DM. This has been done

Reviewer 2

- 1) The authors focus on economic burden in the treatment of diabetes worldwide in the second and third paragraph of Introduction. However, this part is unrelated to the topic of this review. The reviewer would recommend to delete descriptions regarding economic burden of diabetes management, or show previous studies reporting the association of circadian rhythm and melatonin with cost-effectiveness of the management of diabetes.-This has been done
- 2) When discussing the association between circadian rhythm dysregulation and diabetes, the authors should mention about the impact of sleep duration/quality

on glycemic control. The reviewer would suggest that the authors add a paragraph concerning sleep and diabetes to the manuscript. -This has been done

3) The role of gut microbiota in the etiology of diabetes mellitus has been a focus of attention. The authors stated that gut hormones activity is under circadian control on page 16. The reviewer would suggest that the authors investigate whether there is any relationship between gut microbiota and circadian system, and discuss the association of brain-gut axis with melatonin in more detail. Xu et al. [J Pineal Res. 2017 May;62(4).] reported that melatonin may be effective for improving gut microbiota dysbiosis. This has been done