

January 25, 2014

Dear Editor,

Please find enclosed the revised manuscript in Word Journal of Diabetes format for your perusal and action.

Title: Toll-like receptor Expression and Signaling in Human Diabetic Wounds

Author(s): Mohan R.Dasu, Sandra Martin

Name of Journal: *World Journal of Diabetes*

ESPS Manuscript #: 5773

Reviewer-1

Q1. What type of diabetes is mentioned? Type 1? Type 2? Both?

Response: Thank you for the comment. All the DM subjects had type 2 diabetes (T2DM). It is now corrected in the text (page#4, line#1).

Q2. Why is not the P value for MyD88 measurements shown? Yet, in the text, the differences detected are referred to as “significant”

Response: Thank you for pointing out the omission. We now included the P value in the text (page#7, line#15).

Q3. Are control and patients age-matched?

Response: Thank you for the comment. All the patients were aged between 40-65 years of age and were matched within ± 8 years difference due to difficulty in obtaining the appropriate study population.

Q4. There are some inconsistencies in the number of patients studied. In “Materials and Methods” four non-diabetic patients are mentioned, whereas in the table 1, five are shown

Response: Thank you for recognizing the typo. We now corrected the subject number and is now consistent across the text.

Q5. Some of the assertions of the manuscript should be softened and simply state that the results shown suggest the involvement of the markers measured in the pathology of the disease. Thus expression like “... confirming the role of...” or “Thus our data for the first time attest to the...”, are overstatements.

Response: Thank you for the suggestion. We now removed “confirming” and “for the first time” and replaced with “implicating” and “suggest” respectively, per reviewer’s recommendation.

Q6. English usage is correct with some slight modifications - hereodimerization should probably be heterodimerization - Some confusion with opening/closing parentheses in the sentence on page 7 “(Figure 2 depicts significantly increased.....P<0.001) may exist. Opening parenthesis may be placed right before the P value.

Response: Thank you for pointing out the typos. We now corrected it to heterodimerization and also corrected the placement of the parenthesis.

Reviewer 2

Q1. In addition, in the “Abstract”, line 8, “IL-6” should be “IL-1?”; and the result shown in Figure 4 was not summarized in the abstract.

Response: Thank you for pointing out the typo. We now corrected it to IL-1 β and included TBARS in the abstract.

Reviewer-3

Q1.Also, the expression of both IL1 and NFkb is known remain under control of TLR3. Yet, this was not assessed.

Response: Thank you for the comment and the reviewer is right. We measured TLR3 mRNA expression in the study and did not see significant differences (0.7 \pm 0.4 vs 1.1 \pm 0.4 fold change vs non-diabetics). In addition, to maintain the focus of the paper on TLR2 and TLR4, this data was not included.

Q2. In fig.3. the values shown are for IL1b, while in the legend these are referred as IL6/

Response: Again, thank you for pointing the typo. We now corrected it to IL-1 β .

All the changes made in the manuscript are indicated with an underline.

Thank you again for publishing our manuscript in the *World Journal of Diabetes*.

Sincerely yours,

Mohan R.Dasu, PhD
Associate Researcher
Department of Dermatology
U C Davis
3301 C Street
Sacramento CA 95816
Email: rkmdasu@gmail.com