

September 17,2014

Dear editor

Please find enclosed the revised manuscript (file name WJD12733revisedMS.doc)

Title: Nociception at the diabetic foot, an uncharted territory

Author: EA Chantelau

Name of the journal: World Journal of Diabetes

ESPS manuscript Nr. 12733

The MS has been revised, all corrections are marked grey. The responses to the reviewers' suggestions are as follows:

1)Reviewer nr 02459617: there were no comments about the manuscript.

2)Reviewer nr 00875981:

a)"It would be useful to define the word 'limen'" - the word as listed on p 9 means just noticeable difference. The thermal sensory limen is part of the QST protocol of the German Research Network on Neuropathic Pain and was quoted as such. The limen of mechanical pain sensation was not considered in the article and, therefore, was not alluded to.

b)The misprint lager was corrected to larger

c)a section on pathophysiology of nerve fibre destruction was not included, because the exact pathomechanisms of diabetic neuropathy are unknown (to the best of my knowledge). At present, a curative or symptomatic treatment of PLDN is not existing.

d)day-to-day reproducibility of DPPPT or CPPPT: this aspect was alluded to on p 12

e)are these tools helpful in assessing disease progression ? Pinprick pain perception threshold (CPPPT) may be potentially useful for measuring disease progression, as it seem to be very sensitive to small reductions in intraepidermal nerve fibre density (Selim et al., new reference [85])- this was added on page 14.

f)comorbidities affecting psychophysical performance may render QST impossible and, thus, are established exclusion criteria for QST studies

Sincerely,

Ernst A. Chantelau, MD, PhD



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