

November 20, 2014

Dear Editor,

Please find enclosed the edited manuscript in Word format (file name: 14229-edited-2.doc).

Title: Relationship between vascular endothelium and periodontal disease in atherosclerotic lesions: Review article

Authors: Marco Aurélio Lumertz Saffi, Mariana Vargas Furtado, Carisi Anne Polanczyk, Márlon Munhoz Montenegro, Ingrid Webb Josephson Ribeiro, Cassio Kampits, Alex Nogueira Haas, Cassiano Kuchenbecker Rösing, Eneida Rejane Rabelo-Silva.

Name of Journal: World Journal of Cardiology

ESPS Manuscript NO: 14229

The manuscript has been improved according to the suggestions of reviewers: **yes**

1. Format has been updated: **yes, offered the postcode**
2. Revision has been made according to the suggestions of the reviewer

Reviewer 1 (00597793)

(1) This review article purports to examine the association of periodontal disease with ASHD. It turns out that the first 2/3 of the article is a review of the formation of atherosclerosis and has nothing to do with periodontal disease. Given that this article is sent to a cardiology journal I do not believe the first 2/3 of this review is needed. Cardiologists know about inflammation. I suggest presenting just the section on periodontal disease with a brief Introduction that periodontal disease is associated with inflammation proteins. The periodontal section is well written.

Authors' response: Thank you very much for a nice suggestion. Our group has worked with several mechanisms related to atherosclerosis. Within these factors, we associated inflammatory markers and endothelial function assessed by flow-mediated dilation. Thus, we believe that our article brought a major revision for understanding the atherosclerotic disease process and the relationship with periodontal disease. Therefore, other professionals, not only cardiologists, will be able to study this relationship reviewed in this article.

Reviewer 2 (02945927)

(1) The aim of the review is find a relationship between Periodontitis and cardiovascular disease. Major comments: You should generate a table indicating every pro-inflammatory cytokines and adhesion molecules altered in an atherosclerotic plaque. Moreover, another table with inflammatory markers and periodontitis, indicating whether increases or decreases.

Authors' response: Were included in Table 1 (Relationship of pro-inflammatory cytokines and adhesion molecules in atherosclerosis) and Table 2 (Relationship of pro-inflammatory cytokines and adhesion molecules in the effect of periodontal disease).

(2) Why do you explain the manner wide the Shear stress? Is important for you review? What's your propose with this review? Can you propose a cartoon with your idea? Minor comments Add

a reference for Shear stress definition It is not necessary explain what is FMD.

Authors' response: Thank you very much for the important comments. Besides the evaluation of inflammatory markers associated with atherosclerosis, our group has evaluated the flow-mediated dilation as in patients with periodontal disease. For explaining all the mechanisms involved, we believe it is important to mention the shear stress when it comes to endothelial dysfunction.

3. References were corrected: yes, added PubMed citation numbers and DOI citation to the reference list and list all authors.

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

Sincerely yours,



Eneida Rejane Rabelo-Silva

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