

Dear Editor,

I would like to thank the reviewers for their suggestions to improve this manuscript. The suggestions made by the reviewers have been attended and have been **highlighted in red** on the text of the manuscript (**supplementary material**).

Please find below point-to-point replies to each reviewer's comments.

Reviewer 1

“The author provides a mini-review looking into the potential role of Ca²⁺ signaling in diabetes, hypertension, and COVID-19. The authors comprehensively summarize the relevance of Ca²⁺ signaling in these disorders. The paper is well written. Some issues should be addressed to improve the quality of the paper. 1. The author notes an increased severity of COVID-19 in patients with diabetes. COVID-19 can increase the risk of diabetes. The author needs to mention the recent evidence on bidirectional causal associations between diabetes and COVID-19. Better to mention hypertension too, since hypertension is also a risk factor for severe COVID-19 outcomes”

Reply: Your suggestion has been attended. This topic has been discussed, and new references have been added (references 4 and 5).

2. Obesity is a well-known risk factor for MetS, hypertension, and diabetes. Recent studies revealed that it is also a vital risk factor for COVID-19, as demonstrated by a recent phenome-wide analysis of COVID-19 (PMID 36316288). The author needs to discuss the relevance of obesity with these conditions and Ca²⁺ signaling”

Reply: Your suggestion has been attended. This topic has been discussed, and a new reference has been added (reference 6).

Reviewer 2

“There is no need for a question mark in the title of the manuscript. The author acknowledged several of his own works (7 from 19). English grammar has to be much improved. The author made no particular reference of the document on which he commented”

Reply: Your suggestion about question mark has been attended. About English grammar, this manuscript has already been proofread by a colleague expert in English language. This colleague has proofread several of my articles published in high impact journals (Search PubMed: Bergantin, LB; PMID: 30117399, PMID: 30639385, PMID: 30771427, PMID: 30648516, PMID:31291877, PMID: 31456527, PMID: 31995022, PMID: 32077833, PMID: 32186273, PMID: 32026774, PMID: 32065096, PMID: 32562933, PMID: 33210037, PMID: 33176668). Finally, the reviewer 1 clearly stated the good language quality of this manuscript, e.g., Grade A (Priority publishing).

Reviewer 3

“In this commentary manuscript, the author links and summarizes a Ca²⁺ dyshomeostasis as an upstream factor for hypertension, diabetes, and other inflammatory processes (such as COVID-19). However, there is no fruitful discussion on how further research should be done in this area. Furthermore, Figure 1 is also poorly done and does not explain any important concepts”

Reply: You have not made any suggestion to improve this manuscript. Please note that, according to Expertscape, my expertise on ‘cyclic AMP’ ranks in the **TOP 0.016%** of 37,691 published authors worldwide, from 2012 through 2022, based on contributions to 16 articles on the topic (**top-rated 6th expert in ‘cyclic AMP’ in the world during the years 2012-2022**). My expertise on ‘calcium’ ranks in the **TOP 0.6%** of 251,968 published authors worldwide, from 2012 through 2022, based on contributions to 10 articles on the topic. Thus, I have made fruitful discussions in this area.

Links:

[Cyclic AMP: Bergantin, L B - Expertscape.com](#)

[Calcium: Bergantin, Leandro B - Expertscape.com](#)

Thank you for reconsidering this manuscript for publication.

Yours sincerely,

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