

Dear Editor,

Thank you very much for timely arranging the review of our manuscript and sending us your comments and the comments of the reviewers. The comments were highly insightful and helped us to improve the manuscript substantially.

We have revised the manuscript by addressing all the comments. All the amendments are highlighted in the revised manuscript. Point by point responses to the reviewers' comments are listed below this letter.

- 1) Some spelling mistakes throughout manuscript. Please revise.

We revise the manuscript and correct the mistakes

- 2) Please revise the sentence "For this reason drugs blockers inflammatory reaction such as IL-1 blockers.."

we reviewed the sentence

- 3) It strongly proposes a figure with the jack/stat signaling pathway and its effect on cytokines' regulation.

we proceeded to add, as suggested, a figure on the interaction cytokines and JAK pathway

- 4) Please provide more agents against IL-1, IL-6 and jack/stat. Moreover a second table with clinical trials (numbers, specification etc) in other inflammatory diseases beside SARS-CoV-2 will be useful.

we added more agents against IL-1, IL-6 to the review as suggested by the reviewer. we added a second table with registered clinical studies on Sars cov 2 infection.

we decided not to add data from the studies in other infectious diseases, because we believe will lead to confusion not being discussed in the review.