

Reviewer #1:

**Scientific Quality:** Grade C (Good)

**Language Quality:** Grade A (Priority publishing)

**Conclusion:** Minor revision

**Specific Comments to Authors:** This is a general summary of the angiotensin-converting enzyme. Its relations with the actual Covid-19 disease are not yet complete clear: you have to specify and discuss it.

Response: Dear reviewer, you are right. It is an up-to-date summary of the angiotensin-converting enzyme 2. ACE2 had been discovered for more than twenty years. The reason that people concern it again and so hot because the ACE2 is the key cell receptor for the SARS-CoV-2 virus. The paradox of the wide use of ACEI/ARB in many diseases and the probable contribution of the likely increased ACE2 after ACEI/ARB to an increased risk of COVID-19 has puzzled clinicians. Regard to the relationship between ACE2 and Covid-19, some studies had been published. But it is an unsettle question and some result is not consistent. Many researches are ongoing. However, we had cited a part of them and taken three big paragraphs to discuss, which is in red in the review. It would help people understand ACE2 with COVID-19. Anyhow, it is very necessary to further understand the characteristics of ACE2 nowadays. So the aim of this review is try to focus on the biological function of ACE2 and its host regulatory factors and discusses its role in 4 main kinds of disease: cardiovascular, kidney and lung diseases and diabetes. Therefore, we now think that we need change the title into the current one “New Perspectives on Angiotensin-Converting Enzyme 2 and Its Related Diseases”. We think this title might be fit better for the whole review.

**Science editor:**

5 Issues raised: (1) The authors did not provide the approved grant application form(s). Please upload the approved grant application form(s) or funding agency copy of any approval document(s);

Response: As suggested, I had uploaded the relevant document in the system now.

(2) The authors did not provide original pictures. Please provide the original figure documents. Please prepare and arrange the figures using PowerPoint to ensure that all graphs or arrows or text portions can be reprocessed by the editor; and

Response: Yes, you are right. In the word document, it is not the original pictures. Now I had uploaded the PPT version in the system now.

(3) PMID and DOI numbers are missing in the reference list. Please provide the PubMed numbers and DOI citation numbers to the reference list and list all authors of the references. Please revise throughout. 6 Recommendation: Conditional acceptance.

Response: We had revised the reference according your suggestion. PubMed numbers, DOI citation numbers and all authors are included in the reference now.