

Response to “ACE-2 downregulation and incidence of severe acute respiratory syndrome-coronavirus-2 (SARS-CoV-2) infection”

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**To the Editor:** We appreciate the questions raised by Drs. Weir, Glen and McDowell in their letter to the Editor referencing our recent paper about SARS-CoV-2 infection in relation to use of angiotensin converting enzyme inhibitors (ACEIs) and angiotensin receptor blockers (ARBs).<sup>1</sup> They propose that inhibiting ACE-2, the receptor by which SARS-CoV-2 enters cells, might be beneficial in preventing infection or severe outcomes and ask whether we could examine the association between use of aliskiren, a direct renin antagonist thought to downregulate ACE-2, and risk of SARS-CoV-2 infection or severe disease. We looked at our data and found that in the population we studied, aliskiren use was so rare that we would not have adequate power to examine this question. In the past year, patients in our healthcare system were dispensed fewer than 30 prescription fills for aliskiren. For comparison, there were about 150,000 medication dispensings for lisinopril, the most commonly used ACEI. The idea that downregulation of ACE-2 could protect against SARS-CoV-2 infection or severe disease is intriguing and warrants further exploration. We would be interested to know if there are other, more commonly used medications that downregulate ACE-2 that could be the subject of future research.

## References

1. Dublin S, Walker R, Floyd JS, Shortreed SM, Fuller S, Albertson-Junkans L, Harrington LB, Greenwood-Hickman MA, Green BB, Psaty BM. Renin-angiotensin-aldosterone system inhibitors and COVID-19 infection or hospitalization: a cohort study. *Am J Hypertens* 2020;hpaa168. doi: 10.1093/ajh/hpaa168. Online ahead of print.

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