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Adrian Post Literature search; Writing; Review and editing, Robin P.F. Dullaart Literature search; Writing; Review and editing, Stephan J.L. Bakker Literature search; Writing; Review and editing

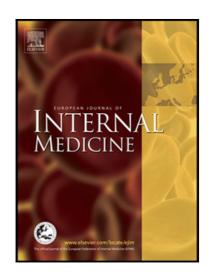
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## Is low sodium intake a risk factor for severe and fatal COVID-19 infection?

Adrian Post<sup>1</sup>, Robin P.F. Dullaart<sup>1</sup> and, Stephan J.L. Bakker<sup>1</sup>

<sup>1</sup>Department of Internal Medicine, University Medical Center Groningen, University of Groningen,

Groningen, the Netherlands.

#### E-mail addresses:

a.post01@umcg.nl; dull.fam@12move.nl; s.j.l.bakker@umcg.nl

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### Corresponding author

Adrian Post

Department of Internal Medicine

University Medical Center Groningen

9700 RB Groningen, the Netherlands

Phone: +316 49653442, Email: a.post01@umcg.nl

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Literature search: Adrian Post, Robin P.F. Dullaart and Stephan J.L. Bakker

Writing: Adrian Post, Robin P.F. Dullaart and Stephan J.L. Bakker

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As of March 16<sup>th</sup>, the number of deaths due to the coronavirus 2019 (COVID-19) outside China has surpassed those in China. Intriguingly, recent calculations that adjust for an incubation period of up to 14 days estimate that the mortality rates from COVID-19 infection in China are nearly three times lower than outside of China<sup>1</sup>. It should be realized that, amongst others differences, China is known for having very high sodium intake compared to other countries in the world <sup>2</sup>. Human pathogenic coronaviruses bind to their target cells through angiotensin-converting enzyme 2 receptor (ACE2), which is expressed in epithelial cells of the lung, intestine, kidney, and blood vessels <sup>3</sup>. Importantly, animal-based studies have demonstrated that the tissue expression of the ACE2 receptor is down-regulated in response to a high dietary sodium intake <sup>4,5</sup>.

Combined, these experimental and epidemiological data led us to hypothesize that low sodium balance may augment cellular damage at a certain virus load and increase the risk of developing severe and fatal COVID-19 infection. If this hypothesis were to be confirmed, it could lead to a conflict because, besides being a target for COVID-19, ACE2 also has anti-inflammatory properties and has been shown to protect against acid-aspiration-induced acute respiratory distress syndrome <sup>6</sup>.

Besides habitual dietary salt intake, more acute changes in sodium balance might also influence ACE2 receptor expression. Intermittent sodium loss, due to either diarrhea, vomiting or perspiration could put patients that acquire COVID-19 infection at higher risk for development of a more severe or fatal course of disease. It seems wise to monitor sodium intake and start with sodium and fluid resuscitation early in the course of more severe COVID-19 infection, and perhaps also to refrain from robust sodium restriction during the current COVID-19 outbreak.

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