# Reply to comment on "Unexpected decline in glycated hemoglobin level after emergency COVID-19 measures in three robust older Japanese women with prediabetes/mild type 2 diabetes"

#### Dear Editor,

Mungmunpuntipantip and Wiwanitkit expressed concerns<sup>1</sup> about the glycated hemoglobin (HbA1c) measurement in our three robust older women who showed a decline in HbA1c after emergency COVID-19 measures.<sup>2</sup> Their first concern was the laboratory's quality control for our HbA1c measurement. The serial analysis of HbA1c value in these three robust older women was performed by a high-quality HbA1c analyzer (HLC-723G11) in our University Hospital. The analysis had sufficiently low imprecision coefficients of variation (CVs), with repeatability imprecision CVs of 1.0% and 0.6% for low and high HbA1c venous samples and between-day imprecision CV of 0.6% for low HbA1c venous samples,<sup>3</sup> which are comparable with, or even lower than the respective imprecision CVs in the report<sup>4</sup> cited in their references.<sup>1</sup> The HbA1c level decreased by 0.3% in each of the three robust older women after emergency COVID-19 measures.<sup>2</sup> Even when adopting the largest error range using venous samples in the report (3.80%),<sup>4</sup> the largest error in the HbA1c decline in our robust older women would be only about  $\pm 0.01\%$  HbA1c. Thus, their concern about the quality control of our HbA1c measurement is unjustified.

They found an increase in blood viscosity in patients with COVID-19, and expressed a second concern about the influence of blood viscosity on HbA1c measurement.<sup>1</sup> However, our three robust older women with unexpected HbA1c decline were, in the first place, not infected with severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2), but just stayed at home during the emergency measures in response to the call by the government to reduce contact with others by at least 70% and by as much as 80%,<sup>5</sup> and refrained from visiting friends with sweets and sugary drinks.<sup>2</sup> Thus, their second concern is also unwarranted.

Older subjects, particularly those with type 2 diabetes mellitus, are vulnerable to COVID-19 as well as the secondary consequences of the pandemic.<sup>6,7</sup> Among older patients with COVID-19, diabetes mellitus, older age  $\geq$ 75 years and male sex were associated with a higher probability of death.<sup>8</sup> Moreover, among patients with both COVID-19 and type 2 diabetes mellitus, HbA1c >8.0%, older age  $\geq$ 65 years and male sex were also associated with a higher probability of death.<sup>9</sup> On the other hand, a higher HbA1c level was reported in non-infected older outpatients with type 2 diabetes after the first emergency COVID-19 measures in Japan, and in those during home confinement related to the COVID-19 lockdown in Italy, probably due to a decrease in physical activity during the measures.<sup>2</sup> Even social distancing itself in the era of the pandemic resulted

in worsening of fasting glucose and HbA1c levels associated with increased body weight in older patients with type 2 diabetes mellitus.<sup>10</sup> Conversely, the first emergency COVID-19 measures in Japan deprived our three robust women of happy times with friends, but brought about an unexpected decline in their HbA1c level.<sup>2</sup>

## **Disclosure statement**

The authors declare no conflict of interest.

#### Data availability statement

Owing to privacy/ethical restrictions data are only available on request.

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