## Comment on: "Effect of vitamin D monotherapy on indices of sarcopenia in community-dwelling older adults: a systematic review and meta-analysis" by Prokopidis et al.

It was interesting to read a synthesis of effects of vitamin D monotherapy on sarcopenia in community-dwelling older adults by Konstantinos Prokopidis et al. (2022).<sup>1</sup> Doctor Prokopidis et al. nicely included 10 randomized controlled trials for meta-analysis on this topic, and most findings reported indicate that vitamin D monotherapy could not improve sarcopenia outcomes in community-dwelling older population. These findings could be understood due to mono-use of vitamin D. and combination of vitamin D with protein intake and exercise would be a better strategy for patients with sarcopenia according to prior experience.<sup>2</sup> We have noticed that the authors tried to extensively confirm the effects of vitamin D monotherapy using a composite outcome (general physical performance), and they also pooled data in an appropriate approach by standardized mean difference (SMD). However, a high heterogeneity  $(I^2 = 71\%)$  may need a further discussion. According to previous evidence, age might be a potential factor due to the associations between them.<sup>3–5</sup>

We did further analysis based on the data set that were reported by Prokopidis et al. (2022),<sup>1</sup> and analysed data in random-effects model with subgroup by mean age. Because older population could be classified into young-old (60 to 69 years old), middle-old (70 to 79 years old) and old-old  $(\geq 80 \text{ years old})$ ,<sup>6</sup> this further analysis took the threshold for subgroup analysis. To depict trend between effects of vitamin D and age, we also used meta-regression. All analysis were carried out by R version 4.1.0 via RStudio version 1.41717. In addition to the reproduced overall pooled results (SMD: -0.02; 95% CI: -0.23 to 0.18), the present analysis found reduced heterogeneity in a subset of middle-old population (Figure 1A). Moreover, our subgroup analysis showed that vitamin D appeared to significantly decrease general physical performance in middle-old population (SMD: -0.15; 95% CI: -0.27 to -0.02), but there was no significant difference between vitamin D monotherapy and placebo in subgroup of young-old population. Result of meta-regression also reflected the phenomenon (particularly the trend in non-linear model due to a lower deviance), although it did not reach statistical significance (Figure 1B). These findings are similar to a previous important study.<sup>7</sup>

In conclusion, this further analysis fosters the understanding of the effects of vitamin D monotherapy for sarcopenia outcome in community-dwelling older population through exploring the influence of age. Vitamin D supplement may have different effect in older population of different age. This hypothesis may explain the divergent conclusion of previous study on vitamin D in preventing falling. Nevertheless, subgroup analysis could not fully explain the heterogeneity in young-old population. Regression would be a better method to identify the association between effects of vitamin D monotherapy and age, while our analysis has no power to find the trend due to very limited numbers of studies included in the meta-regression. Wherefore, large-scale studies are still needed to reveal how the effects of vitamin D on sarcopenia outcomes in community-dwelling older population, and identify which age group should take vitamin D cautiously.

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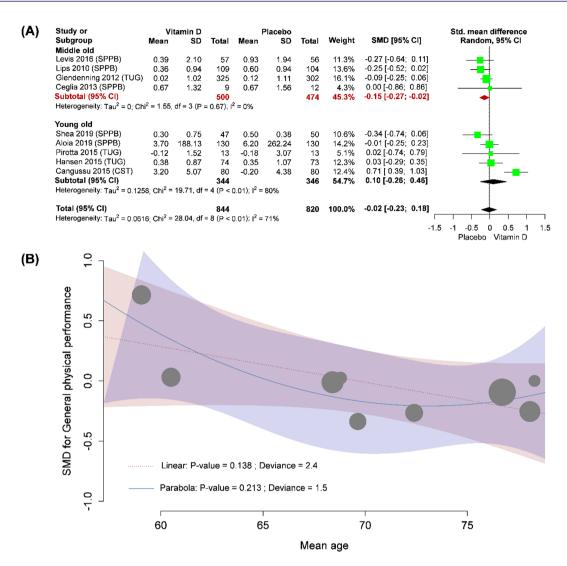


Figure 1 General physical performance (A) between vitamin D and placebo, and (B) association between effects of vitamin D and age. CI, confidence interval; SD, standard deviation; SMD, standardized mean difference.

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