#### ASBMR

# Vitamin D: A single initial dose is not bogus if followed by an appropriate maintenance intake

#### To the Editor:

We read the review by Mazess and colleagues<sup>(1)</sup> in *JBMR Plus* with interest. We have several points for comment and discussion.

First, a bolus dose is a single dose of a substance given over a short period, also called a loading dose.<sup>(2)</sup> However, in the review,<sup>(1)</sup> the term "bolus" dose refers to the nutrient cholecalciferol, vitamin D3 supplement as a longer-term micronutrient, in larger quantities than accepted daily doses with intervals of over a month duration between intakes.

Second, although the authors rightly critique overly long intervals between vitamin D3 supplementation intakes (ie, >1-month intervals), they were mistaken to refer to these as "bolus doses" and so mischaracterize bolus dosing as "bogus."

In those with coronavirus disease(s) (e.g., COVID-19), sepsis or other medical emergencies, achieving and mainteaning serum 25-hydroxyvitamin D [25(OH)D] concentration of above 50 ng/ mL is crucial for achieving proper immune function<sup>(3)</sup> (also toreplete body stores) within days with a single bolus dose of vitamin D<sub>3</sub>, such as 5 to 10 mg (200,000 to 400,000 IU), or within 4 hours using a single, 0.5-mg to 1-mg oral dose of calcifediol (0.014 mg/ kg body weight).<sup>(4)</sup> However, to maintain the boosted serum 25 (OH)D concentrations, D<sub>3</sub> supplements should be taken daily or once a week in larger quantities (taking less frequently than this, is less effective).<sup>(5)</sup>

Third, the review's critique of the Murai study should not be considered a general critique of bolus D3 dosing. This trial resulted in poor clinical outcomes due to the faulty study design of using a bolus dose of D<sub>3</sub> in seriously ill patients. Earlier administration of vitamon D<sub>3</sub> and/or the use of rapidly acting calcifediol, would have been more effective in patients with advanced COVID-19 and in sepsis.<sup>(6)</sup>

Finally, the circulatory 1,25-dihydroxyvitamin D [1,25-(OH)<sub>2</sub>D; calcitriol] levels and 24-hydroxylase enzyme activity used in this review as indicators of immune system functions are not relevant to vitamin D status or to the critical, autocrine and paracrine signaling system of immune cells.

# **Conflict of Interest**

The authors declares no conflicts of interest.

### **Peer Review**

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# References

- Mazess R, Bischoff-Ferrari HA, Dawson-Hughes B. Vitamin D: bolus is bogus—a narrative review. JBMR Plus. 2021;5(12):e10567.
- 2. National Cancer Institute. Definition of the bolus dose. https://www.cancer.gov/search/results?swKeyword=bolus+dose. 2022.
- Quraishi SA, Bittner EA, Blum L, Hutter MM, Camargo CA Jr. Association between preoperative 25-hydroxyvitamin D level and hospitalacquired infections following Roux-en-Y gastric bypass surgery. JAMA Surg. 2014;149(2):112-118.
- Suñé Negre J, Azpitarte IO, Barrios PDA, Hernández Herrero G. Calcifediol soft capsules. 2016. https://patents.google.com/ patent/WO2016124724A1/en. W.W.A.2016-08-11. U.S. Patent WO2016124724A1.
- Mocanu V, Stitt PA, Costan AR, et al. Long-term effects of giving nursing home residents bread fortified with 125 microg (5000 IU) vitamin D(3) per daily serving. *Am J Clin Nutr.* 2009;89(4):1132-1137.
- Murai IH, Fernandes AL, Sales LP, et al. Effect of a single high dose of vitamin D3 on hospital length of stay in patients with moderate to severe COVID-19: a randomized clinical trial. JAMA. 2021;325(11): 1053-1060.

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