



A comparison of capsule and oral spray solution as a method of delivering vitamin D₃ and raising vitamin D status: a wintertime randomised, open-label crossover study

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Vitamin D deficiency is highly prevalent in the UK and Ireland and is defined as a total 25-hydroxyvitamin D (25[OH]D) concentration below 30 nmol/L with respect to bone health⁽¹⁾. Owing to the UK and Ireland's northerly latitudes (50–58 and 51–55°N respectively) as well as the limited range of naturally occurring and fortified dietary sources of vitamin D, supplementation is often regarded as advisable in order to optimise wintertime vitamin D status. Interventions typically use capsules as a peroral method of delivery. This study aimed to compare the efficacy of two forms of supplemental vitamin D₃; liquid capsules or oral spray solution, at increasing total 25(OH)D concentrations in healthy adults.

In total, 22 participants (males $n = 10$ and females $n = 12$) were independently randomised to receive 3000IU (75µg) vitamin D₃ daily for 4 weeks in either a capsule or oral spray form during wintertime (Oct–Feb). Following a 10-week washout, participants crossed-over onto the opposite treatment for a final 4 weeks. Height (cm) was measured at baseline while weight (kg) and fasted blood samples were obtained before and after each supplementation phase. Total 25(OH)D was quantified using LCMS-MS and intact parathyroid hormone (PTH) concentration was measured by ELISA. Dietary vitamin D intake was estimated using a validated food frequency questionnaire⁽²⁾.

Time point Measure	Treatment							
	Capsules				Oral spray			
	Pre Mean	SD	Post Mean	SD	Pre Mean	SD	Post Mean	SD
Age, years	25	6	25	6	25	6	25	6
Weight, kg	72.61	15.42	72.48	15.35	69.84	14.46	69.93	14.65
BMI, kg/m ²	24.79	3.61	24.64	3.70	24.04	3.39	24.02	3.36
Total 25(OH)D, nmol/L	60.78	25.26	88.36 ^a	22.64	58.88	25.44	85.34 ^a	20.77
PTH, g/dL	54.03	30.22	53.80	26.39	49.49	19.53	48.79	20.44

^asignificantly different from pre-intervention $P < 0.05$ (paired t test)

Overall, baseline mean \pm SD total 25(OH)D concentration averaged 59.76 ± 29.88 nmol/L, representing clinical sufficiency. Prior to hypothesis testing, a time by treatment interaction and potential carryover effects were ruled-out ($P = 0.107$ and $P = 0.681$, respectively). Subsequently, analysis of covariance determined that there was no significant difference in mean \pm SD change from baseline, with respect to total 25(OH)D concentrations, between oral spray and capsule supplementation (26.46 ± 23.91 versus 27.58 ± 15.93 nmol/L respectively, $P = 0.995$). Dietary vitamin D intake averaged 6.25 ± 6.24 µg/day, falling short of the current 10µg/day reference nutrient intake. Our findings advocate oral spray vitamin D₃ supplementation as an equally effective alternative to capsules. This may have major implications for micronutrient delivery in those with malabsorption syndromes; as vitamin D₃ administered by oral spray bypasses the intestine via buccal, sublingual and palatal membrane absorption sites in the oral cavity. This supplementation method will also prove advantageous for those with difficulty swallowing such as the elderly, young children and babies.

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1. Ross AC, Manson JE, Abrams SA *et al.* *J Clin Endocrinol Metab.* 2011; **96**:53–58
2. Weir RR, Carson EL, Mulhern MS *et al.* *J Hum Nutr Diet.* 2015; **29**:255–261