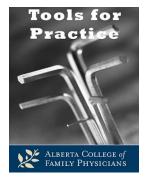
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Vitamin D and Respiratory Tract Infections: Does the sun's vitamin chase the cold?

<u>Clinical Question</u>: Can regular vitamin D supplementation reduce the frequency, duration, or severity of respiratory tract infection (RTI)?

<u>Bottom-line</u>: Regular use of vitamin D does not reduce the frequency, duration, or severity of RTI in western populations. Infrequent benefits seen in a few studies are at high risk of bias and/or involved children with profound deficiency (example 17.5 nmol/L) in developing countries.

# **Evidence:**

- Three systematic reviews<sup>1-3</sup> with 4-11 Randomized Controlled Trials (RCTs) with 1,668-5,660 patients. Vitamin D supplementation:
  - Statistically significantly reduced RTI in two meta-analyses [Odds Ratio 0.64 (0.49-0.84)<sup>1</sup> and 0.58 (0.42-0.81)<sup>2</sup>] but not another [Relative Risk 0.98 (0.93-1.03)].<sup>3</sup>
    - Why the difference? Numerous large biases, especially the former two: Using odds ratios for common problems exaggerates effects, including selectively reported outcomes, combining unrelated studies, including secondary analyses, inconsistent results, and publication bias.<sup>1,2</sup>
  - Examining individual RCTs:
    - Any RTI:
      - 162 US adults: No difference.4
      - 164 Finnish military recruits: No difference.<sup>5</sup>
      - 140 Immuno-compromised patients: Improved non-validated RTI score (not clinically interpretable).<sup>6</sup>
      - 247 Mongolian children with profound vitamin D deficiency (level=17.5 nmol/L): 0.35 less RTI over three months.<sup>7</sup>
    - Cold and Flu (mostly cold):
      - 322 New Zealander adults: No effect in any outcome.<sup>8</sup>
        - This is the highest quality study.
    - Flu:
      - 430 Japanese children mean age 10: No difference.<sup>9</sup>
    - Pneumonia:

- 453 Afghanistan children age <3 years: Reduced risk of one repeat pneumonia but not multiple pneumonias.<sup>10</sup>
- 3,060 Afghanistan children age <1 year: No difference (suggesting earlier results spurious).<sup>11</sup>
- Three RCTs of other conditions looked at RTI secondarily:
  - Two found no difference,<sup>12,13</sup> but the weakest (smallest RCT with grossly under-reported RTI examined retrospectively) found possible reduced cold/flu frequency.<sup>14</sup>

## **Context:**

- Cohort studies suggest patients with low vitamin D levels get more RTI. 15
  - However, low vitamin D status is associated with many ills from weight gain to mortality but vitamin D RCTs rarely find clinical improvements.<sup>16,17</sup>
  - Vitamin D is likely a surrogate marker for ill health.<sup>16</sup>
- Cold prevention likely lies with physical interventions like hand-washing. 18

#### **Authors:**

G Michael Allan MD CCFP, Frank Martino MD CCFP

## **Disclosure:**

Authors do not have any conflicts to disclose.

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