

**ORIGINAL RESEARCH**

# Patterns of Clinical Care Subsequent to Nonindicated Vitamin D Testing in Primary Care

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**Background:** Exponential increases in nonindicated, low-value vitamin D testing have been reported over the past 15 years. Downstream effects of such testing have not been well quantified.

**Methods:** The purpose of this study was to describe patterns of vitamin D testing within primary care of a large regional health system and to explore downstream health service utilization subsequent to nonindicated testing. Instances of vitamin D testing in 2015 were obtained by an electronic health record-automated search. A subset of patients for whom vitamin D testing was classified as nonindicated was identified, and vitamin D-related service utilization was tracked for 24 months.

**Results:** Of the 77,836 adult primary care patient records identified in 2015, vitamin D tests were conducted on 8,042 (10.3%), with 24.3% of tests yielding abnormal results. In the nonindicated test subset (n = 574), substantial clinical variability was illustrated by 85 care pathways and 26 vitamin D prescriptions. Forty-five percent of abnormal vitamin D lab tests were not followed up with repeat vitamin D tests. Vitamin D-related services (laboratory tests, imaging, and prescriptions) occurred at an average rate of 1.6 services per patient during the 24 months following nonindicated vitamin D testing. Some of these services were also classified as nonindicated.

**Conclusions:** Evidence of a health service cascade following nonindicated vitamin D testing exists. Opportunities for improved consistency and quality of care related to vitamin D were observed in our health system. These results may inform clinical pathways related the prevention, evaluation, and treatment of low vitamin D. (J Am Board Fam Med 2020;33:569–579.)

**Keywords:** Electronic Health Record, Low Value Care, Overuse, Primary Health Care, Quality Improvement, Uncertainty, 25-hydroxyvitamin D

## Introduction

Over 25% (\$760 to \$932 billion annually) of US health care spending is considered unnecessary and wasteful.<sup>1</sup> Low value care, which is defined as clinical services that confer minimal or no benefit or may cause harm, is a substantial component of unnecessary health care expenditures.<sup>2,3</sup> The Choosing

Wisely Campaign is an initiative aimed at reducing low value care and highlighting clinical practices inconsistent with the evidence.<sup>4,5</sup> Initially founded in 2012 by the American Board of Internal Medicine Foundation, Choosing Wisely includes recommendations from over 80 professional medical societies.<sup>4</sup>

Three medical societies have identified a Choosing Wisely recommendation of *Do not order population-based screening for vitamin D*.<sup>6</sup> This recommendation aligns with a 2014 US Preventive Services Task Force recommendation statement that advises against vitamin D screening in asymptomatic community-dwelling adults due to

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**Table 1. Indicators for Vitamin D Testing**

Category of Indication*	ICD-10 Text Diagnosis	ICD-10 Code(s) <sup>†</sup>
Vitamin D deficiency or insufficiency, osteomalacia, rickets	Rickets, active	E55.0
	Vitamin D deficiency, unspecified	E55.9
	Sequelae of rickets	E64.3
	Adult osteomalacia	M83
	Adult osteomalacia due to malnutrition	M83.3
	Disorder of bone density and structure, unspecified	M85.9
Osteoporosis, osteopenia	Disorder of bone, unspecified	M89.9
	Osteoporosis with current pathological fracture	M80
	Osteoporosis without current pathological fracture	M81
Pathological fracture or stress fracture	Stress fracture	M84.3
	Pathological fracture	M84.4
Malabsorption or inflammatory bowel disease	Crohn's disease [regional enteritis]	K50
	Ulcerative colitis	K51
	Celiac disease, tropical sprue, blind loop syndrome, pancreatic steatorrhea, other malabsorption due to intolerance	K90.0-K90.4
	Other intestinal malabsorption	K90.89
	Intestinal malabsorption, unspecified	K90.9
Parathyroid, calcium, or phosphate disorder	Postsurgical malabsorption, not elsewhere classified	K91.2
	Idiopathic hypoparathyroidism	E20.0
	Hypoparathyroidism, unspecified	E20.9
	Secondary hyperparathyroidism	E21.0-E21.5
	Hypocalcemia	E83.51
	Hypercalcemia	E83.52
Chronic kidney disease	Chronic kidney disease, stages 3–5; end-stage renal disease	N18.3-18.5; N18.6

\*Indicators for vitamin D testing based on Choosing Wisely criteria.<sup>1</sup>

<sup>†</sup>Patients with a diagnosis of any of these ICD-10 codes were excluded from the subset of patients identified as having nonindicated vitamin D testing in 2015.

ICD-10, International classification of diseases, tenth revision.

insufficient evidence of benefits and harms.<sup>7</sup> The US Endocrine Society and the National Academy of Medicine (formerly known as the Institute of Medicine) also recommend against vitamin D testing in low-risk populations.<sup>8,9</sup> Choosing Wisely and others do recognize, however, that vitamin D testing is indicated in patients at high risk for abnormal vitamin D or related complications (ie, osteoporosis or osteopenia, parathyroid or calcium-related disorder, or chronic kidney disease of  $\geq$  stage 3) (Table 1).<sup>6,10</sup>

At \$100 to \$300 per test, vitamin D testing is itself a low to moderately priced laboratory test.<sup>11</sup> However, the large volume of tests regularly conducted makes vitamin D testing a significant economic investment.<sup>10,12</sup> It has been estimated that over 1 billion vitamin D tests are ordered annually worldwide<sup>13</sup> and that test ordering has increased substantially over the past 15 years.<sup>14–18</sup> Between 25% and 77% of these tests follow a pattern of

nonindicated screening rather than targeted testing of high-risk patients.<sup>11,19–22</sup> Six million vitamin D tests labeled as “low value” were conducted in commercial and Medicare-insured Americans in 2014, tallying over \$800 million. In addition, \$24 million was spent on vitamin D testing considered nonindicated just in the state of Virginia in 2016.<sup>23</sup> Recently, insurance providers have implemented stricter guidelines on vitamin D test reimbursement.<sup>24</sup>

Along with financial burden, overuse of screening and diagnostic tests has been associated with increased risk of patient harm.<sup>25–28</sup> Further research is needed to examine downstream effects and consequences of nonindicated vitamin D testing. The purpose of this study was to describe vitamin D test ordering patterns and explore downstream consequences of nonindicated vitamin D test ordering within primary care of a regional health system.

## Methods

### Study Design

An automated search of the electronic health record (EHR) database for a regional health system was performed to identify instances of vitamin D testing in adult, continuous care patients in 2015. Vitamin D testing was defined as a single 25-hydroxyvitamin D [25(OH)D] test. We also identified a subset of vitamin D tests that were ordered without clinical indication and tracked vitamin D-related laboratory testing, imaging services, and prescriptions incurred during the 24 months following the index vitamin D test. For the purpose of this study, nonindicated vitamin D testing was defined as vitamin D tests that do not meet Choosing Wisely criteria for indicated services (Table 1).<sup>6</sup> This study was approved by the institutional review board of Carilion Clinic (IRB-18-274). The Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) Statement checklist for observational studies was used in the preparation of this manuscript.<sup>29</sup>

### Setting

Carilion Clinic is a nonprofit comprehensive health care system that serves over 1 million patients in Southwest Virginia. Adult primary care services at Carilion Clinic are delivered by 226 family medicine clinicians and 22 internal medicine clinicians (physicians, physicians assistants, and nurse practitioners) in 54 different facilities. Epic (Epic Systems, Wisconsin, US) has served as the EHR platform for Carilion Clinic since 2008, and Quest Diagnostics (New Jersey, US) has served as the laboratory services provider for Carilion Clinic since 2014. Vitamin D test analyses are conducted using an immunoassay methodology certified by the Centers for Disease Control and Prevention Vitamin D Standardization Certification Program. Carilion Clinic assigns 30 to 100 ng/mL as the normal

reference range for 25(OH)D (Table 2). Within our health system, ICD-10 codes are assigned at the time that laboratory tests are ordered.

### Data Extraction

From Carilion Clinic's EHR, we identified adult patients ( $\geq 18$  years of age) who received ongoing care ( $\geq 1$  visit annually) from 1 or more primary care clinicians between January 1, 2013 and December 31, 2017. Data were extracted for any of these continuous primary care patients for whom an outpatient initial vitamin D test (ie, first test within 12 months) was conducted between January 1, 2015 and December 31, 2015. In addition to vitamin D test date and test results, we also extracted the following data: age, gender, race/ethnicity, body mass index (BMI), insurer, and diagnoses code(s) associated with the vitamin D test.

A subset of patients for whom the initial vitamin D test was considered nonindicated was established through the identification of patients who did not have diagnoses shown in Table 1, had no previous vitamin D test(s) documented anywhere in their history, and had not received a vitamin D prescription in the 12 months before their index vitamin D test. The following additional vitamin D-related service data documented in the 24 months subsequent to the nonindicated vitamin D test were extracted for this subset: laboratory tests [25(OH)D, 1,25(OH)2D, calcium, phosphorus, alkaline phosphatase, and parathyroid hormone (PTH)], prescriptions (vitamin D preparations  $>400$  IU, osteoporosis medications, and calcimimetic medications), imaging services (bone mineral density), and diagnoses.

### Data Analysis

Data were analyzed using IBM Statistical Package for the Social Sciences (SPSS) version 26.0 (Chicago, IL). Descriptive statistics (means, standard deviation, and frequencies) were calculated on all data. Differences between means were evaluated using the unpaired *t* test or one-way analysis of variance with Tukey's multiple comparisons test. For categorical data,  $\chi^2$  and Fisher's exact tests were used. Multivariate linear regression analysis was used to evaluate the contribution of patient factors to vitamin D test results. Poisson regression analysis was used to evaluate the contribution of patient factors to number of services incurred.

**Table 2. Vitamin D Test Result Reference Range**

Category	25(OH)D (ng/mL)*
Low	0 to 29.9
Vitamin D deficient	0 to 19.9
Vitamin D insufficient	20 to 29.9
Normal	30 to 99.9
High	$>100$

\*Vitamin D test result reference range based on the reference range established by Carilion Clinic and Quest Diagnostics<sup>2</sup>.

Statistical significance was set at an  $\alpha$  level of 0.05. Plotly R Package version 4 (Quebec, Canada) was used to create an alluvial diagram.

## Results

We identified 77,836 continuously enrolled adult primary care patient records for 2015. Vitamin D tests were conducted on 8042 of these patients (10.3%). The nonindicated vitamin D test subset included 574 patients. Patients in the nonindicated vitamin D test subset were younger, more likely to be male, more likely to be commercially insured or self-pay, and less likely to be insured by Medicare compared with all patients who had vitamin D testing (Table 3).

### *Vitamin D Test Results*

Vitamin D test results were similar among the nonindicated and indicated vitamin D test groups (Table 4), but the nonindicated vitamin D test subset included a greater proportion of abnormal test results (39.5%) than those who had indicated tests (24.3%) ( $P < .001$ ) (Table 4). Vitamin D test results did not differ by patient age. White patients had 18% higher vitamin D test results than black or African American, Asian, and Hispanic patients ( $P < .001$ ). Patients with underweight ( $< 18.5 \text{ kg/m}^2$ ) or normal BMI ( $18.5$  to  $25 \text{ kg/m}^2$ ) had higher (17% and 15%, respectively) vitamin D test results than patients with a BMI of  $> 25 \text{ kg/m}^2$  ( $P < .001$ ) as did those with commercial insurance or self-pay status compared with other insurers (Medicare, Medicaid, and other) ( $P < .001$ ).

### *Downstream Services following Nonindicated Vitamin D Testing*

A total of 4437 vitamin D-relevant laboratory, prescription, and imaging services were ordered for patients who had nonindicated vitamin D testing over the 24-month observation period. Because the volume of calcium ( $n = 2184$ ) and alkaline phosphatase ( $n = 1412$ ) tests ordered was disproportionate to other laboratory tests and because they are part of common test panels (basic metabolic panel and comprehensive metabolic panel, respectively), they were omitted from primary analyses. Thus, we observed a total of 841 vitamin D-relevant laboratory ( $n = 466$ ), prescription ( $n = 344$ ), and imaging ( $n = 31$ ) services during the 24 months following a nonindicated vitamin D test. The provision of these services followed 85 different clinical pathways (ie, steps and timing of vitamin D-related care; data not shown).

Based on Poisson analysis, gender, race, age, insurer, month of test, and index vitamin D test result category (deficient, insufficient, normal, and high) did not predict the number of services incurred. Figure 1 shows the flow of vitamin D-relevant services observed in the nonindicated vitamin D test subset based on results of the index vitamin D test.

### *Laboratory Testing Subsequent to Nonindicated Vitamin D Testing*

Follow-up vitamin D testing was conducted in 200 patients (34.8%) in the nonindicated vitamin D test subset. The time lapse between initial and follow-up test(s) varied broadly (2 to 718 days), but 19 (5.9%) follow-up vitamin D tests took place sooner than 8 weeks after the nonindicated test. The number of follow-up tests conducted over 24 months was 1 (129 patients), 2 (37 patients), 3 (23 patients), 4 (9 patients), 5 (1 patient), or 7 (1 patient). No follow-up vitamin D testing was conducted in 61.7% of patients with abnormal vitamin D test results. Of the 320 total follow-up vitamin D tests conducted, 194 (60.6%) occurred in patients with normal vitamin D test results in their most recent vitamin D test. Patients with insufficient initial vitamin D test results had a 2-fold greater likelihood of follow-up vitamin D tests than patients with deficient test results and a 5-fold greater likelihood than those with normal test results ( $P = .022$ ).

Other vitamin D-relevant laboratory tests included PTH ( $n = 281$ ), phosphorus ( $n = 123$ ), and 1,25(OH)<sub>2</sub>D ( $n = 62$ ). Abnormal test results were observed with 33.3% (PTH), 23.6% (phosphorus), and 20.0% [1,25(OH)<sub>2</sub>D] of these tests. There was no relationship between initial vitamin D test result and likelihood of having a 1,25(OH)<sub>2</sub>D test. Most 1,25(OH)<sub>2</sub>D tests (92%) were conducted in patients who did not have abnormal PTH or elevated calcium test results.

### *Prescriptions Subsequent to Nonindicated Vitamin D Testing*

A total of 275 prescriptions for oral vitamin D supplements were provided to 112 (20%) patients. Nine different vitamin D prescriptions were provided in a total of 26 different treatment regimens (Table 5). Seventy-three (65%) prescriptions were for ergocalciferol (vitamin D<sub>2</sub>) and 39 (35%) were for cholecalciferol (vitamin D<sub>3</sub>). There was no relationship between vitamin D test result and type or

**Table 3. Demographic and Clinical Characteristics of Patients Who Had Vitamin D Testing in 2015**

Characteristic	Patients Who Had Vitamin D Test with Indication (N = 7468)*	Nonindicated Vitamin D Test Subset (N = 574)*
Age (years)		
Means (range)	65.7 ± 13.5 (19.8 to 101.3)	56.9 ± 11.1 <sup>†</sup> (19.9 to 95.3)
18 to 29.9	55 (0.7%)	20 (3.5%)
30 to 39.9	244 (3.3%)	54 (9.4%)
40 to 49.9	658 (8.8%)	104 (18.1%) <sup>‡</sup>
50 to 59.9	1388 (18.6%)	165 (28.7%) <sup>‡</sup>
60 to 69.9	2154 (28.8%)	126 (22.0%) <sup>§</sup>
70 to 79.9	1884 (25.2%)	68 (11.8%) <sup>‡</sup>
80 to 89.9	949 (12.7%)	31 (5.4%) <sup>§</sup>
90 to 99.9	135 (1.8%)	6 (1.0%)
>100	1 (0.1%)	0 (0.0%)
Sex		
Male	1779 (23.8%)	227 (39.5%) <sup>‡</sup>
Female	5689 (76.2%)	347 (60.5%) <sup>‡</sup>
Race/ethnicity		
White	6792 (90.9%)	526 (91.6%)
Black	522 (7.0%)	35 (6.1%)
Asian	53 (0.7%)	3 (0.5%)
Hispanic	22 (0.3%)	2 (0.3%)
Other	79 (1.1%)	8 (1.4%)
BMI (kg/m <sup>2</sup> )		
Overall mean (range)	30.1 ± 7.5 <sup>  </sup> (9.9 to 74.4)	30.7 ± 9.0 <sup>¶</sup> (17.2 to 56.8)
<18.5	73 (1.9%)	1 (0.3%)
18.5 to 24.9	899 (23.0%)	70 (22.4%)
25.0 to 29.9	1252 (32.1%)	99 (31.7%)
30.0 to 34.9	873 (22.4%)	71 (22.8%)
35.0 to 39.9	418 (10.7%)	44 (14.8)
>40.0	388 (9.9%)	27 (8.7)
Insurer		
Medicare	3564 (47.7%)	98 (17.1%) <sup>†</sup>
Medicaid	108 (1.4%)	12 (2.1%)
Commercial	2092 (28.0%)	240 (41.8%) <sup>‡</sup>
Self-pay	1675 (22.4%)	220 (38.3%) <sup>†</sup>
Other	29 (0.38%)	4 (0.70%)
Top 5 diagnoses associated with vitamin D test		
	Vitamin D deficiency: 4202 (56.3%)	Fatigue: 274 (48%)
	Osteoporosis/osteopenia: 2276 (30.5%)	Wellness exam/screening: 113 (20%)
	Chronic kidney disease: 336 (4.5%)	Pain: 90 (16%)
	Malabsorption: 300 (4.0%)	Rheumatological condition: 28 (5%)
	Fracture: 24 (0.3%)	Depression: 25 (4%)

\*Values are n (%) unless otherwise indicated.

<sup>†</sup>P < .001.

<sup>‡</sup>P < .01.

<sup>§</sup>P < .05

<sup>||</sup>n = 3904.

<sup>¶</sup>n = 312.

BMI, body mass index.

dosage of vitamin D prescribed. For example, there were 11 25(OH)D test results of <12 ng/mL. The following vitamin D prescriptions were provided to

these patients: high-dose vitamin D<sub>2</sub> (n = 2), high-dose vitamin D<sub>3</sub> (n = 2), moderate-dose vitamin D<sub>3</sub> (n = 2), and no vitamin D prescription (n = 5).

**Table 4. Results of Initial Vitamin D Tests in Primary Care Patients in 2015**

Result	Patients Who Had Vitamin D Test with Indication (N = 7468), n (% of Indicated Tests)	Nonindicated Vitamin D Test Subset (N = 574), n (% of Nonindicated Tests)
Vitamin D deficient (0 to 19.9 ng/mL)	469 (6.3%)	49 (8.6%)
Vitamin D insufficient (20 to 29.9 ng/mL)	1302 (17.5%)	171 (30.0%)*
Normal vitamin D (30 to 99.9 ng/mL)	5658 (75.7%)	352 (61.3%)*
High vitamin D (>100 ng/mL)	39 (0.5%)	2 (0.5%)

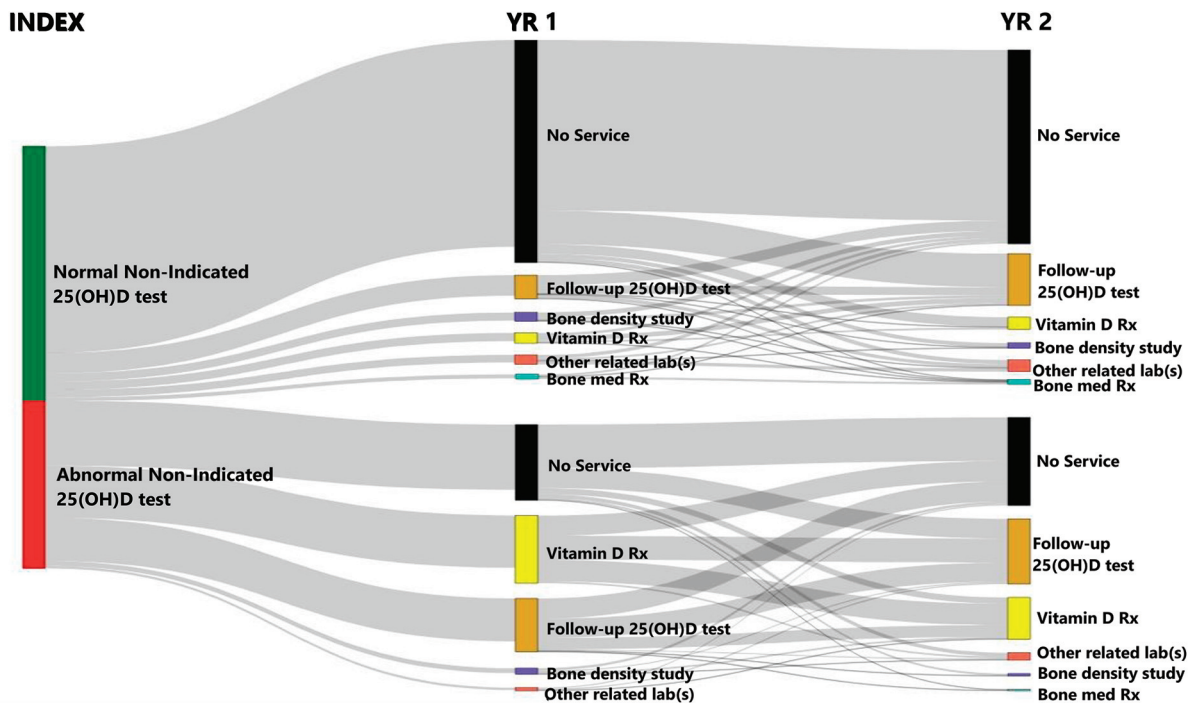
\* $P < .01$ .

Of the 112 patients who received a vitamin D prescription, 82 (73%) had at least 1 follow-up vitamin D test. The type of vitamin D prescription (vitamin D<sub>2</sub> or D<sub>3</sub>) was a significant predictor of change in vitamin D test result ( $R^2 = 0.854$ ,  $P < .001$ ) (Figure 2). Patients prescribed vitamin D<sub>2</sub> experienced a 68% decline in vitamin D test results (range, -710% to 48% change), whereas patients prescribed vitamin D<sub>3</sub> experienced a 43% improvement in vitamin D test results (range, -180% to 430%). There was no significant interaction between type of prescription and time. Half (50%) of patients with low initial vitamin D test results achieved normal test results during the 24-

month observation. The likelihood of achieving normal vitamin D results was significantly higher in patients prescribed moderate-dose (1000 to 2000 IU/day) or high-dose ( $\geq 2000$  IU/day) vitamin D<sub>3</sub> than in those prescribed low-dose (400 to 1000 IU/day) vitamin D<sub>3</sub> ( $P = .030$ ). There were no high (>100 ng/mL) vitamin D test results following the provision of vitamin D prescriptions.

Calcium (without vitamin D) was prescribed to 21 patients (5%), and bone resorption inhibitors were prescribed to 15 patients (3%) who had nonindicated vitamin D tests. Calcimimetic medications (eg, cinacalcet) were not prescribed to any patients in the subset.

**Figure 1. Vitamin D-relevant services incurred in the 24 months following a nonindicated vitamin D test. An alluvial diagram showing the flow of vitamin D-relevant service utilization for 2 years following a nonindicated vitamin D test in 2015.**



**Table 5. Vitamin D Prescriptions Provided to the Subset of Patients Who Had Nonindicated Index Vitamin D Tests in 2015**

Form	Dose	Frequency	Duration	n	Prescription Category
Ergocalciferol/vitamin D <sub>2</sub> (n = 73)	50,000 IU	Weekly	4 weeks	2	High-dose vitamin D <sub>2</sub>
	50,000 IU	Weekly	8 weeks	35	
	50,000 IU	Weekly	12 weeks	20	
	50,000 IU	Weekly	16 weeks	9	
	50,000 IU	Weekly	20 weeks	4	
	50,000 IU	Weekly	24 weeks	2	
	50,000 IU	Weekly	48 weeks	1	
Cholecalciferol/vitamin D <sub>3</sub> (n = 39)	50,000 IU	Daily	2 months	1	High-dose vitamin D <sub>3</sub>
	50,000 IU	Weekly	12 weeks	1	
	10,000 IU	Daily	4 months	1	Moderate-dose vitamin D <sub>3</sub>
	5,000 IU	Daily	3 months	5	
	5,000 IU	Daily	2 months	1	
	4,000 IU	Daily	3 months	1	
	2,000 IU	Daily	10 months	1	
	2,000 IU	Daily	6 months	3	
	2,000 IU	Daily	4 months	3	
	2,000 IU	Daily	3 months	4	
	2,000 IU	Daily	2 months	3	
	2,000 IU	Daily	1 month	1	
	1,000 IU	Daily	12 months	1	Low-dose vitamin D <sub>3</sub>
	1,000 IU	Daily	8 months	1	
	1,000 IU	Daily	6 months	4	
	1,000 IU	Daily	4 months	2	
	1,000 IU	Daily	3 months	1	
	400 IU	Daily	6 months	1	
	400 IU	Daily	3 months	2	
	600 mg calcium + 400 IU	Daily	3 months	2	

*Imaging Subsequent to Nonindicated Vitamin D Testing*

Vitamin D-related imaging studies were conducted on 30 patients (5%) during the 24 months following nonindicated vitamin D testing. One patient underwent 2 imaging studies, whereas the remainder had a single imaging study. The majority of imaging studies (83%) consisted of multisite dual energy radiograph absorptiometry (DXA), and the remainder consisted of axial-only DXA.

*Diagnoses Subsequent to Nonindicated Vitamin D Testing*

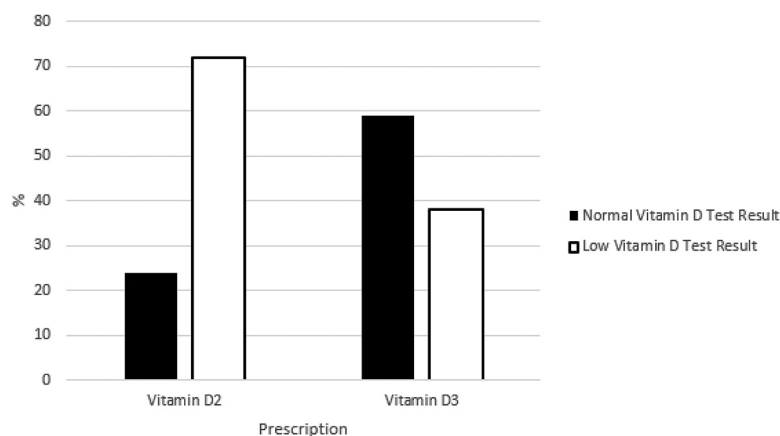
During the 24 months following a nonindicated vitamin D test, a new diagnosis of vitamin D deficiency or vitamin D insufficiency was documented for 356 patients (62%). Of these patients, 107 (21%) had only normal vitamin D test results during the 24-month observation period. Other vitamin D-related diagnoses documented were

osteoporosis (8%), osteopenia (1%), osteoporotic fracture (1%), pathologic fracture (0.5%), stress fracture (0.5%), inflammatory bowel disease (5%), malabsorption (2%), and parathyroid/calcium/phosphate disorder (1%). There was no difference in the likelihood of receiving these diagnoses based on initial vitamin D status.

**Discussion**

Amid controversy in vitamin D research and clinical guidelines, vitamin D testing and prescribing have increased substantially in the past 15 years.<sup>10</sup> Nonindicated testing has been shown to represent a substantial proportion of this testing and is considered a source of low value health care.<sup>6</sup> One purpose of this study was to explore downstream consequences of nonindicated vitamin D testing within primary care of our Southwest Virginia health system. Overall, we observed inconsistency

**Figure 2. Proportion of low and normal vitamin D test results following vitamin D2 and vitamin D3 prescriptions**  
**\*The highest vitamin D test result attained during the 24 months following a nonindicated index vitamin D test.**



in care, opportunities to improve care, and evidence of a low-value service cascade during the 24 months following a nonindicated vitamin D test.

### **Patterns of Care**

Inconsistency in vitamin D-related care has been reported previously<sup>30–32</sup> and existed in the current study. Numerous care pathways and vitamin D treatment regimens were observed. In addition, the number of follow-up vitamin D tests, time between vitamin D tests, and the 25(OH)D concentration associated with a vitamin D deficiency diagnosis were highly variable.

We observed the highest inconsistency in care following insufficient vitamin D test results. Defined as 25(OH)D of 20 to 29.9 ng/mL in the present study (Table 2), insufficient test results were associated with the highest rate of follow-up vitamin D tests (double that of low test results). However, few patients with insufficient test results were given vitamin D prescriptions. It is possible that over-the-counter vitamin D supplements were recommended for patients with insufficient test results, but because only one-third of these patients showed improved vitamin D test results within the 24-month observation period, issues of documentation, adherence, and effectiveness are all possible. Of note, the definition of vitamin D insufficiency is controversial and not even recognized by all practice guidelines. For instance, the Endocrine Society recognizes 25(OH)D of 20 to 29.9 ng/mL as insufficient,<sup>8</sup> but the National Academy of Medicine does not recognize a vitamin D insufficient category and states that 25(OH)D of 20 ng/mL meets

the needs of 97.5% of the population.<sup>9</sup> It is also important to consider that many professional organizations have highly variable recommendations related to indicators for vitamin D testing,<sup>10</sup> which is challenging for clinicians to navigate.<sup>33–35</sup> We observed higher average vitamin D test results ( $41.1 \pm 9.3$  ng/mL) and a lower prevalence of severe vitamin D deficiency (<1%) than previous reports.<sup>36–38</sup> As there are numerous factors that influence vitamin D status, these differences could be based on variability in geography, age, race, adiposity, medication use, socioeconomic status, overall health status, or other factors. For example, our cohort included almost exclusively white patients (who typically have higher vitamin D than individuals with darker skin pigmentation) and patients with lower BMI than other cited studies (higher BMI is associated with lower vitamin D).<sup>24,25,39,40</sup>

A few differences in characteristics of patients who had nonindicated initial vitamin D tests compared with those who had indicated vitamin D tests were identified. The difference in age (nonindicated test patients were younger and less likely to be insured by Medicare) may be due to the exclusion of patients who were recorded as already taking vitamin D supplements. More than 50% of Americans over 60 years of age have been reported to take vitamin D supplements.<sup>41</sup> The greater proportion of self-pay and commercially insured patients in the nonindicated test cohort is likely a factor of this younger age-group that is less likely to be insured by Medicare. The higher rate of abnormal vitamin D test results in the nonindicated test subset may reflect clinicians' clinical instinct related

to the possibility of low vitamin D. However, it is unclear if identification and treatment of low vitamin D is clinically valuable in patients without specific indications.<sup>7</sup>

### **Quality of Care**

We identified some potential opportunities to improve quality of care related to vitamin D. For example, nearly half of low vitamin D test results were not followed up by repeat vitamin D testing (Figure 1). Monitoring 25(OH)D following low test results and at least 12 weeks of treatment is a common practice recommendation.<sup>8,42,43</sup> In addition, two-thirds of vitamin D prescriptions were for vitamin D<sub>2</sub> rather than vitamin D<sub>3</sub>, which has greater bioavailability than vitamin D<sub>2</sub> and is more effective in raising 25(OH)D.<sup>44,45</sup> As this finding is somewhat recent, it is not reflected in major vitamin D guidelines, which describe both forms as suitable treatment options.<sup>8,9</sup>

One-fifth of patients given a new diagnosis of vitamin D deficiency or insufficiency did not have low vitamin D test results. This may reflect the existing controversy around 25(OH)D values.<sup>33</sup> Clinicians may have also assigned such a diagnosis to obtain insurance reimbursement for tests, which has been reported previously.<sup>34</sup>

### **Health Services Cascade**

Abnormal test results have been described as triggering a cascade process by which a number of referrals and additional investigations arise.<sup>46</sup> The volume of vitamin D-relevant services (an average of 1.6 services per patient) that followed a nonindicated vitamin D test suggests that a similar cascade occurs within our health system. Another consideration is that, during 2015, our health system had a limited endocrinology service and did not have a nephrology service, so referrals for these services were made outside of our health system. Thus, we are likely missing vitamin D-relevant health services that were not documented in our EHR.

Some of the services provided subsequent to a nonindicated vitamin D test can themselves be described as low value. For example, more than half of the 320 follow-up tests ordered (up to 7 follow-up tests in 24 months) did not meet our criteria (based on Choosing Wisely) for an indicated vitamin D test. In addition, the majority of 1,25(OH)<sub>2</sub>D tests were conducted in patients who did not fulfill

the Choosing Wisely indications for this test (hypercalcemia and/or abnormal PTH).<sup>4</sup>

The financial impact of nonindicated vitamin D testing can be significant. Based on the charge for an outpatient vitamin D test at Carilion Clinic (~\$278 in 2015), the sum of all vitamin D charges for tests ordered by primary care clinicians in this health system in 2015 is estimated at \$2,946,800. Over \$240,000 was charged for vitamin D testing within the nonindicated test subset. Downstream health services and associated resources for these nonindicated tests may result in additional financial burden. In addition, the overall culture of low-value care is in itself considered harmful. Consequences such as increased false-positive test results, enhanced health risk related to follow-up services, emotional impact to patients, increased resources required to administer services, and decreased opportunity for higher value care have been observed following low-value health services.<sup>26,29</sup>

A strength of this study is the use of EHR data to explore downstream impacts of a common nonindicated laboratory test within a large health system, similar to that which has been performed with other low-value care services.<sup>47,48</sup> We are not aware of other published studies that have examined downstream consequences of low-value vitamin D tests. Although individual chart review was not performed in this study, Isaac et al.<sup>40</sup> recently observed only minor differences in low-value care services identified via automated EHR search and individual chart review.

We acknowledge some limitations in the current study. By nature, EHR research makes assumptions about clinician behavior without necessarily knowing intent. To that end, our results provide a broad description of health service patterns without consideration of details that may have factored into clinician decisions and behaviors. Some tests that we classified as nonindicated may actually have been indicated, and vice versa. We were also unable to account for over-the-counter vitamin D supplements (unless reported in patients' medication list), diet, and sun exposure. Finally, we acknowledge that service chronology does not necessarily reflect downstream effects but instead interprets the high volume of vitamin D-relevant services as suggestive of a causal association. Overall, our results provide one of the first attempts to capture downstream health service use following nonindicated vitamin D testing. The broad patterns we observed generate hypotheses for future investigations.

## Conclusions

In 2015, vitamin D tests were ordered for 10% of adult primary care patients within our Southwest Virginia health system, with approximately one-quarter yielding abnormal results. In a subset of these patients who had nonindicated vitamin D tests, there seems to be high variability in care pathways and some opportunities to improve care quality. We also observed evidence of a low-value service cascade during the 24 months following a nonindicated vitamin D test. Taken together, these results provide insight into primary care clinicians' practice patterns in light of evolving research and conflicting guidelines related to the clinical management of vitamin D status. Possible interventions to improve care and reduce inconsistency related to vitamin D services could include clinician education and feedback, development of clinical pathways related to the identification and treatment of vitamin D-related abnormalities, modifications to EHR default settings and prompts, and patient education. Further research on the downstream impacts of nonindicated vitamin D testing is needed.

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