

Proximal muscle strength as a predictor of vitamin D insufficiency in elderly

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ABSTRACT

Objectives: This study aims to evaluate the association of serum vitamin D level with proximal muscle strength, tone, elasticity, and stiffness in elderly.

Patients and methods: Between September 2017 and January 2018, a total of 109 participants (21 males, 88 females; mean age: 71.2±4.6 years; range, 65 to 85 years) were included in the study. The proximal muscle strength was evaluated by MicroFET® 3 device. The muscle tone, elasticity, and stiffness were measured using the MyotonPRO® digital palpation device. Serum 25-hydroxyvitamin D [25(OH)D] level was tested by high-performance liquid chromatography. A receiver operating characteristic (ROC) curve was performed to evaluate the potential role of MicroFET® 3-measured proximal muscle strength in the quantification of vitamin D status.

Results: Vitamin D sufficient participants had a higher proximal muscle strength ($p<0.001$). Quadriceps and hamstring elasticity at the non-dominant site were significantly higher in vitamin D sufficient group ($p<0.05$). The ROC analysis indicated that the deltoid muscle strength had the potential of determining vitamin D insufficiency with moderate accuracy (area under the curve=0.744; 95% confidence interval: 0.643-0.845; $p<0.001$).

Conclusion: Proximal muscle strength, elasticity, and physical performance are associated with vitamin D status. Proximal muscle strength measured by a hand-held dynamometer can be used as a predictor of hypovitaminosis D in elderly.

Keywords: Geriatrics, gerontology, muscle strength, muscle tension, muscle tone, vitamin D.

Vitamin D is an essential molecule for human beings. Vitamin D, either as ergocalciferol (vitamin D₂) or cholecalciferol (vitamin D₃), can regulate cell metabolism and physiology by means of vitamin D receptors (VDRs).^[1,2] Since VDRs are vastly expressed in musculoskeletal system tissues, vitamin D is highly involved in the muscle and bone metabolism.^[3]

Vitamin D deficiency is regarded as an important public health issue. Decreased vitamin D level contributes to several comorbidities including malignancy, coronary heart disease, neurocognitive disorders, autoimmune-autoinflammatory diseases,

and musculoskeletal disorders.^[4,5] The detrimental effects of vitamin D deficiency are more prominent in elderly. The most widely accepted theory about the sensitivity of elderly is that the expression of the VDRs declines with age.^[6] Furthermore, increasing age is related to decreased exposure to sunlight, major organ pathologies/failures, and inadequate oral intake of vitamin D.^[5] Thus, the prevalence of hypovitaminosis D can reach up to 90 to 100% of elderly population.^[3,5]

A lack of vitamin D may lead to impaired muscle functions partly related to its impact on the biological properties of the muscle tissue.^[3,7] Prolonged

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vitamin D deficiency or insufficiency may cause type 2 fiber atrophy, fatty infiltration, fibrosis, and increased glycogen granules in elderly.^[6,8] In the light these data, we, in the present study, aimed to evaluate the relation of vitamin D level with the strength, tone, elasticity, and stiffness of proximal muscles.

PATIENTS AND METHODS

This cross-sectional study was conducted at the Department of Physical Medicine and Rehabilitation, Faculty of Medicine, Çukurova University between September 2017 and January 2018. A total of 200 individuals aged ≥ 65 years were assessed for eligibility. Exclusion criteria were as follows: (i) major organ failure, (ii) malignancy, (iii) malabsorptive conditions, (iv) neurological disability, and (v) history of endocrinological and/or gastroenterological surgery. Finally, a total of 109 participants (21 males, 88 females; mean age 71.2 ± 4.6 years; range, 65 to 85 years) were included in the study. A written informed consent was obtained from each participant. The study protocol was approved by the Ethics Committee of Çukurova University Faculty of Medicine (Date of approval: 8-September-2017, Approval number: 68/3). The study was conducted in accordance with the principles of the Declaration of Helsinki.

Demographic characteristics of all participants variables were recorded. Serum 25-hydroxyvitamin [25(OH)D] level was measured using high-performance liquid chromatography (HPLC). The serum samples were obtained in the morning after overnight fasting. In accordance with the guidelines of the Endocrine Society, the level of 25(OH)D was classified as sufficient (≥ 30 ng/mL), insufficient (21-29 ng/mL), and deficient (< 20 ng/mL).^[9]

The proximal muscle strength was evaluated by the MicroFET® 3 hand-held dynamometer (Hoggan Health Industries Inc., UT, USA). All measurements were done by a single physician and in the sitting position. For deltoid muscle, upper extremity was positioned in 90° of shoulder abduction and internal rotation. The dynamometer was placed on the distal humerus nearby the lateral epicondyle. The strength of biceps muscle was tested, while the arm was at side and the forearm was at 90° of flexion. The dynamometer was located on the volar side of the distal forearm close to the wrist. For triceps muscle, the elbow and the forearm were positioned at full extension in the coronal plane. The dynamometer was cited on the dorsal side of the distal forearm. The quadriceps strength was evaluated, while the

knee was at full extension and the dynamometer was placed on the anterior aspect of the distal tibia. For hamstring muscle, the knee positioned at 90° flexion, and the dynamometer was put on the posterior side of the distal tibia. Maximal voluntary isometric muscle contraction from elbow flexion and extension, knee flexion and extension, and shoulder abduction were measured three times. All measurements were based on the break technique. Accordingly, the physician attempted to overpower the efforts of the subject, while maximal voluntary isometric muscle contraction. The value where the examiner could break the participant's strength was regarded as the isometric muscle strength. The average strength of three consecutive measurements was calculated for each muscle and peak force was given in pounds.^[10]

Mechanical properties (tone, elasticity, and stiffness) of the proximal muscles were evaluated using the MyotonPRO® (Myoton AS, Tallinn, Estonia) digital palpation device. The measurements were performed, while the participants were lying down in a relaxing position following a resting period of 10 min. The MyotonPRO® device was set to the multi-scan mode and to a pre-pressure of 0.18 Newtons (N). Deltoid, biceps, and quadriceps muscles were evaluated in the supine position. On the other hand, triceps and hamstring muscle measurements were performed in the prone position. The MyotonPRO® was positioned perpendicular to the skin over the most bulky part of the muscle. A mechanical impulse at a force of 0.40 N was applied to the muscle for 15 microsec. The damped oscillations were recorded by an accelerometer. Resulting numerical values were calculated to depict the viscoelastic and biomechanical properties of the muscle. The average of two consecutive measurements was recorded for each tone, elasticity, and stiffness.^[11,12] Muscle tone is described by the frequency (Hz) of the oscillations. Muscle elasticity is characterized by the logarithmic reduction of the oscillations. Muscle stiffness (N/m) represents the resistance of the muscle during contraction.^[13]

Physical performance was evaluated by the Timed Up and Go (TUG) test. The participants were instructed to stand up from the armless chair without holding anywhere, come back to the chair after walking three meters with the normal pace, and sit down to the chair. All participants were allowed to wear comfortable shoes and not to use any mobility aids during the test. Walking duration was measured by a chronometer and recorded as sec.^[14,15]

Statistical analysis

Statistical analysis was performed by using the IBM SPSS version 20.0 (IBM Corp., Armonk, NY, USA). Descriptive data were expressed in mean \pm standard deviation (SD), median (min-max) or number and frequency. Data distribution was tested using the Shapiro-Wilk test. Comparative analysis was performed using the Mann-Whitney U test and Kruskal Wallis test. Correlation of continuous variables was analyzed by the Spearman's rank test. Further post-hoc analysis was performed for two-group comparisons. A receiver operating characteristic (ROC) curve was used to evaluate the accuracy of proximal muscle strength in determining of vitamin D insufficiency. A *p* value of <0.05 was considered statistically significant.

RESULTS

Of a total of 200 elderly individuals, 91 were excluded from the study due to chronic kidney disease ($n=43$), history of gastroenterological disorders ($n=37$), neurological disability ($n=25$), and ongoing malignancy ($n=9$). Demographic characteristics of all participants ($n=109$) included in the study are shown in Table 1.

The median 25(OH)D level was 21.7 ng/mL. Of the participants, 30.3% and 42.2% had vitamin D insufficiency and deficiency, respectively. When the sample was categorized into three as vitamin D deficient, vitamin D insufficient, and

	n	%	Median	Min-Max
Age (years)			71	65-85
Sex				
Male	21	19.3		
Female	88	80.7		
Height (cm)			158	142-180
Weight (kg)			72	40-102
Body mass index (kg/m ²)			27.9	17.5-41.9
Min: Minimum; Max: Maximum.				

vitamin D sufficient groups, the body mass index was significantly higher in vitamin D deficient group ($p=0.005$). On the other hand, there was no significant difference among the groups in terms of age, sex, height, employment status, smoking, and alcohol consumption.

The proximal muscle strength showed a statistically significant difference among vitamin D groups ($p<0.001$). Vitamin D deficient group had the lowest values for each tested muscle strength (Table 2). In terms of physical performance, the median duration of TUG test was 12.6 sec and 12.1 sec in vitamin D deficient and insufficient groups, respectively ($p=0.043$). On the other hand, proximal muscle biomechanical properties (tone, elasticity, and stiffness) were not different among the groups, except for the elasticity of

	Deficient		Insufficient		Sufficient		<i>p</i>
	Median	Q1-Q3	Median	Q1-Q3	Median	Q1-Q3	
Deltoid							
Dominant	6.1	5.4-6.5	6.8	5.8-7.3	7.4	6.8-7.6	<0.001
Non dominant	5.5	5.1-6.2	6.1	5.4-6.9	6.9	6.4-7.2	<0.001
Biceps							
Dominant	6.4	5.8-6.8	6.9	6.1-7.5	7.2	6.2-7.2	<0.001
Non dominant	5.9	5.3-6.3	6.3	5.8-6.9	6.9	5.0-7.7	<0.001
Triceps							
Dominant	5.6	4.9-5.9	6.4	5.8-6.9	6.4	6.0-6.8	<0.001
Non dominant	5.3	4.0-7.0	5.9	5.3-6.5	6.1	5.8-6.2	0.001
Quadriceps							
Dominant	6.5	5.8-7.1	7.0	6.5-7.8	7.2	7.0-7.4	<0.001
Non dominant	6.0	5.3-6.6	6.5	6.0-7.2	6.8	6.4-7.3	0.001
Hamstring							
Dominant	6.2	5.8-6.5	6.8	6.1-7.3	7.1	6.5-7.4	0.001
Non dominant	5.8	5.3-6.2	6.4	5.4-6.9	6.6	6.1-7.0	0.001
Values are given in median (Q1-Q3).							

quadriceps and hamstring muscle (only at the non-dominant site), which was significantly higher in vitamin D sufficient participants ($p < 0.05$ for quadriceps and hamstring muscles, post hoc = 0.043) (Table 3).

The ROC analysis was performed to evaluate the ability of proximal muscle strength to predict vitamin D status in elderly. Accordingly, the area under the curve (AUC) of deltoid muscle strength for determining

TABLE 3
Comparison of biomechanical properties among vitamin D groups

	Deficient		Insufficient		Sufficient		Total		<i>p</i>
	Median	Q1-Q3	Median	Q1-Q3	Median	Q1-Q3	Median	Q1-Q3	
Deltoid									
Tone									
Dominant	15.0	13.9-17.4	16.5	14.1-18.0	15.5	14.0-17.6	15.5	14.0-17.6	0.465
Non dominant	15.1	12.8-16.5	15.2	13.4-16.7	15.1	13.4-16.5	15.1	13.4-16.5	0.760
Elasticity									
Dominant	1.20	1.02-1.34	1.22	1.02-1.39	1.20	1.02-1.34	1.20	1.02-1.34	0.663
Non dominant	1.19	1.10-1.34	1.18	1.10-1.28	1.18	1.06-1.29	1.18	1.06-1.29	0.232
Stiffness									
Dominant	242.0	190.5-293.5	245.0	200.5-302.0	246.0	197.5-299.5	246.0	197.5-299.5	0.799
Non dominant	227.0	198.5-301.8	256.0	191.0-277.5	238.0	198.0-284.5	238.0	198.0-284.5	0.976
Biceps									
Tone									
Dominant	14.0	13.0-14.9	13.4	12.6-14.2	13.7	12.6-14.8	13.7	12.6-14.8	0.209
Non dominant	13.3	12.3-14.0	12.4	12.0-13.5	13.0	12.1-14.0	13.0	12.1-14.0	0.180
Elasticity									
Dominant	1.40	1.22-1.57	1.39	1.15-1.60	1.40	1.19-1.58	1.40	1.19-1.58	0.903
Non dominant	1.36	1.25-1.54	1.38	1.26-1.55	1.35	1.24-1.53	1.35	1.24-1.53	0.398
Stiffness									
Dominant	237.5	211.8-259.5	221.0	198.0-241.0	232.0	208.5-252.0	232.0	208.5-252.0	0.111
Non dominant	231.5	207.5-248.0	222.0	209.5-242.5	224.0	207.0-244.5	224.0	207.0-244.5	0.441
Triceps									
Tone									
Dominant	12.6	11.0-13.6	13.2	11.5-14.6	12.8	11.2-14.2	12.8	11.2-14.2	0.269
Non dominant	12.4	11.0-13.7	12.5	11.0-14.2	12.3	11.0-14.0	12.3	11.0-14.0	0.320
Elasticity									
Dominant	1.96	1.71-2.40	1.94	1.38-2.25	1.94	1.66-2.32	1.94	1.66-2.32	0.660
Non dominant	1.74	1.55-2.02	1.74	1.49-1.98	1.74	1.55-2.03	1.74	1.55-2.03	0.784
Stiffness									
Dominant	241.5	214.0-288.8	249.0	205.5-272.0	248.0	215.5-284.0	248.0	215.5-284.0	0.621
Non dominant	251.5	224.8-275.8	244.0	216.0-285.5	253.0	221.5-275.0	253.0	221.5-275.0	0.824
Quadriceps									
Tone									
Dominant	12.9	12.4-14.4	13.3	11.9-15.2	13.0	11.9-14.4	13.0	11.9-14.4	0.912
Non dominant	13.1	11.9-14.1	13.0	11.9-14.2	13.0	11.9-14.1	13.0	11.9-14.1	0.959
Elasticity									
Dominant	1.38	1.11-1.67	1.55	1.21-1.82	1.46	1.21-1.75	1.46	1.21-1.75	0.160
Non dominant	1.30	1.10-1.54	1.55	1.27-1.78	1.43	1.24-1.71	1.43	1.24-1.71	0.021
Stiffness									
Dominant	224.5	196.5-247.8	225.0	197.0-279.5	225.0	196.0-256.5	225.0	196.0-256.5	0.697
Non dominant	226.5	196.5-255.3	228.0	209.0-254.5	228.0	204.0-255.0	228.0	204.0-255.0	0.519
Hamstring									
Tone									
Dominant	13.7	12.3-15.0	12.9	11.9-15.3	13.3	12.3-14.7	13.3	12.3-14.7	0.606
Non dominant	12.5	11.4-14.0	12.6	11.4-14.3	12.5	11.4-13.4	12.5	11.4-13.4	0.836
Elasticity									
Dominant	1.47	1.30-1.73	1.59	1.42-1.88	1.51	1.40-1.85	1.51	1.40-1.85	0.131
Non dominant	1.44	1.22-1.61	1.55	1.30-1.79	1.45	1.35-1.67	1.45	1.35-1.67	0.033
Stiffness									
Dominant	227.0	203.8-255.3	238.0	209.0-258.0	234.0	206.5-256.0	234.0	206.5-256.0	0.736
Non dominant	220.5	187.5-241.5	224.0	119.0-261.5	223.0	194.0-247.5	223.0	194.0-247.5	0.508

Values are given in median (Q1-Q3).

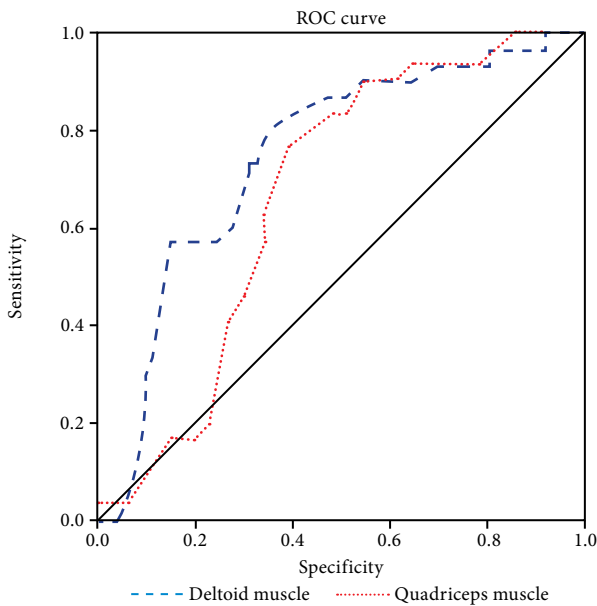


Figure 1. ROC curves of deltoid and quadriceps muscle strength in determining vitamin D insufficiency (AUC for deltoid= 0.744; AUC for quadriceps= 0.667).

ROC: Receiver operating characteristics; AUC: Area under the curve.

vitamin D insufficiency was 0.744 ($p < 0.001$). The best cut-off value for deltoid muscle strength was calculated as 7.0 lb. Using this cut-off value, diagnostic sensitivity and specificity of the deltoid muscle strength were 68.4% and 73.3%, respectively. On the other hand, the ROC curve of quadriceps muscle strength (cut-off value=7.1 lb) showed lower accuracy in predicting vitamin D status (AUC=0.667; $p = 0.007$; sensitivity=65.8%; specificity=63.3%) (Figure 1).

DISCUSSION

Vitamin D is a secosteroid hormone essential for calcium and phosphate metabolism.^[16] Vitamin D deficiency is a common condition affecting almost one billion individuals in the world.^[17] Over the past decade, there has been an extensive amount of research on this topic.^[18-21] The results from these studies have shown that vitamin D deficiency is more common in elderly, compared to younger adults.^[20,21] Consistent with the literature, the current study showed that 72.5% of elderly had vitamin D insufficiency [25(OH)D level < 30 ng/mL]. Age-related decreases in dietary intake, exposure to sunlight, gastrointestinal absorption, and hydroxylation capacity are some of the underlying causes of vitamin D insufficiency/deficiency in elderly population.^[22]

There are several detrimental effects of decreased vitamin D level on human metabolism.^[23] One of the

most prominent health effects of vitamin D deficiency is seen in the musculoskeletal system.^[24] It has a strong negative impact on bone health. Rickets, osteomalacia, osteopenia, and osteoporosis are conditions associated with vitamin D deficiency. Vitamin D also affects muscle mass, muscle strength, and neuromuscular performance.^[24,25] It may also have an impact on the biomechanical properties of the muscle. With this hypothesis in mind, we evaluated the potential effect of vitamin D status proximal muscle biomechanical properties including tone, elasticity, and stiffness in our study. However, other than the elasticity component, there was no relationship between the muscle biomechanical properties and vitamin D status. Elasticity of quadriceps and hamstring muscle at the non-dominant extremity was significantly higher in vitamin D sufficient individuals. The literature search reveals no study evaluating the relationship between vitamin D status and muscle biomechanical properties using the MyotonPRO® device. On the other hand, there is a limited number of data regarding the impact of age/sex on the biomechanical properties of muscles. Agyapong-Badu et al.,^[26] in their study, evaluated tone, elasticity, and strength of the biceps and rectus femoris muscles. These three biomechanical parameters were compared between young adults (20-35 years of age) and elderly (65-82 years of age). Elderly participants revealed higher stiffness and lower elasticity in biceps and rectus femoris muscles than young adults. On the other hand, tone was higher in biceps muscle in elderly with no difference in rectus femoris muscle between the groups. The authors also showed no impact of sex on muscle elasticity in elderly group. However, the present study demonstrated that proximal muscle elasticity, tone, and stiffness were higher in men than those in women. Further studies are warranted to clarify the potential role of age, sex, and vitamin D status on the biomechanical properties of muscles in elderly.

The present study confirmed once again that hypovitaminosis D is an important determinant of muscle strength in elderly. The vitamin D deficient group had the lowest values for each tested muscle strength by MicroFET® 3. With regard to the physical performance, participants with a 25(OH)D level of < 20 ng/mL revealed a longer median duration of TUG test than those with a vitamin D level of > 20 ng/mL. Houston et al.,^[27] in their cross-sectional study, evaluated physical performance among elderly by Short Physical Performance Battery and hand grip strengths using a hand dynamometer. Physical performance scores were significantly lower in the

vitamin D deficient group, compared to participants with 25(OH)D level of >10 ng/mL. In terms of hand grip strength, there was a significant difference between vitamin D insufficient and sufficient groups. A multi-center, retrospective study compared muscle strength between postmenopausal women with optimal vitamin D level and those with a 25(OH)D level of <30 ng/mL.^[28] The handgrip strength and lower extremity isokinetic knee extensor strength were significantly lower in participants with suboptimal vitamin D level. Additionally, muscle strength was found to be significantly correlated with 25(OH)D level.^[28]

Given the close relationship between proximal muscle strength and vitamin D level in older adults, we further postulated that proximal muscle strength could be used to predict vitamin D deficiency in daily clinical practice. With this aim, further ROC analysis was carried out to determine whether muscles properties (strength, tone, elasticity and stiffness) had any diagnostic value in determining the vitamin D status of elderly individuals. Myotonometric measures including tone, elasticity, and stiffness were found to have no diagnostic role in estimating the vitamin D level. However, the proximal muscle strength measured by MicroFET[®] 3 hand-held dynamometer revealed a predictive value with moderate (AUC=0.744) and low (AUC=0.667) diagnostic accuracies for deltoid and quadriceps muscles, respectively. It was found that a deltoid muscle strength of 7.0 lb showed 68.4% sensitivity and 73.3% specificity in discriminating between sufficient (>30 ng/mL) and insufficient (<30 ng/mL) levels of vitamin D. To the best of our knowledge, this is the first study to evaluate the muscle strength as a diagnostic tool in determining vitamin the status. There are many detrimental clinical consequences of vitamin D insufficiency/deficiency in elderly. One can consider screening the vitamin D level of every single individual and to tailor supplementation accordingly. However, given the high costs of serum 25(OH)D testing and lack of enough evidence, several authorities do not recommend routine screening in elderly.^[29,30] On the other hand, while supplementing the patient without a priori testing, it is important to avoid high levels of vitamin D, since serum vitamin D level has a U-shaped relationship with several health conditions with its overdose, leading to an increased fall risk and fractures.^[31,32] Therefore, it would be of great value to anticipate hypovitaminosis D using a cheap and practical method in daily clinical practice. Accordingly, many researchers have attempted to find alternative ways to evaluate vitamin D level. Annweiler

et al.^[33,34] developed a clinical diagnostic tool for the identification of older community-dwellers with hypovitaminosis D. However, none has evaluated the predictive role of proximal muscle strength so far.

In conclusion, there is a close relationship between the muscle strength and vitamin D status. Proximal muscle strength measurement results are valuable in terms of estimating vitamin D status in elderly. Given the moderate diagnostic accuracy of deltoid muscle test results, deltoid muscle strength measured by a hand-held dynamometer can be used as a predictor tool for hypovitaminosis D in elderly population.

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