# Assessment of Medication Adherence and Factors Contributing to Non-Adherence to Calcium and Vitamin D as Mainstay in Treatment and Prophylaxis of Osteoporosis

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#### ABSTRACT

Introduction: Calcium and vitamin D are the cornerstone in treatment and prevention of osteoporosis as recommended by several guidelines. The efficacy of this combination regimen is limited due to poor adherence to the prescribed medication. Aim: This study aimed to assess patient's adherence and persistence to Calcium and vitamin D supplements and to address reasons for non-adherence. Methods: This is a prospective quantitative survey, included patients who were attending a secondary care rheumatology clinic in a teaching hospital. Recruitment period was between April and June 2014. Patients were verbally asked before distributing the questionnaire about the duration of use of Calcium and vitamin D supplements, and the reason for prescribing it. Results: Regarding adherence score, the difference between calcium and vitamin D groups was not statistically significant (p=0.175). About one-third of patients in the two groups showed low adherence; 31% (53/171) and 38.2% (128/335) in calcium and vitamin D groups respectively. Age showed to influence adherence as the difference in scores between the two groups was significant (p=0.001). The younger patients (18-39 years) reported lower adherence to both medications. The educational status did not show a significant effect on

adherence. The most frequently reported reason for non-adherence in both groups was "Forgetting to take medication" (29.5%, 89/302). About 25% of the patients stated that the reasons behind their non-adherence are multi-factorial. **Conclusion:** Low rates of adherence to Calcium and vitamin D supplements were reported among patients in the two treatment groups. The difference in adherence rate among these two groups was not significant.

Key words: Calcium; vitamin D; patient's adherence; osteoporosis

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### **INTRODUCTION**

Osteoporosis has been defined by many international organizations, one of the most accept definitions is the World Health Organization (WHO) definition on 1994 which is "a disease characterized by low bone mass and micro-architectural deterioration of bone tissue, enhanced bone fragility and a consequent increase in fracture risk" (Kanis et al.). The increased life expectancy and the aging population have led to a high prevalence of osteoporosis, making it a primary health concern associated with high burden. Evidence is suggesting that the prevalence of osteoporosis will continue to rise and escalate. According to the International Osteoporosis Foundation (IOF) fast fact sheet (2011); about 50% of Asian and Caucasian women and 25% of Asian and Caucasian men of the age group of 50 years and above shall experience an Osteoporosis-related fracture in their lifetime. Osteoporosis highly impacts the quality of life of the affected population. About the local population in Qatar,<sup>[1-5]</sup> stated that "Osteoporosis is common among menopausal Qatari women and should be considered a matter of public concern". This was concluded from a cross-sectional study that aimed to establish a reference value for Qatari women and to compare it with values of the other Arab countries and the universal values.

The management of osteoporosis involves several approaches, including medication, lifestyle changes, and fall prevention.<sup>[5-7]</sup> The mainstay of the pharmacologic treatment includes calcium (1200 mg daily) and vitamin D supplements. These may be used with other osteoporosis medications, like; Bisphosphonates, parathyroid hormone (Teriparatide), the selective estrogens Receptor Modulators (SERM) (Raloxifene), calcitonin (no more used),<sup>[8-10]</sup> Although the adherence to calcium and vitamin D is the rate limiting step in the treatment response, yet, there no available literature about the adherence level of calcium and vitamin d supplementation in the Middle East generally and Qatar especially. In this study, we will shed some light on the adherence and persistence of Qatar's population to Ca/Vitamin D supplementation, and bridges the gap in the literature about the real causes for non-adherence in Qatar's population.

Medications' efficacy, adherence, and persistence are all important determinants of the treatment outcomes. Therefore, in this study, we aim to assess the adherence to Ca/Vitamin D supplements and to investigate the reasons for non-adherence. This effort is intended to establish ways to overcome the identified barriers enhance the efficacy and contribute to a better life quality for osteoporotic patients in Qatar.

#### **MATERIALS AND METHODS**

The study proposal has been reviewed and approved by the institutional review board oversees research at Hamad Medical Corporation research and ethics committee, the research Id number is (14134/14). A questionnaire was given to all patients attending the rheumatology clinic in a Tertiary teaching hospital at Hamad Medical Corporation, during the period from April to June 2014. The populations of this study were patients 18 years old and above, receiving Calcium and vitamin D as treatment or prophylaxis of osteoporosis for one year or more and were willing to participate in the study. Patients were excluded from the study who did not accept to answer the questionnaire, short term prescription or taking vitamin D and calcium for other indications than osteoporosis indications.

Patients were approached in the waiting areas, briefed about the survey idea and asked verbally to participate in the study before receiving the questionnaire. A consent form was obtained from each patient

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Demographical data and participant's characteristics were analyzed using descriptive statistics (frequencies, Percentages, mean  $\pm$  standard deviation) utilized for the response. Chi-square test was used to assess the effect of different factors. Reliability and validation test of tools were done, and SPSS software was used to figure the data.

## RESULTS

This study included 445 patients who met the inclusion criteria. Out of these, 128 patients were taking calcium (28%), 148 patients were taking vitamin D (33%), and 169 patients were taking vitamin D and calcium together (39%) as illustrated in Table 1.

Patients have stratified into 3 age-groups; group A (18-39 years), B (40-59 years), and C (>60 years). Group B was dominant in this study, representing around 45% in both groups (45.3% in Calcium group and 44.9% in Vitamin D group). The participant's education level was college or higher for almost half the population (46% Calcium group, 47.9% Vitamin D group and 36% Calcium and Vitamin D group). There was no statistically significant difference between Calcium and vitamin D groups regarding adherence score (p=0.673) [Figure 1].

More than one-third of participants in three groups showed low adherence score; 38.2% of patients in Vitamin D group and 26.6% in Calcium and Vitamin D group, and 31% in Calcium group. Medium adherence score was 33%, 33.7% and 43.4% for Calcium group, Vitamin D group and Calcium and Vitamin D group, respectively [Figure 1]. The adherence score between all ages showed no statistical difference (p=0.053).

Forget to take the medications achieved the highest percentage (19.1%) among the reasons for non-adherence in all groups (15.6%, 17.0% and 23.6% for Calcium group, Vitamin D group and Calcium and Vitamin D group respectively). On the other hand lack of benefits showed the least percentage (1.5%) as a reason of non-adherence among studied patients (0% in Calcium group and 2% Vitamin D group and Calcium and Vitamin D group). 31.4% of patients described multi-reasons of non-adherence through all groups like; cost, side effects, availability, poly-pharmacy, condition improved and forget to take medications [Figure 2].

The result showed that the low adherence in college was lower than non- college within the age 18- 39 years old while it was vice versa in other age groups [Tables 2 and 3].

# DISCUSSION

This study was designed to assess adherence and persistence to calcium and vitamin D supplements in patients with osteoporosis, and characterize the reasons for non-adherence. Adherence was as low as 31.4% in all groups, and the main reason for non-adherence was reported by our patients was "forgetting to take the medication", which is comparable to the results obtained from study evaluated

adherence among post-postmenopausal women over 10 years of treatment, sharing the same main reason of non-adherence which was forgetfulness.<sup>[11,12]</sup> However, another study revealed a higher adherence level to calcium and vitamin D among postmenopausal women, 71% of them were adherent at baseline which decreased at three months follow- up to reach 68%.<sup>[13]</sup>

Surprisingly, the results showed low adherence level among the age group A (18-39 years), who are attending the college, when compared to other age groups. However, highest rate of adherence was observed in the non-collage group among the age groups B (40 -59 years) and C (>60 years). This age and attitude can be explained that the younger patients in better health status, than their comparator of other age groups, and are not aware of the important of the disease or being engaged with their activities which indicates future need for studies to investigate the reasons for better understanding of the problem and to participate actively in solve this problem. Also, in this study there were some other reported reasons like; unpalatable taste, unable to swallow large tablets, side effect, availability, poly-pharmacy, and improved general condition.

This study is the first of its type in Qatar, and its findings are important as they aid clinicians in improving health outcomes and enhancing patient's adherence via providing an insight into the current problem of non-adherence to a therapy that is considered the mainstay of osteoporosis treatment. It also shows the reasons patients are nonadherent, which would help in directing counseling of this patient population to address these reasons. The issue of non-adherence to chronic therapies has been recognized by the World Health Organization (WHO) as an issue that severely compromises health outcomes, in particular with the growing population of elderly patients and the increase in the burden of chronic disease. The WHO highlights the need for evolution of healthcare system to contain this problem, especially in the developing countries. Numerous studies have been conducted to assess adherence to chronic medications and evaluate the etiology of non-adherence. Adherence to chronic medications







Figure 2: Reasons for nonadherence

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	Calcium group	Vitamin D group	<b>Calcium and Vitamin D group</b>		
Characteristics	Proportion, n (%)	Proportion, n (%)	Proportion, n (%)		
Total Patient No.	atient No. 128 (46%) 148 (54%)		169 (39%)		
iender					
Male	10 (7.8 %)	17 (11.4 %)	25 (15%)		
Female	118 (92.2 %)	131 (88.6 %)	144 (85%)		
lge					
(A) 18-39 years	27 (21 %)	37 (25 %)	10 (6%)		
(B) 40-59 years	58 (46 %)	66 (44.9 %)	79 (47%)		
(C) >60 years	43 (33 %)	45 (30.1 %)	80 (47%)		
ducation Level					
College level	59 (46 %)	70 (47.9 %)	60 (36%)		
Non-college level	69 (54 %)	78 (52.1 %)	109 (64%)		

Table 2: Level of adherence and reasons for non-adherence distribution among studied groups

	Calcium group	Vitamin D group	Calcium and Vitamin D	
	Proportion, n (%)	Proportion, n (%)	Proportion, n (%)	
evel of adherence				
Low adherence	39 (31 %)	56(38.2 %)	45(26.6 %)	
Medium adherence	43 (33.9 %)	50 (33.7 %)	74(43.4 %)	
High adherence	46 (35.1 %)	42 (28.1 %)	50 (30 %)	
leason of low and medium adherence				
Forgot to take	20 (15.6 %)	25 (17 %)	40 (23.6 %)	
Too many medication	8 (6 %)	6 (4 %)	20 (11.8%)	
Condition improved	8 (6 %)	9 (6 %)	4 (2 %)	
Lack of Benefits	0 (0 %)	3 (2 %)	4 (2 %)	
Cost	3 (2.3 %)	2 (1.3 %)	2 (1.3 %)	
Side effect	4 (3 %)	5 (3.3 %)	5 (3 %)	
Multi Reason	39 (30.4 %)	56 (37.8 %)	45(26.6 %)	

Table 3: The adherence level among different age groups and educational level

Age	(A) 18-39 years		(B) 40-59 years		(C) >60 years	
	College level (n)	Non-college level (n)	College level (n)	Non-college level (n)	College level (n)	Non-college level (n)
Low adherence	25	5	25	35	24	26
Medium adherence	15	13	25	49	20	36
High adherence	14	2	24	45	17	45

was reported by Yaw and colleagues to be globally weak, especially when these drugs are being used to treat asymptomatic conditions for the sake of preventing future adverse health outcomes. Looking into osteoporosis prophylactic therapies specifically,<sup>[14]</sup> reports that concomitant use of H2 receptor blocker medications or proton pump inhibitors were inversely associated with adherence. Studies were variable in their findings regarding the relationship between glucocorticoids utilization and adherence to osteoporosis medications. Smoking was also found to contribute to non-adherence. On the other hand, experiencing previous fractures and being aware of the results of bone density tests improved patients' adherence to osteoporosis medications.[14-16] incorporated the Extended Health Belief Model to explain patient's attitudes and beliefs towards osteoporosis medications, which may aid clinicians in understanding the motives behind adherence or the lack of it. The main components of this model are the perceived effectiveness of drugs, perceived benefits of taking them, disease threat, concerns about/distrust in medications, and the perceived barriers to taking medications.

This is a cross-sectional study, and data was primarily collected for the purpose of the study. Our study carries the usual limitations of an observational study. Selection bias arises as one of the main concerns of an observational study, and we tried to overcome this by recruiting a high number of participants to guarantee that they are as representative of our population as possible. Ecological fallacy is another concern, and hence clinicians must be careful interpreting results of observational studies and making conclusions about individual patients based on data drawn from a patient population.

#### CONCLUSION

Low adherence was high among vitamin D groups, however; there was no significant difference in medications adherence between the two groups. Forget to take medication was the main reason for nonadherence. This study shows the current adherence status among such patients towards their therapy and promotes future intervention to improve the adherence, and further studies to be done to improve patient's adherence and eventually their health outcome.

#### REFERENCES

- Conti F, Piscitelli P, Italiano G, Parma A, Caffetti MC, Giolli L, et al. Adherence to Calcium and Vitamin D supplementations: results from the ADVICE Survey. Clinical Cases in Mineral and Bone Metabolism 2012;9:157.
- Grant AM, Avenell A, Campbell MK. Oral vitamin D3 and calcium for secondary prevention of low-trauma fractures in elderly people (Randomised Evaluation of Calcium Or vitamin D, RECORD): a randomised placebo-controlled trial. Lancet 2005;365:1621.
- Hong H. Effects of calcium intake, milk and dairy product intake, and blood vitamin D level on osteoporosis risk in Korean adults: analysis of the 2008 and 2009 Korea National Health and Nutrition Examination Survey. Nutr Res Pract 2013;7:409-17.
- Higashiguchi M. Calcium intake and associated factors in a general Japanese population: baseline data of NIPPON DATA80/90 and the National Nutrition Survey. NIPPON DATA80/90. Research Group J Epidemiol 2010;20:3:S549-56.
- Iof bonehealth (2017) Facts and Statistics. International Osteoporosis Foundation. Available at: https://www.iofbonehealth.org/facts-statistics [Accessed 14 Mar 2015].
- Jackson RD, LaCroix AZ, Gass M. Calcium plus vitamin D supplementation and the risk of fractures. N Engl J Med 2006;354:669.
- Porthouse J, Cockayne S, King C. Randomised controlled trial of calcium and supplementation with cholecalciferol (vitamin D3) for prevention of fractures in primary care. BMJ 2005;330:1003.

- Reid IR, Ames RW, Evans MC. Long-term effects of calcium supplementation on bone loss and fractures in postmenopausal women: a randomized controlled trial. Am J Med 1995;98:331-5.
- Reid IR. The role of calcium and vitamin D in the prevention of osteoporosis. Endocrinol Metab Clin North Am 1998;27:389-98.
- Schousboe JT. Adherence with medications used to treat osteoporosis: behavioral insights. Curr Osteoporos Rep 2013;11:21-9.
- Touskova T, Vytrisalova M, Palicka V, Hendrychova T, Chen YT, Fuksa L, et al. Patterns of Nonadherence to Supplementation with Calcium and Vitamin D in Persistent Postmenopausal Women Are Similar at Start and 1 Year Later: A Qualitative Longitudinal Study. Frontiers in Pharmacology 2016;7.
- Włodarek D. Calcium intake and osteoporosis: the influence of calcium intake from dairy products on hip bone mineral density and fracture incidence - a population-based study in women over 55 years of age. Public Health Nutr 2012;10:1.
- 13. World Health Organization. Adherence to long-term therapies: Policy for action 2001.
- Yeaw J, Benner JS, Walt JG, Sian S, Smith DB. Comparing adherence and persistence across 6 chronic medication classes. J Manag Care Pharm 2009;15:728-40.
- Badawi A. Prevalence of vitamin D insufficiency in Qatar: a systematic review. Journal of Public Health Research 2012;1:e36.
- Kanis JA, Melton LJ, Christiansen C, Johnston CC, Khaltaev N. The diagnosis of osteoporosis. J Bone Miner Res 1994;9:1137-41.