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RESEARCH

Please cite this paper as: Ajwah I, Albalawi E, Albalawi Z, Alhawiti W, Alahmari N, Albalawi H, Alanazi T, Alenezi B, Alwabsi W, Alenezi M, Alotaibi O, Rayyan H. Vitamin D supplementation as a fall prevention method: A systematic review. AMJ 2020;13(4):148–155.

https://doi.org/10.35841/1836-1935.13.4.148-155

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ABSTRACT

Background

Falls are common occurrences events leading to remarkable morbidity and mortality. Vitamin D supplementation is often recommended to prevent falls, although vitamin D trials conclude contradictory results.

Aims

This review aimed to summarize the available RCTs that studied the effect of vitamin D administered on falls in older adults.

Methods

Pub Med, Google Scholar, and EBSCO databases were systematically search for relevant articles. The terms vitamin D, falls, ergocalciferols, 25-hydroxy vitamin D, and 1, 25-dihydroxy vitamin D were used. out of Three hundred and four, only eleven fulfilled the inclusion criteria.

Results

In the majority of the randomized control trial, vitamin D supplementation provides protection against falls among the elderly population.

Conclusion

The majority of the studies concluded that vitamin D alone or in combination with calcium was a successful fall reduction method among the elderly.

Key Words

Vitamin D, falls, ergocalciferols

What this study adds:

1. What is known about this subject?

Studies have shown that vitamin D supplementation improves muscle strength and function, thereby reducing the risk of falls.

2. What new information is offered in this study?

Falls reduction can be achieved by daily vitamin D supplementation in doses range between 700-1100IU. While lower daily or higher intermittent dose provide no benefit.

3. What are the implications for research, policy, or practice?

This review did not lead to definite conclusion; an adequate dose of vitamin D supplementation may reduce falls and further study is needed for the definite conclusion.

Background

According to the World Health Organization, among the elderly, falls are common occurrences events leading to remarkable morbidity and mortality. In the United States, following road traffic injuries, falls considered the second



leading cause of unintentional injury.¹

Elderly population particularly at increased risk of vitamin D deficiency as a result of several factors, including but not limited to low dietary intake and diminished sunlight exposure for these reasons, hypovitaminosis D is very frequent in the elder.^{2,3} The association between low level of vitamin D level and increased risk of falls were linked in several studies. For example, the prevalence of low vitamin D in non-faller elderly is ranging between 40–50 per cent while it is reaching up to 70 per cent in fallers.^{4,5}

Vitamin D supplementation has a beneficial effect on muscle strength and balance. Thereby, reduce the occurrence of falls and prevent it is associated morbidity.⁶ On the other hand, insufficient evidence to support the fact that vitamin D supplements especially alone (without supplemental calcium) may have a role in falls prevention.⁷ Given these controversial results, we conducted a systematic review to evaluate the effect of vitamin D administered on falls in older adults.

Method

A systematic electronic search was conducted including the Pub Med, Google Scholar, and EBSCO using the following terms in different combinations vitamin D, falls, ergocalciferols, 25-hydroxy vitamin D, and 1, 25-dihydroxy vitamin D.

Randomized controlled trials in full text that assessed the effects of vitamin D supplementation on risk of falls were included, while abstracts were not included.

Three hundred and four articles were identified, only eleven of them fulfilled the inclusion and exclusion criteria. The abstracts and full texts were screened independently by two authors (HO, AI). The authors extracted the data, and then the author's names, year and region of publication, the study type, period of study, and the result were reported. Tables 1 and 2 The PRISMA Chart was used in the current survey (Figure 1).

Results

Our literature search identified 304 records, 28 of which were reviewed as full-text articles for inclusion. After further exclusions based on our selection criteria, 11 provided sufficient information that was included in the current review. In total, these randomized controlled trials included 6,978 older adults; Study sizes ranged from 122 to 3,717 participants, with the duration of follow-up ranging from one month to 36 months.

Studies prove vitamin D supplementation as falls prevention methods:

A) Vitamin D alone:

Supplementation of vitamin D alone is proven to reduce the fall occurrence was concluded in a double-blind, placebocontrolled trial carried out by Cangussu et al.,⁸ With interventional to control group ratio of 1:1, where the interventional group received a daily dose of 1,000IU of vitamin D.

The dose-dependent effect of vitamin D supplementation on the reduction of falls was evaluated by Broe et al.,⁹ Where four different doses of vitamin D (200IU, 400IU, 600IU, or 800IU) were compared. A lower falls number was noted in participants receiving 800IU of vitamin D.

B) Vitamin D plus calcium

Bischoff et al.,¹⁰ compare the effect of vitamin D supplementation plus calcium versus calcium alone for falls reduction, where the participant randomized to either receive 1.2g calcium plus 800IU cholecalciferol or 1.2g calcium. At the end of 12 weeks treatment period, falls reduced by 49 per cent in favour of vitamin D plus calcium supplementation. Similarly, a three-year double-blinded placebo-controlled trial carried out by conducted by the same author Bischoff et al,.¹¹ Participants received either 700IU of cholecalciferol plus 500mg of calcium citrate malate per day or placebo. Supplementation reduces the odds of falling in ambulatory older women by at least 46 per cent.

In Prince, et al study.¹² A 19 per cent relative risk reduction in the interventional group (ergocalciferol, 1000IU/d plus calcium citrate, 1000mg/d) was concluded from a1-year population-based, double-blind, randomized controlled trial. Alike, 27 per cent at month 12 and 39 per cent at month 20 decrease in the number of subjects with first falls were noted in Pfeifer et al study.¹³

Should Older People in Residential Care Receive Vitamin D to Prevent Falls? In order to answer this question, Flicker et al.¹⁴ conduct two years randomized, placebo-controlled double-blind, trial in which interventional group receive ergocalciferol, initially 10,000IU given once weekly and then 1,000IU daily. Study authors finally conclude that vitamin D supplementation can reduce the number of falls in the elderly.



Studies deny vitamin D supplementation as falls prevention methods:

A)-Vitamin D alone:

Law et al.,¹⁵ Conduct a randomized controlled trial of cluster design. Where the interventional group received ergocalciferol a daily dose of 1,100IU. After 10 months of follow-up, they found non-significant differencebetween the two study arms.

This result was reinforced in a recent study by Uusi-Rasi et al.,¹⁶ participants were randomly allocated to four study groups, including placebo without exercise, vitamin D (800IU/d) without exercise, placebo and exercise, and vitamin D (800IU/d) and exercise. Conclude that vitamin D does not affect the rate of falls.

The effect of high intermittent dose of vitamin D on fall prevention examined as well by Glendenning et al.¹⁷ In which participants receive oral cholecalciferol 150,000IU every three months and at nine months of the trial, no additive benefit of vitamin D was seen.

B) Vitamin D plus calcium:

Burieigh et al.,¹⁸ evaluate the benefit of adding 800 IU of vitamin D to 1.2g of calcium in a randomized, double-blind, controlled study among hospitalized patients, aimed to determine whether vitamin D supplementation has an effective role in falls prevention in older hospital inpatients. Vitamin D did not reduce the number of fallers. Routine supplementation cannot be recommended to reduce falls in this group.

Discussion

The current review summarizes eleven randomized control trials evaluated the hypothesized possible beneficial effect of vitamin D in isolation as well as in combination with calcium supplement on falls reduction.

A significant reduction was concluded in seven studies. Despite the different dosage of vitamin D were used in these studies (200–1000IU), a significant reduction was associated with a dose of 700IU and higher.⁸⁻¹⁴

As an example, Broe et al., studied different doses of vitamin D supplement and only participants who receive 800IU of vitamin D, had a lower incidence of falls compared to those who take a lower dose.⁹

Many evidences suggest that vitamin D in combination with calcium supplementation has a beneficial role in falls prevention. Four out of seven studies combined calcium to

the vitamin D supplement.¹⁰⁻¹³

As an illustration, Bischoff et al. found a reduction of falls with vitamin D - calcium supplement in the community elderly.^{10,11} But in these studies, the effect of calcium and vitamin D could not be separated. In another systemic review, Haiting and Qingjiang found that combined calcium plus vitamin D supplementation are statistically significantly associated with a reduction in fall risks across various populations. Finally, the current evidence for the effect of supplementary vitamin D alone on fall outcomes is limited.¹⁹ On the other hand, in this review, four studies found that no reduction in risk of falls neither with vitamin D alone nor in combination with calcium.¹⁵⁻¹⁸

In one of these studies. Glendenning et al used higher doses of vitamin D (cholecalciferol 150,000IU every three months) and found that it is ineffective or deleterious on the risk of falls.¹⁷ This result was also supported in a randomized control trial conducted by Bischoff et al., in 2016 when they compared low versus high doses of vitamin D. The study showed an increased risk of falls were associated with the usage of higher doses.²⁰

The results of the pooled studies are limited and cannot be generalized, as the studies participants were mainly community dwelling and hospitalized elderly.

To summarize, controversial results were found between studies. However. These contradictory results may be related to the variation in the baseline serum vitamin D levels between studies participants. Furthermore, the usage of different vitamin D doses among the interventional group.

Conclusion

In conclusion, the majority of the studies concluded that vitamin D alone or in combination with calcium was a successful fall reduction method among the elderly.

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ACKNOWLEDGEMENTS

We deeply thank Mrs. Amal Bedaiwi for her grateful advice and assistance in keeping our progress on schedule.

CONFLICTS OF INTEREST

The authors declare that they have no competing interests.

FUNDING

None



Figures and Tables

Table 1: Randomized control trials with proven effect of vitamin D supplementation to reduce the risk of falls

Author—Year Country	Methods	Results
	Study design: Randomized, double-blind, placebo-controlled trial	Study completer: 140 Women
Cangussu et al. ⁸	Inclusion criteria: Date of last menstruation at least 12 months before, aged 50 to 65 years, history of falls in the last 12 months, and with bone mineral density (BMD) higher than S1.5 SD.	Primary objective: The occurrence of falls and falls recurrence was higher in the placebo group compared to the VITD group (<i>P</i> <0.001).
2015	Interventional group: 1,000IU/day/orally	Limitation: Representative study
Brazil	Follow-up: 9 months	
	Primary objective: To evaluate the effect of isolated vitamin D supplementation on the occurrence of falls	Conclusion: Isolated vitamin D supplementation for 9 months resulted in a lower incidence of falls.
	Study design: Double-blind, placebo- controlled trial	Study completer: 124 participants
Broe et al. ⁹	Inclusion criteria: Life expectancy of 6 months, ability to swallow medication.	Primary objective: Low rate of falls among residents in
2007	Interventional group: Vitamin D (200IU, 400IU, 600IU, 800IU)	800IU vitamin D supplement group.
Massachusetts	Follow-up: 5 months	Limitation: use on multivitamins and result cannot be generalized.
	Primary objective: occurrence of falls.	Conclusion: Adequate vitamin D supplementation in elderly nursing home residents could reduce the numbe of falls experienced by this high fall risk group.
	Study design: Randomized double-blind trial	Study completer: 122 Participants
Bischoff et al. ¹⁰	Inclusion criteria: People age 60 or older and able to walk 3m with or without a walking aid, without primary hyperparathyroidism, renal insufficiency, hypocalcaemia, hypercalciuria and fracture or stroke in the last 3 months.	Primary objective:
2003	Interventional group: 1200mg calcium plus 800IU cholecalciferol	Number of falls reduced by 49% in the Interventional group.
Switzerland	Follow-up: 6 - week pre-treatment period and a 12 - week treatment period	Limitation: High drop-out rate and result cannot be generalized.
	Primary objective: The number of falls per person during the treatment period	Conclusion: Vitamin D reduces the risk of falls.
	Study design: Randomized, double-blind placebo-controlled trial	Study completer: 445 individuals
Bischoff-Ferrari et al. ¹¹	Inclusion criteria healthy ambulatory men and women 65 years or older and living in the community.	Primary objective: Cholecalciferol-calcium significantly reduced the odds of falling in women, but not in men.
2006	Interventional group: Cholecalciferol (vitamin D3; 700IU/d) plus calcium (500mg/d).	Limitation: The result cannot be generalized.
Switzerland	Follow-up: 3 years	Conclusion: Long-term dietary cholecalciferol-calcium supplementation reduces the odds of falling in ambulatory older women by 46%.
	Primary objective: occurrence of falls.	



	Study design: Population-based, double- blind, randomized controlled trial	Study completer: 302 Participants
Prince et al. ¹²	Inclusion criteria: Women aged 70 to 90 years living in Perth, Australia, with a serum 25-hydroxyvitamin D concentration of less than 24.0ng/mL and a history' of falling in the previous year.	Primary objective: Ergocalciferol therapy reduced the risk of having at least 1 fall over 1 year
2008	Interventional group: Ergocalciferol, 1,000IU/d plus calcium citrate 1000mg/d.	Limitation: inequality between the study arms.
Australia	Follow-up: 12 months	Conclusion: Ergocalciferol in addition to the calcium result in a 19% reduction in the relative risk of falling.
	Primary objective: occurrence of falls.	
	Study design: Double-blind, controlled trial.	Study completer: 242 subjects
	Inclusion criteria: Healthy ambulatory women and men 70 years of age or older with 25-(OH)D serum level below 78nmol/l	Primary objective: calcium plus vitamin D resulted in a significant decrease in the number of falls.
Pfeifer et al. ¹³	Interventional group: 1000mg of calcium plus 800 IU of vitamin D per day.	Limitation: The result cannot be generalized.
	Follow-up: 20 months	Conclusion: Combined calcium and vitamin D supplementation proved superior to calcium alone in reducing the number of falls.
	Primary objective: occurrence of falls.	
	Study design: Randomized, placebo- controlled double-blind, trial.	Study completer: 625
Flicker et al. ¹⁴	Inclusion criteria: serum 25-hydroxyvitamin D levels between 25 and 90nmol/L.	Primary objective: Incident rate ratio for falls of 0.63, an odds ratio (OR) forever falling of 0.70.
Australia	Interventional group: Ergocalciferol, initially 10,000IU given once weekly and then 1,000IU daily.	Limitation: High drop-out rate.
2005	Follow-up: 24 months	Conclusion: Vitamin D supplements for 2 years can reduce the incidence of falls in the elderly.
	Primary objective: Falls and fractures occurrence.	

Table 2: Randomized Control Trial that showed no role of vitamin D in reduces the risk of falls

Author—Year Country	Methods	Results
	Study design: Randomized controlled trial of cluster design	Study completer: 3,717 participating residents
Malcolm Law et al. ¹⁵	Inclusion criteria: Age ≥60 years old and Not taking calcium/vitamin D or drugs that increase bone density.	Primary objective: Falls among the study arms not statistically significant.
2006	Interventional group: Ergocalciferol 2.5mg every 3 months.	Limitation: Reporting error.
British	Follow-up: 10 months	Conclusion: No evidence that vitamin D prevents fractures or falls in elderly people in care home accommodation
	Primary objective: Falls and fracture.	



	Study design: Randomized, double-	Study completer: 370 participants
	blind, placebo-controlled trial.	
Kirsti Uusi-Rasi et al. ¹⁶	Inclusion criteria: At least 1 fall during the previous year, no use of vitamin D supplements, and no contraindication to exercise.	Primary objective: No falls reduction.
2015	Interventional group: Four study groups, including placebo without exercise, vitamin D (800IU/d) without exercise, placebo and exercise, and vitamin D (800IU/d) and exercise.	Limitation: Elevated baseline 25 (OH) D levels.
Finland	Follow-up: 2 years.	Conclusion: vitamin D does not affect the rate of falls.
	Primary objective: monthly reported falls.	
	Study design: Randomized, double-blind, placebo-controlled, prospective, parallel study	Study completer: 686 participants
Glendenning et al. ¹⁷	Inclusion criteria: age over 70 years, registration with a general practitioner, and likelihood, in the investigators' opinion, of attending four study visits over 9 months	Primary objective: Faller rates in the two groups did not differ
2012	Interventional group: Cholecalciferol 150,000IU of vitamin D every 3 months	Limitation: The result cannot be generalized.
Australia	Follow-up : 9 months	Conclusion: Intermittent large doses of vitamin D are ineffective or have a deleterious effect on falls
	Primary objective: Falls occurrence.	
	Study design: Prospective, randomized, double-blind, controlled study.	Study completer: 205 Participants.
Burleigh et al. ¹⁸	Inclusion criteria: Patients transferred or admitted into the general assessment and rehabilitation wards in an acute geriatric unit were screened for eligibility for trial entry.	Primary objective: no reduction in the number of falls
2007	Interventional group: vitamin D 800IU plus calcium 1,200mg	Conclusion: Population of geriatric hospital inpatients, vitamin D did not reduce the number of fallers. Routine supplementation cannot be recommended to reduce falls in this group.
	Follow-up: 1 month	
	Primary objective: Falls occurrence.	



Figure 1: Flow diagram through the different phases of the systematic review (PRISMA flowchart)

