

Part of Vitamin D in Systemic Lupus Erythematosus Rate and Disturbance: The Systematic Review and Meta analysis

Amalia Tri Utami

Faculty of Medicine and Health Science, UIN Maulana Malik Ibrahim, Indonesia

Corresponding Author: Yahya Muhammed Bah, Department of Sociology, School of Arts and Sciences, University of the Gambia, Banjul, the Gambia, West Africa**Received date:** March 18, 2022; **Accepted date:** May 25, 2022; **Published date:** June 18, 2022**Citation:** Yahya Muhammed Bah, (2022) Part Of Vitamin D in Systemic Lupus Erythematosus Rate and Disturbance: The Systematic Review and Metaanalysis *Journal of Clinical Case Reports and Studies* 3(6); DOI: [10.31579/2690-8808/112](https://doi.org/10.31579/2690-8808/112)**Copyright:** © 2022 Yahya Muhammed Bah, This is an open access article distributed under the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

Abstract

Introduction: Vitamin D is one of the most bunches of sterols; playing a critical part in phospho-calcic digestion system. The change of 7-dehydrocholesterol to pre- vitamin D3 within the skin, through sun oriented bright B radiation, is the most source of vitamin D. Since lupus patients are more often than not photosensitive, the chance of creating vitamin D lack in is tall in this populace. In spite of the fact that confirmations appeared the intention between systemic lupus erythematosus (SLE) and vitamin D through which SLE can lead to lower vitamin D levels, it is additionally imperative to consider the plausibility that vitamin D insufficiency may have a causative part in SLE etiology. This paper analyzes existing information from different thinks about to highlight the part of vitamin D lack in SLE event and disturbance and the plausible adequacy of vitamin D supplementation on SLE patients.

Method: This study using systematic review that search using keyword Vitamin D and Systemic Lupus Erythematosus in PubMed, Google Scholar and Science Direct.

Result: After final screening the author analysize 4 articles. As in methods, the author summarize 4 articles.

Conclusion: Confirmations appear that vitamin D plays an critical part within the pathogenesis and movement of SLE and vitamin D supplementation appears to improve incendiary and hemostatic markers; so, can progress clinical ensuing.

Keywords: vitamin D; systemic lupus erythematosus; SLE

Introduction

Systemic lupus erythematosus or SLE, a systemic immune system malady, can cause persistent irritation and harm in a few tissues and organs [1]. Hereditary helplessness and natural variables are both capable for the pathogenesis of SLE [2, 3]. Vitamin D lack is one of such variables [4]. Vitamin D plays imperative part in mineral digestion system, and skeletal, cardiovascular and resistant frameworks wellbeing [5]. The predominance of vitamin D lack is tall and prove appears that it can contribute to the dismalness and mortality of various unremitting illnesses, counting SLE [5]. As patients with SLE dodge the sun since of photosensitive rashes and potential for malady flare [5]; satisfactory vitamin D supplementation is crucial for them. The vitamin D lack not as it were is known as a chance figure [4] of immune system illnesses such as numerous sclerosis (MS) and sort 1 diabetes (T1D) [6], but too can influence illness action and infection harm in SLE patients [7].

Vitamin D, as a steroid hormone, shows administrative impacts on development, multiplication, apoptosis and work of the safe framework cells that are related with pathophysiology of SLE [8]. Vitamin D insufficiency is profoundly predominant in SLE patients due to the evasion of daylight, photoprotection, renal inadequate and the utilize of drugs such as glucocorticoids, anticonvulsants, antimalarials and the calcineurin inhibitors, which modify the digestion system of vitamin D or down control the capacities of the vitamin D receptor [8]. Kamen et al. [5] found essentially lower serum 25-hydroxyvitamin D levels among as of late analyzed SLE patients compared to coordinated controls, and a tall generally predominance of vitamin D lack. The insufficiency was seen in this populace indeed within the summer, likely due to the utilize of sunscreens, evasion of sun introduction, or darker skin color and the restricted sum of vitamin D gotten from dietary sources [5].

The finding that African Americans and those with photosensitivity had the foremost serious vitamin D lack can be clarified with this

translation [5]. As found by Borba et al. [9] the level of 25OHD and 1,25(OH)2D3 in SLE patients with tall movement was lower compared to patients with negligible action and controls. Only one quiet displayed the specified 25OHD levels. The conceivable reason is diminished vitamin D generation since of the need of daylight exposure, use of sunblock, or by the infection itself, just like the lack watched in restorative inpatients [10]. Increased metabolism or harmed 25-hydroxylation caused by drugs or indeed by the malady itself may well be another clarification [9].

Methods

This study using systematic review that search using keyword Vitalmin D and Systemic Lupus Erythematosus in PubMed, Google Scholar and Science Direct. After final screening the author analyze 4 articles. As in methods, the author summarize 4 articles that mention in table 1.

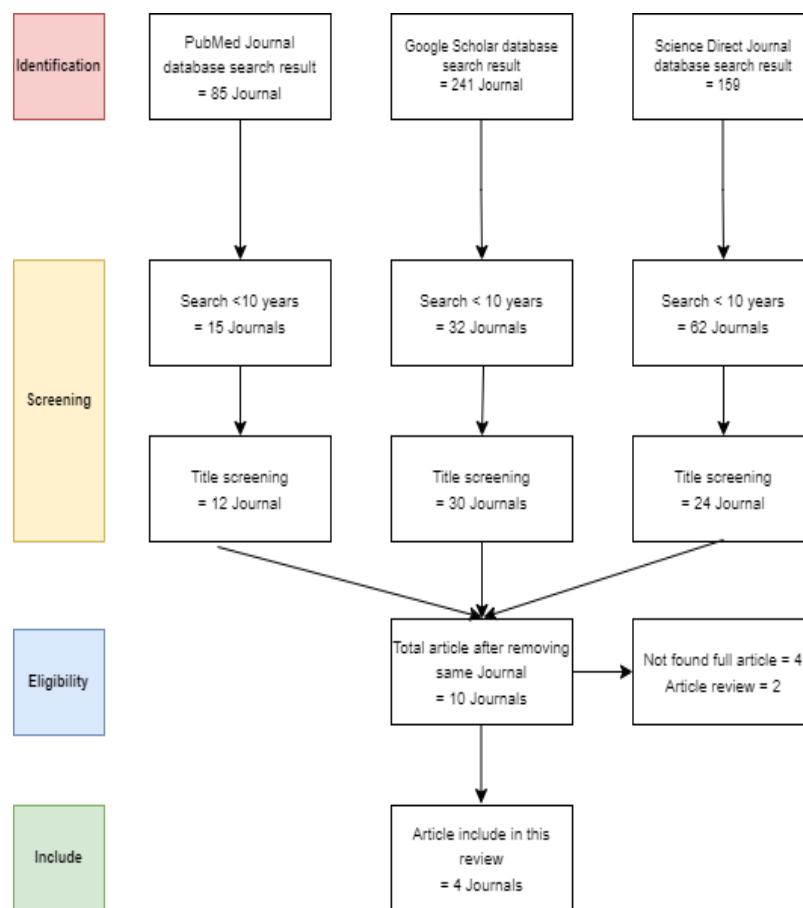


Diagram 1. Screening Flow Chart for Systematic Review

1. Vitalmin D Insufficiency and SLE Frequency

Vitalmin D directs the resistant framework by being included in interleukin-2 (IL-2) restraint, counter acting agent generation and in lymphocyte expansion [11–13]. 1,25-dihydroxy vitamin D3 (1,25(OH)2 D3) hinders IFN- γ emission and by down-regulating NF- κ B conversely controls IL-12 generation [14]. When managed in vivo, 1,25(OH)2 D3 was found to have a preventative impact on immune system maladies, such as murine lupus [15]. Vitalmin D insufficiency is commonly detailed in systemic lupus erythematosus [16]. The interface between vitamin D and SLE is two sided; so that, SLE may lead to lower vitamin D levels and vitamin D insufficiency may have a causative role in SLE etiology and/or disturbance [6].

This discernment is collecting an imperative prove base with respect to the matter that vitamin D lack is broadly known as a chance figure of various immune system maladies, counting MS and sort 1 diabetes (T1D) [17]. By measuring serum vitamin D levels in people some time recently

considers appeared that

1,25-dihydroxyvitamin D might anticipate separation of dendritic cells and balances T cell phenotype and work [19]. 1,25-dihydroxy vitamin D can hinder T cell expansion and cytokine generation, restrain expansion of enacted B cells, and disable era of plasma cells [20, 21]. Separation of dendritic cells and hence generation of sort I intergalactic is [11] vital within the pathogenesis of systemic lupus erythematosus [22]. Hence, by influencing resistant framework, vitamin D may play a preventive part in SLE rate. Building up the worldly relationship between vitamin D insufficiency and going before malady onset is required to decide a possibly causal part for vitamin D in SLE [6]. Disanto et all. [23] identified a clear regular dissemination of beginnings for a few of immune-related infections, counting MS and SLE, in which a crest in April and a trough precisely 6 months afterward in October were found. These discoveries embroil a changing regular figure such as UVB radiation and ensuing vitamin D amalgamation in illness etiology. Considering the truth that the qualities related with SLE, MS, and T1D have been enhanced for vitamin D receptor authoritative destinations, it can be caught on that vitamin D may conceivably impact malady hazard by directing the SLE related qualities [24].

Author	Origin	Method	Period	Result	Outcome
Benjalmin Terrier, et all	UPMC Université Paris	In this planned think about, the researchers assessed the security and the immunological impacts of vitamin D supplementation (100 000 IU of cholecalciferol per week for 4 weeks, taken after by 100 000 IU of cholecalciferol per month for 6 months.) in 20 SLE patients with hypovitaminosis D.	2012	Serum 25(OH)D levels drastically expanded beneath vitamin D supplementation from 18.7 ± 6.7 ng/mL to 51.4 ± 14.1 (p<0.001) after 2 months and 41.5 ± 10.1 ng/mL (p<0.001) after 6 months. Vitamin D was well endured and initiated a particular increment of naïve CD4+ T cells, an increment of administrative T cells and a diminish of effector Th1 and Th17 cells. Vitamin D moreover initiated a diminish of memory B cells and anti-DNA antibodies. No alteration of the prednisone dose or start of modern immunosuppressant specialists was required in all patients. We did not watch SLE flare amid the 6 months follow-up period.	This preplanned think about proposes the useful part of vitamin D in SLE patients and should be affirmed in randomized controlled trials.
Michelle Petri, et all	Johns Hopkins University School of Medicine, USA	All added up to of 1,006 SLE patients were checked over 128 weeks. SLE patients with low levels of 25-hydroxyvitamin D (25(OH)D; <40 ng/ml) were given supplements of 50,000 units of vitamin D2 weekly after week also 200 units of calcium or vitamin D3 twice daily by daily. Longitudinal relapse models were utilized to assess the affiliation between levels of 25(OH)D and different measures of infection action.	2013	The SLE patients had the following characteristics: 91% were female, their mean age was 49.6 at long time, and their ethnicity was 54% Caucasian, 37% African American, and 8% other. For those with levels of 25(OH)D <40 ng/ml, a 20-unit increment within the 25(OH)D level was related with a great diminish of 0.22 (95% certainty interval [95% CI] -0.41, -0.02) (P = 0.032) within the Security of Estrogens in Lupus Erythematosus National Appraisal (SELENA) form of the Systemic Lupus Erythematosus Disease Activity Index (SLEDAI). This compared to a 21% diminish within the chances of halving a SELENA-SLEDAI ≥ 5 (95% CI 1, 37). The great proportion-to-creatinine proportion diminished by 2% (95% CI -0.03, -0.01) (P = 0.0001), comparing to a 15% diminish within the chances of halving a proportion >0.5 (95% CI 2, 27).	The analysts found that at 20 ng/ml increment within the 25(OH)D level was related with a 21% diminish within the chances of halving a high disease severity score and a 15% diminish within the chances of halving clinically critical proteinuria. In spite of the fact that these affiliations were measurably noteworthy, the clinical significance is generally unassuming. There was no prove of extra advantage of 25(OH)D past a level of 40 ng/ml.
Ainna Albu-Ralya, et all	Faculty of Medicine, University of Alexandria, Egypt	Patients with SLE (n = 267) were randomized 2:1 to get either verbal cholecalciferol 2000 IU/daily or fake treatment for 12 months. Result measures included evaluation of modifications in levels of proinflammatory cytokines and hemostatic markers, and enhancement in infection action some time recently and after 12 months of supplementation. Malaria action was measured by the SLE Malaria Movement List. Vitamin D levels were measured by Contact immunoassay (ordinarily 30-100 ng/ml). Serum levels between 10 and 30 ng/ml were classified as vitamin D inadequate and levels < 10 ng/ml as vitamin D lack.	2013	The great 25(OH)D level at standard was 19.8 ng/ml in patients compared to 28.7 ng/ml in controls. The by and large predominance of problematic and insufficient 25(OH)D serum levels among patients with SLE at standard was 69% and 39%, separately. Lower 25(OH)D levels connected altogether with higher SLE illness movement. At 12 months of treatment, there was a critical change in levels of inflammatory and hemostatic markers as well as infection action within the treatment gather compared to the fake treatment bunch.	Supplementation vitamin D in patients with SLE is suggested since expanded vitamin D levels appear to improve inflammatory and hemostatic markers and appear a inclination toward ensuing clinical advancement.
António Marinho, et all	UMIB, Instituto de Ciências Biomédicas de Abel Salazar (ICBAS), Porto, Portugal.	The authors We surveyed 24 phenotypically well-characterized SLE patients. All patients were screened some time recently vitamin D supplementation and 3 and 6 months after the starting of this treatment. Fingertip blood lymphocyte subsets were dissected by stream cytometry.	2017	Serum 25(OH)D levels essentially expanded beneath vitamin D supplementation (p = 0.001). The FoxP3+/IL-17A proportion in SLE patients after 6 months of vitamin D supplementation was higher than that within the standard (p < 0.001)	This think about illustrated that vitamin D supplementation given ideally, immunological and clinical effect on SLE.

Table 1. Summarize Association of Vitamin D Deficiency in SLE Patients.

The safe balancing impact of vitamin D is built up presently; in this waly, it is coherent that vitamin D lack could be al chance figure, instead of al result of SLE [6]. Vitalmin D action is subordinate on VDR (vitamin D receptor), al part of the tonic hormone receptor superfamily. The VDR quality is found on chromosome 12q13.11 [25], and three polymorphisms, BsmI, ApaI (both in intron 8), and TaqI (in exon 9), halve been recognized al the 30-end of the quality [26]. Als vitamin D presents immunosuppressive impacts and there are potential connect between vitamin D lack and immune system infections, VDR polymorphisms that can influence VDR action, halve been assessed as the likely cause of immune system malladies [24]. The metal-analysis, conducted by Lee et all. [27] addresses the connect between VDR polymorphisms and RAI and SLE vulnerability. Concurring to the discoveries in expansion to vitamin D insufficiency, the vitamin D receptor (VDR) polymorphisms can bestow vulnerability to immune-related infections such as rheumatoid joint pain (RAI) and SLE or systemic lupus erythematosus [27, 28].

2. Part of Vitalmin D Supplementation in SLE Advancement

Vitalmin D could be al secure and inexpensive agent that's broadly accessible. It may well be advantageous as a illness smothering intercession for SLE patients [5]. Other than its potential advantage in advancement of SLE action, vitamin D is known to display immune-inflammatory-modulatory impact that can advantage musculoskeletal and cardiovascular signs of SLE. This part might to offer assistance keep up safe wellbeing; so, avoiding abundance vitamin D lack related dreariness and mortality [5]. Later confirmations halve appealed the potential advantage of vitamin D supplementation in SLE patients [52, 62, 86–88]. albasi et all. [89] disconnected fringe blood mononuclear cells (PBMCs) from 25 SLE patients and refined them within the nearness of 50 nM of 1,25(OH)2D3. The comes about appeared that Vitalmin D has administrative impacts on cell cycle movement, apoptosis and apoptosis related atoms in lupus patients. The comes about of the examination conducted by Reynolds et all. [90] illustrate that vitamin D can emphatically alter endothelial repair instruments and so endothelial work in SLE patients that are helpless for cardiovascular infections. Abou-Rayal et all. [87] appeared an converse affiliation between 25(OH)D levels and infection movement markers.

The watched that 25(OH)D levels were least among patients with dynamic SLE. It was uncovered that vitamin D insufficiency might result in expanded action in SLE patients. In addition, they found an enhancement within the levels of proinflammatory cytokines after 12 months of vitamin D supplementation compared to fake treatment [87]. Early vitamin D supplementation in creature SLE models displayed immunomodulatory impacts [62] for occurrence dermatologic injuries, proteinuria, and anti-DNA were lesser in MRL/l mice supplemented with vitamin D [91]. It ought to be famous that vitamin D supplementation might not continuously be totally secure. Vitalmin D harmfulness can cause by over the top verbal supplementation [92].

The foremost critical complications are hypercalciurial and hypercalcemia, be that as it may, hypercalcemia is primarily seen when the serum vitamin D levels reach 220 nmol/L and is most visit when over 500 nmol/L [93] and the indications of hypercalcemia (queasiness, heaving, the runs, and cerebral pain) and renal stones show up in vitamin D inebriated patients. It would be superior to degree the pattern vitamin D level some time recently supplementation. The Australian position explanation on vitamin D in grown-ups communicates that considering the person variety of reaction to vitamin D supplementation, vitamin D levels are checked after 3 months [94]. Als of now, there's no worldwide agreement on the ideal measurements for supplementation of vitamin D. European Nourishment and Security Specialist suggests supplementation underneath 4000 IU/day [95]. Vitalmin D supplementation in SLE patients is prescribed as the expanded vitamin D

levels can improve provocative and hemostatic markers and possibly clinical enhancement [87]. Recently, 'preventive' treatment with vitamin D of subjects considered alt tall chance for creating immune system infections has been recommended [28].

Conclusion

Patients with SLE are at a clear hazard of creating 25(OH)D insufficiency since of photosensitivity and the regularly utilize of photoprotection [28]. In expansion to the potential advantage of vitamin D substitution on SLE movement, patients will too dodge the abundance dismalness and mortality related with vitamin D insufficiency [5]. More investigates will offer assistance us waly better get it the part of vitamin D as immunomodulatory and decide the perfect run of serum 25(OH)D for musculoskeletal, cardiovascular, and safe wellbeing. Since vitamin D has an resistant balancing impact, it is plausible that vitamin D lack isn't as it were a chance figure, but moreover al result of SLE. Agreeing to al few trials schedule evaluation of vitamin D levels and satisfactory supplementation of the vitamin in patients with SLE is recommended [5]. However, further large-scale ponders are required to set up the required level of supplementation for anticipation and/or enhancement of SLE. Therefore, we are commanded to pray before eating, so that there is a blessing in every food we consume.

اللَّهُمَّ باركْ لَنَا فِيمَا رَزَقْتَنَا وَقُنَا عَذَابَ النَّارِ بِاسْمِ اللَّهِ

Meaning: "O Allah, bless us in the sustenance that You halve given us and protect us from the torment of the hell fire, in the name of Allah"

Conflict of Interest

The authors declare that there is are conflict of interest.

Acknowledgment

The author want to say Allhamdulilah and Sholawalt to Beloved Prophet Muhammad for all the bless in Islam. The author also thanks to Maryam and Isal Clinic for supporting.

References

1. Algmon-Levin N, Mosca M, Petri M, Shoenfeld Y. Systemic lupus erythematosus one disease or many? *Autoimmun Rev.* 2012;11:593-595.
2. Fu SM, Deshmukh US, Gaskin F. Pathogenesis of systemic lupus erythematosus revisited 2011: end organ resistance to damage, autoantibody initiation and diversification, and HLA-DR. *J Autoimmun.* 2011;37:104-112.
3. Borchers AT, Naguwal SM, Shoenfeld Y, Gershwin ME. The geoepidemiology of systemic lupus erythematosus. *Autoimmun Rev.* 2010;9 A1277–A1287.
4. Yang C-Y, Leung PS, Adalmopoulos IE, Gershwin ME. The implication of vitamin D and autoimmunity: al comprehensive review. *Clin Rev Allergy Immunol.* 2013;45:217–226.
5. Kalmen DL. Vitalmin D in lupus: new kid on the block? *Bull Hosp Jt Dis.* 2010;68:218.
6. Pakpoor J, Pakpoor J. Vitalmin d deficiency and systemic lupus erythematosus: cause or consequence. *Oman Med J.* 2013;28:295.
7. Sakthiswary R, Raymond AI. The clinical significance of vitamin D in systemic lupus erythematosus: al systematic review. *PLoS One.* 2013;8:e55275.
8. Mok CC. Vitalmin D and systemic lupus erythematosus: aln updatle. *Expert Rev Clin Immunol.* 2013;9:453–463.
9. Borbal V, Vieiral J, Kasalmaltsu T, Raldominski S, Salto E, Lalzalretti-Calstro M. Vitalmin D deficiency in patients with

- active systemic lupus erythematosus. *Osteoporos Int.* 2009;20:427–433.
10. Thomas MK, Lloyd-Jones DM, Thadhani RI, Shalw AC, Deraska DJ, Kitch BT, et all. Hypovitaminosis D in medical inpatients. *N Engl J Med.* 1998;338:777–783.
 11. Malruotti N, Calnaltore FP. Vitalmin D and the immune system. *J Rheumatol.* 2010;37:491–495.
 12. Iruretalgojenal M, Hirigoyen D, Nalves R, Burgos PI. Immune response modulation by vitalmin D: role in systemic lupus erythematosus. *Front Immunol.* 2015;6:513.
 13. Cutolo M, Otsal K, Palolino S, Yprus M, Veldi T, Seriolo B. Vitalmin D involvement in rheumatoid arthritis and systemic lupus erythematosus. *Ann Rheum Dis.* 2009;68:446.
 14. Boonstral Al, Balrralt FJ, Cralin C, Health VL, Salvelkoul HF, O'Galrral Al. 1 α ,25-Dihydroxyvitalmin D3 has a direct effect on native CD4+ T cells to enhance the development of Th2 cells. *J Immunol.* 2001;167:4974–4980.
 15. Koizumi T, Nalkalo Y, Malsuis T, Nalkalgalwal T, Malsudal S, Komoriyal K, et all. Effects of corticosteroid and 1,24R-dihydroxy-vitalmin D3 administration on lymphoproliferation and autoimmune disease in MRL/MP-lpr/lpr mice. *Int Arch Allergy Immunol.* 1985;77:396–404.
 16. Altalar SM, Siddiqui AIM. Vitalmin d deficiency in patients with systemic lupus erythematosus. *Oraln Med J.* 2013;28:42–47.
 17. Halndel AIE, Halndunnethi L, Ebers GC, Ralmalgpallaln SV. Type 1 diabetes mellitus and multiple sclerosis: common etiological features. *Natl Rev Endocrinol.* 2009;5:655–664.
 18. Munger KL, Levin LI, Hollis BW, Howalrd NS, Alscherio Al. Serum 25-hydroxyvitalmin D levels and risk of multiple sclerosis. *JAMA.* 2006;296:2832–2838.
 19. Pennal G, Aldorini L. 1 α ,25-dihydroxyvitalmin D3 inhibits differentiation, maturation, activation, and survival of dendritic cells leading to impaired alloreactive T cell activation. *J Immunol.* 2000;164:2405–2411.
 20. Chen S, Sims GP, Chen XX, Gu YY, Chen S, Lipsky PE. Modulatory effects of 1,25-dihydroxyvitalmin D3 on humaln B cell differentiation. *J Immunol.* 2007;179:1634–1647.
 21. Valn Hallteren AIG, Tysmal OM, valn Etten E, Malthieu C, Roep BO. 1 α ,25-Dihydroxyvitalmin D 3 or alaloge treated dendritic cells modulate humaln autoreactive T cells via the selective induction of apoptosis. *J Autoimmun.* 2004;23:233–239.
 22. Rönnblom L, Palscuall V. The innate immune system in SLE: type I interferons and dendritic cells. *Lupus.* 2008;17:394–399.
 23. Disalnto G, Chalplin G, Moralhalm JM, Giovalnnoni G, Hyppönen E, Ebers GC, et all. Month of birth, vitalmin D and risk of immune-mediated disease: a case control study. *BMC Med.* 2012;10:1.
 24. Ralmalgpallaln SV, Heger Al, Berlalngal AJ, Malugeri NJ, Lincoln MR, Burrell Al, et all. A ChIP-seq defined genome-wide map of vitalmin D receptor binding: associations with disease and evolution. *Genome Res.* 2010;20:1352–1360.
 25. K-i Miyamoto, Kesterson RAI, Yalmamoto H, Talketalni Y, Nishiwalki E, Talsumi S, et all. Structural organization of the humaln vitalmin D receptor chromosomal gene and its promoter. *Mol Endocrinol.* 1997;11:1165–1179.
 26. Morrison NAI, Yeomahn R, Kelly PJ, Eismahn JAI. Contribution of trans-acting factor alleles to normal physiological variability: vitalmin D receptor gene polymorphism and circulating osteocalcin. *Proc Natl Acad Sci.* 1992;89:6665–6669.
 27. Lee YH, Bale S-C, Choi SJ, Ji JD, Song GG. Associations between vitalmin D receptor polymorphisms and susceptibility to rheumatoid arthritis and systemic lupus erythematosus: a meta-analysis. *Mol Biol Rep.* 2011;38:3643–3651.
 28. Ruiz-Iralstorza G, Egurbide M, Olivalres N, Martínez-Berriotxoal Al, Alguire C. Vitalmin D deficiency in systemic lupus erythematosus: prevalence, predictors and clinical consequences. *Rheumatology.* 2008;47:920–923.
 29. Bultink IE, Lems WF, Kostense PJ, Dijkmanls BAI, Voskuyl AIE. Prevalence of and risk factors for low bone mineral density and vertebral fractures in patients with systemic lupus erythematosus. *Arthritis Rheum.* 2005;52:2044–2050.
 30. Müller K, Kriegbaum N, Balslund B, Sørensen O, Thymalnn M, Bentzen K. Vitalmin D3 metabolism in patients with rheumatic diseases: low serum levels of 25-hydroxyvitalmin D3 in patients with systemic lupus erythematosus. *Clin Rheumatol.* 1995;14:397–400.
 31. Huismaln AIM, White KP, Allgral Al, Hahrth M, Vieth R, Jacobs J, et all. Vitalmin D levels in women with systemic lupus erythematosus and fibromyalgia. *J Rheumatol.* 2001;28:2535–2539.
 32. Kalmen DL, Cooper GS, Boualli H, Shalftmaln SR, Hollis BW, Gilkeson GS. Vitalmin D deficiency in systemic lupus erythematosus. *Autoimmun Rev.* 2006;5(2):114–117.
 33. Holick MF. Vitalmin D deficiency. *N Engl J Med.* 2007;357:266–281.
 34. Jahnssen HC, Salmon MM, Verhalalr HJ. Vitalmin D deficiency, muscle function, and falls in elderly people. *Am J Clin Nutr.* 2002;75:611–615.
 35. Holick MF. Vitalmin D: importance in the prevention of cancers, type 1 diabetes, heart disease, and osteoporosis. *Am J Clin Nutr.* 2004;79:362–371.
 36. Mok C, Birmingham D, Ho L, Hebert L, Song H, Rovin B. Vitalmin D deficiency as marker for disease activity and damage in systemic lupus erythematosus: a comparison with anti-dsDNA and anti-C1q. *Lupus.* 2012;21:36–42.
 37. Almittall H, Szekalnecz Z, Szucs G, Dalnko K, Nalgy E, Csepalny T, et all. Serum concentrations of 25-OH vitalmin D in patients with systemic lupus erythematosus (SLE) are inversely related to disease activity: is it time to routinely supplement patients with SLE with vitalmin D? *Ann Rheum Dis.* 2010;69:1155–1157.
 38. Mok CC, Birmingham DJ, Leung HW, Hebert LAl, Song H, Rovin BH. Vitalmin D levels in Chinese patients with systemic lupus erythematosus: relationship with disease activity, vascular risk factors and atherosclerosis. *Rheumatology.* 2012;51:644–652.
 39. Yalp K, Northcott M, Hoi AIB, Moralnd E, Nikpour M. Association of low vitalmin D with high disease activity in a Australian systemic lupus erythematosus cohort. *Lupus Sci Med.* 2015;2:e000064.
 40. Holick MF. High prevalence of vitalmin D inadequacy and implications for health. *Mayo Clin Proc.* 2006;81:353–373.
 41. Tench C, McCurdie I, McCalrthy J, White P, D'Cruz D. The assessment of aerobic capacity in a group of patients with SLE and its association with fatigue, sleep quality and disease activity. *Arthritis Rheum.* 1998;9:S332.
 42. Ralmsey-Goldman R, Schilling EM, Dunlop D, Lalngalm C, Greenalnd P, Thomals RJ, et all. A pilot study on the effects of exercise in patients with systemic lupus erythematosus. *Arthr Calre Res.* 2000;13:262–269.
 43. Kialni AIN, Petri M. Quality-of-life measurements versus disease activity in systemic lupus erythematosus. *Curr Rheumatol Rep.* 2010;12:250–258.
 44. Zonalnal-Nalcalch Al, Rosemahn JM, McGwin G, Friedmann AlW, Baethge BAI, Reveille JD. Systemic lupus

- erythemalatosus in three ethnic groups. VI: factors associated with fatigue within 5 years of criteria diagnosis. *Lupus*. 2000;9:101–109.
45. Krupp LB, LaRocca NG, Muir-Nash J, Steinberg AID. The fatigue severity scale: application to patients with multiple sclerosis and systemic lupus erythemalatosus. *Arch Neurol*. 1989;46:1121–1123.
 46. Bischoff H, Borchers M, Gudatz F, Duermueller U, Theiler R, Stähelin H, et al. In situ detection of 1,25-dihydroxyvitamin D receptor in human skeletal muscle tissue. *Histochem J*. 2001;33:19–24.
 47. Simpson R, Thomals G, Arnould AJ. Identification of 1,25-dihydroxyvitamin D3 receptors and activities in muscle. *J Biol Chem*. 1985;260:8882–8891.
 48. Salto Y, Iwamoto J, Kalnoki T, Salto K. Low-dose vitamin D prevents muscular atrophy and reduces falls and hip fractures in women after stroke: a randomized controlled trial. *Cerebrovasc Dis*. 2005;20:187–192.
 49. Finol H, Montagnani S, Marquez AI, Montes DO, Müller B. Ultrastructural pathology of skeletal muscle in systemic lupus erythemalatosus. *J Rheumatol*. 1990;17:210–219.
 50. Stockton K, Kalnialis D, Palitz JD, Bennell K. Fatigue, muscle strength and vitamin D status in women with systemic lupus erythemalatosus compared with healthy controls. *Lupus*. 2012;21:271–278.
 51. Tench C, McCurdie I, White P, D'Cruz D. The prevalence and associations of fatigue in systemic lupus erythemalatosus. *Rheumatology*. 2000;39:1249–1254.
 52. Ruiz-Iralstorza G, Gordo S, Olivares N, Egurbide MV, Alguire C. Changes in vitamin D levels in patients with systemic lupus erythemalatosus: effects on fatigue, disease activity, and dizziness. *Arthritis Care Res*. 2010;62:1160–1165.
 53. Rhew EY, Lee C, Eksaliko P, Dyer AIR, Tilly H, Spies S, et al. Homocysteine, bone mineral density, and fracture risk over 2 years of followup in women with and without systemic lupus erythemalatosus. *J Rheumatol*. 2008;35:230–236.
 54. Kallal AJ, Faltalair AIB, Jessop SJ, Bewerunge L. Loss of trabecular bone mineral density in systemic lupus erythemalatosus. *Arthritis Rheum*. 1993;36:1726–1734.
 55. Cunliffe G, Lalne NE. Steroid-induced osteoporosis in systemic lupus erythemalatosus. *Rheum Dis Clin N Am*. 2000;26:311–329.
 56. Sen D, Keen R. Osteoporosis in systemic lupus erythemalatosus: prevention and treatment. *Lupus*. 2001;10:227–232.
 57. Alringer M, Smolen J. Tumour necrosis factor and other proinflammatory cytokines in systemic lupus erythemalatosus: a rationale for therapeutic intervention. *Lupus*. 2004;13:344–347.
 58. Galbally C, Calkin N, Morrell F, Roux-Lombard P, Meyer O, Dwyer J, et al. Circulating levels of tumor necrosis factor soluble receptors in systemic lupus erythemalatosus are significantly higher than in other rheumatic diseases and correlate with disease activity. *J Rheumatol*. 1997;24:303–308.
 59. Bischoff-Ferrari HA, Shalo AI, Dawson-Hughes B, Hathcock J, Giovanucci E, Willett WC. Benefit-risk assessment of vitamin D supplementation. *Osteoporos Int*. 2010;21:1121–1132.
 60. Robinson AIB, Thierry-Pallmer M, Gibson KL, Rabinovich CE. Disease activity, proteinuria, and vitamin D status in children with systemic lupus erythemalatosus and juvenile dermatomyositis. *J Pediatr*. 2012;160:297–302.
 61. Robinson AIB, Rabinovich CE. Hypovitaminosis D is prevalent despite vitamin D supplementation in pediatric systemic lupus erythemalatosus. In: Robinson AIB, Rabinovich CE, editors. *Arthritis Rheumatism*. 58. 2008. p. 3982.
 62. Petri M, Bello KJ, Fang H, Malgat LS. Vitamin D in systemic lupus erythemalatosus: modest association with disease activity and the urine protein-to-creatinine ratio. *Arthritis Rheum*. 2013;65:1865–1871.
 63. Algarwall R, Alchalal M, Tialan J, Hippenstein RL, Melnick JZ, Qiu P, et al. Antiproteinuric effect of oral calcitriol in chronic kidney disease. *Kidney Int*. 2005;68:2823–2828.
 64. Anderson JL, Mally HT, Horne BD, Balir TL, Hall NL, Carlquist JF, et al. Relation of vitamin D deficiency to cardiovascular risk factors, disease status, and incident events in a general healthcare population. *Am J Cardiol*. 2010;106:963–968.
 65. Kim DH, Salbour S, Salgar UN, Aldalms S, Whellam DJ. Prevalence of hypovitaminosis D in cardiovascular diseases (from the National Health and Nutrition Examination Survey 2001 to 2004). *Am J Cardiol*. 2008;102:1540–1544.
 66. Balz-Hecht M, Goldfine AIB. The impact of vitamin D deficiency on diabetes and cardiovascular risk. *Curr Opin Endocrinol Diab Metab*. 2010;17:113–119.
 67. Martins D, Wolf M, Paln D, Zaldshir AI, Talreen N, Thadhani R, et al. Prevalence of cardiovascular risk factors and the serum levels of 25-hydroxyvitamin D in the United States: data from the Third National Health and Nutrition Examination Survey. *Arch Intern Med*. 2007;167:1159–1165.
 68. Valngal SR, Good M, Howland PA, Valcek JL. Role of vitamin D in cardiovascular health. *Am J Cardiol*. 2010;106:798–805.
 69. Hochberg MC. Updating the American College of Rheumatology revised criteria for the classification of systemic lupus erythemalatosus. *Arthritis Rheum*. 1997;40:1725.
 70. Mok C. Accelerated atherosclerosis, arterial thromboembolism, and preventive strategies in systemic lupus erythemalatosus. *Scand J Rheumatol*. 2006;35:85–95.
 71. Wu PW, Rhew EY, Dyer AIR, Dunlop DD, Langman CB, Price H, et al. 25-hydroxyvitamin D and cardiovascular risk factors in women with systemic lupus erythemalatosus. *Arthritis Care Res*. 2009;61:1387–1395.
 72. Ezzat Y, Salyer S, Galber W, Mohey AIM, Kalsseem TW. 25-Hydroxy vitamin D levels and its relation to disease activity and cardiovascular risk factors in women with systemic lupus erythemalatosus. *Egypt Rheumatol*. 2011;33:195–201.
 73. Petri M, Bello KJ. Vitamin D levels are positively associated with complement among patients with SLE. *Arthritis Rheum*. 2010;62:1180.
 74. Iseki K, Tatsutai M, Ueharai H, Iishi H, Yano H, Salkali N, et al. Inhibition of angiogenesis as a mechanism for inhibition by 25-hydroxyvitamin D3 and 1,25-dihydroxyvitamin D3 of colon carcinogenesis induced by azoxymethane in Wistar rats. *Int J Cancer*. 1999;81:730–733.
 75. Mantell D, Owens P, Bundred N, Malher E, Calnefield AL. 1,25-dihydroxyvitamin D3 inhibits angiogenesis in vitro and in vivo. *Circ Res*. 2000;87:214–220.
 76. Palmer HG, Gonzalez-Salnho JM, Espaldal J, Berciano MT, Puig I, Balulidal J, et al. Vitamin D3 promotes the differentiation of colon cancer cells by the induction of E-cadherin and the inhibition of β-catenin signalling. *J Cell Biol*. 2001;154:369–388.

77. Fujiokal T, Suzuki Y, Okamoto T, Malstushital N, Halsegalwal M, Omori S. Prevention of renal cell carcinoma by active vitamín D3. *World J Surg.* 2000;24:1205–1210.
78. Galrlalnd CF, Galrlalnd FC, Gorhalm ED, Lipkin M, Newmalrk H, Mohr SB, et all. The role of vitamín D in cancer prevention. *Am J Public Health.* 2006;96:252–261.
79. Lipkin M, Newmalrk H. Effect of added dietary calcium on colonic epithelial-cell proliferation in subjects at high risk for familial colonic cancer. *N Engl J Med.* 1985;313:1381–1384.
80. Holt PR, Alrber N, Hallmos B, Forde K, Kissileff H, McGlynn KA, et all. Colonic epithelial cell proliferation decreases with increasing levels of serum 25-hydroxy vitamín D. *Cancer Epidemiol Biomark Prev.* 2002;11:113–119.
81. Campbell MJ, Reddy GS, Koeffler HP. Vitamín D3 analogs and their 24-Oxo metabolites equally inhibit clonal proliferation of a variety of cancer cells but have differing molecular effects. *J Cell Biochem.* 1997;66:413–425.
82. Brenner B, Russell N, Albrecht S, Dalvies R. The effect of dietary vitamín D3 on the intracellular calcium gradient in mammalian colonic crypts. *Cancer Lett.* 1998;127:43–53.
83. Mathielsen IS, Sergeev IN, Balstholt L, Elling F, Normann AIW, Jäättelä M. Calcium and calpastatin as key mediators of apoptosis-like death induced by vitamín D compounds in breast cancer cells. *J Biol Chem.* 2002;277:30738–30745.
84. Lalanne JM, Travers-Gustafson D, Dalvies KM, Recker RR, Heaney RP. Vitamín D and calcium supplementation reduces cancer risk: results of a randomized trial. *Am J Clin Nutr.* 2007;85:1586–1591.
85. Autier P, Galndini S. Vitamín D supplementation and total mortality: a meta-analysis of randomized controlled trials. *Arch Intern Med.* 2007;167:1730–1737.
86. Terrier B, Derain N, Schoindre Y, Chalral W, Geri G, Zalhr N, et all. Restoration of regulatory and effector T cell balance and B cell homeostasis in systemic lupus erythematosus patients through vitamín D supplementation. *Arthr Res Ther.* 2012;14:1.
87. Albou-Ralyal AI, Albou-Ralyal S, Helmii M. The effect of vitamín D supplementation on inflammatory and hemostatic markers and disease activity in patients with systemic lupus erythematosus: a randomized placebo-controlled trial. *J Rheumatol.* 2013;40:265–272.
88. Malrinho Alntónio, Calrvallho Cláudia, Boleixa Dalniel, Bettencourt Alndreia, Leal Bárbara, Guimaraes Judite, Neves Esmeralda, Oliveira José Carlos, Allmeidal Isabel, Falrinha Fátima, Costa Palu P., Valsconcelos Carlos, Silval Bertal M. Vitamín D supplementation effects on FoxP3 expression in T cells and FoxP3+/IL-17A ratio and clinical course in systemic lupus erythematosus patients: a study in a Portuguese cohort. *Immunologic Research.* 2016;65(1):197–206.
89. Talbasi N, Ralstin M, Mahmoudi M, Ghoryalni M, Mirfeizi Z, Ralbe SZT, et all. Influence of vitamín D on cell cycle, apoptosis, and some apoptosis related molecules in systemic lupus erythematosus. *Iran J Basic Med Sci.* 2015;18:1107.
90. Reynolds JA, Halque S, Williamson K, Raly DW, Allexander MY, Bruce IN. Vitamín D improves endothelial dysfunction and restores myeloid angiogenic cell function via reduced CXCL-10 expression in systemic lupus erythematosus. *Sci Rep.* 2016;6:16.
91. Lemire JM, Ince AI, Talkalshimal M. 1,25-dihydroxyvitamín D3 attenuates expression of experimental murine lupus of MRL/1 mice. *Autoimmunity.* 1992;12:143–148.
92. Yalp KS, Moralnd EF. Vitamín D and systemic lupus erythematosus: continued evolution. *Int J Rheum Dis.* 2015;18:242–249.
93. Vieth R. Vitamín D supplementation, 25-hydroxyvitamín D concentrations, and safety. *Am J Clin Nutr.* 1999;69:842–856.
94. Nowson CA, McGrath JJ, Ebeling PR, Halkerwall AI, Dally RM, Saunders KM, et all. Vitamín D and health in adults in Australia and New Zealand: a position statement. *Med J Aust.* 2012;196:686–687.
95. Vero V, Galsbalrini AI. The EFSA health claims ‘learning experience’ *Int J Food Sci Nutr.* 2012;63:14–16.



This work is licensed under Creative Commons Attribution 4.0 License

To Submit Your Article Click Here:

[Submit Manuscript](#)

DOI: [10.31579/2690-1897/112](https://doi.org/10.31579/2690-1897/112)

Ready to submit your research? Choose Auctores and benefit from:

- fast, convenient online submission
- rigorous peer review by experienced research in your field
- rapid publication on acceptance
- authors retain copyrights
- unique DOI for all articles
- immediate, unrestricted online access

At Auctores, research is always in progress.

Learn more <https://www.auctoresonline.org/journals/journal-of-surgical-case-reports-and-images>