



RESEARCH

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Association of vitamin D levels with anthropometric and adiposity indicators across all age groups: a systematic review of epidemiologic studies

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Abstract

Objectives: It has not been established whether vitamin D deficiency is associated with anthropometric state; therefore, this systematic review examined the relationship between serum vitamin D levels with anthropometrics and adiposity across different ages.

Methods: Studies that examined vitamin D deficiency with adiposity measures in different age groups were searched in the PubMed, Scopus, Embase, and Google Scholar databases until November 2023. Two investigators independently reviewed titles and abstracts, examined full-text articles, extracted data, and rated the quality in accordance with the Newcastle–Ottawa criteria.

Results: Seventy-two studies, with a total of 59,430 subjects, were included. Of these studies, 27 cross-sectional studies and one longitudinal study (with 25,615 participants) evaluated the possible link between 25(OH)D serum concentrations and anthropometric/adiposity indices in the pediatric population. Forty-two cross-sectional studies and two cohort investigations (with 33,815 participants) investigated the relationship between serum 25(OH)D levels and adiposity measures in adults and/or the elderly population. There is evidence supporting links between vitamin D deficiency and obesity, and revealed an inverse association between vitamin D and adiposity indicators, specifically in female subjects. However, the effects of several confounding factors should also be considered.

Conclusion: Most published studies, most of which were cross-sectional, reported a negative association between vitamin D and female adiposity indicators. Therefore, serum vitamin D levels should be monitored in overweight/obese individuals.

Keywords: vitamin D; 25(OH)D; anthropometric; adiposity

Introduction

Obesity results from excess fat accumulation and a positive energy balance, contributing to various chronic diseases and reduced life expectancy (1). Approximately 650 million adults, roughly 13% of the global adult population, were overweight or obese, with 340 million children and adolescents aged 5–19 years classified as overweight or obese in 2016 (2).

Micronutrient deficiencies, notably hypovitaminosis D, are common in obese patients (3, 4). Extensive observational studies like NHANES Framingham have linked obesity to an increased risk of hypovitaminosis D (5, 6). Vitamin D serves various functions, including maintaining calcium homeostasis and bone health, while also influencing metabolic processes, immunity, cellular proliferation, differentiation, among other effects such as antiinflammatory, antiatherogenic, cardioprotective, and neuroprotective impacts (7, 8). A global prevalence of widespread vitamin D deficiency has been identified, with deficiency rates rising by 13%, insufficiency rates reaching 40%, and notably higher rates observed in Asian countries (9, 10).

Obesity is commonly linked with reduced vitamin D levels regardless of various factors such as age, gender, season, study region, or smoking status (11). The coexistence of obesity and hypovitaminosis D represents a dual public health concern globally, prompting the need for investigating the underlying pathophysiology of this relationship. Mechanisms contributing to low vitamin D levels in obesity involve volumetric dilution, sequestration into adipose tissue, limited sunlight exposure, and reduced vitamin D synthesis in adipose tissue and the liver (11). Studies have suggested that low vitamin D levels may influence adipose tissue differentiation and growth, impacting obesity through gene expression regulation or by modulating parathyroid hormone (PTH), calcium, and leptin (11, 12, 13). While several observational studies have explored the link between vitamin D status and body weight, comprehensive evaluations of the relationship between serum vitamin D levels and anthropometric and adiposity indicators in both adults and children are lacking.

Hence, in this groundbreaking systematic review, we significantly contribute to the existing literature by taking a comprehensive and inclusive approach to evaluate the intricate relationship between serum 25(OH)D levels and adiposity. Unlike prior studies that primarily focused on specific age groups or relied on limited adiposity measures such as body mass index (BMI) and waist circumference (WC), our research spans diverse age groups and considers a broader set of indicators, including BMI, WC, HC (hip circumference), WHR (waist-to-hip ratio), and body fat mass percentage. Our findings reveal intriguing patterns

across the life span, adding a valuable dimension to the understanding of vitamin D deficiency in the context of obesity. This holistic evaluation provides a nuanced perspective on the association between serum 25(OH)D levels and various aspects of adiposity, offering a more comprehensive overview compared to previous reviews. This research is important as it seeks to fill the existing gap in knowledge concerning the intricate association between vitamin D levels and obesity. By conducting a systematic review, the study intends to shed light on the underlying mechanisms and implications of this relationship, thereby contributing to the development of effective public health strategies.

Methods

We conducted a systematic review of studies that assessed the relationship between serum vitamin D levels with anthropometric and adiposity indices in children, adolescents, adults, and the elderly. Serum 25(OH)D was used as a proxy measure for vitamin D levels.

Search strategy

The PubMed, Scopus, Embase, and Google Scholar databases were used to identify relevant publications. Two authors (BA and SA) independently searched papers published until November 2023 using ('25-hvdroxy vitamin D' OR 'vitamin D' 'cholecalciferol' OR '25(OH)D') AND ('BMI' OR 'body mass index' OR 'weight' OR 'obese' OR 'obesity' OR 'waist' OR 'waist circumference' OR 'adiposity' OR 'adipose', OR 'fat') as keywords. No restrictions were imposed on publication time or language. The reference lists of relevant articles were also reviewed by the authors to determine whether any publications were missing. All of the studies included in this systematic review were published in English. Data extraction was done independently by two investigators (BA and MN). In the event of any disagreement, three authors (BA, MN, and FH) discussed it among them to resolve the disagreement. Owing to the differences in the comparisons of the included studies (differences in exposures, outcomes, participants, and settings), diversity of applied statistical tools in the comparisons of the included studies, and lack of data that could be pooled, we performed a qualitative systematic review. The systematic review was conducted following the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA 2020) Statement (14).

Table 1 shows the PICOS (population, intervention/exposure, comparator, outcome, and setting) items used to conduct the systematic review. Owing to the methodological approach, no ethical approval was required.

Table 1 PICOS (population, intervention/exposure, comparator, outcome, and setting) criteria used to perform the systematic review.

PICOS	Criteria	
Population	Healthy general population across all age groups	
Intervention/exposure	Serum 25(OH)D	
Comparator	Statistical tools (OR, HR, RR)	
Outcome	Anthropometric/adiposity indices	
Setting	Observational studies	

OR, odds ratios: HR, hazard ratios: RR, relative risk.

Eligibility criteria

Publications with abstracts that suggested vitamin D levels were investigated in relation to anthropometric and adiposity variables were reviewed in full. Studies met the inclusion criteria if they: i) had observational design; (ii) were carried out in apparently healthy individuals (without chronic diseases, such as diabetes, liver diseases, cancer, or chronic kidney disease); and (iii) used serum 25(OH)D levels as a proxy for vitamin D state. However, clinical trials, reviews, editorials, and studies on nonhuman models, were excluded. Sex and age ranges were not strictly defined in this systematic review.

Study selection

Each title and abstract collected during the initial search was independently evaluated by two authors after removing duplicates. To ensure that eligibility and exclusion criteria were met, the two authors assessed full-text articles. The researchers consulted each other whenever they disagreed.

Data extraction and quality assessment

The following information was recorded in a data mining sheet: first author, publication year, country, ethnicity, design of the study, sample size, sex of participants, age, study population, method of 25(OH)D measurements, cut offs for vitamin D status, anthropometric indices investigated in the study and their cut off points, adjustments, and main findings. We assessed the quality of observational studies using the Newcastle–Ottawa Scale (NOS) (15).

Results

Literature search and study selection process

From databases, 19392 studies were initially found. Following the removal of 9247 duplicate articles, 7342 were excluded after scanning the titles/abstracts as they had no relevance to the present systematic review. After careful screening of 2803 full texts, we also

excluded 2731 more studies because they evaluated the relationship between serum levels of 25(OH)D with an outcome other than anthropometric/adiposity indices, were clinical trials, animal or *in vitro* studies in design, editorial, and reviews, or the participants of the studies were unhealthy (with chronic diseases, such as diabetes, cancer, or chronic kidney disease). Ultimately, 72 studies (12, 13, 16, 17, 18, 19, 20, 21, 22, 23, 24, 25, 26, 27, 28, 29, 30, 31, 32, 33, 34, 35, 36, 37, 38, 39, 40, 41, 42, 43, 44, 45, 46, 47, 48, 49, 50, 51, 52, 53, 54, 55, 56, 57, 58, 59, 60, 61, 62, 63, 64, 65, 66, 67, 68, 69, 70, 71, 72, 73, 74, 75, 76, 77, 78, 79, 80, 81, 82, 83, 84, 85) with 59,430 participants in total, published between 1981 and 2023, could be considered for the systematic review. Figure 1 illustrates the flowchart for selecting studies.

Characteristics of the studies conducted in children/adolescent population

The current systematic review identified 27 crosssectional studies (16, 17, 18, 19, 20, 21, 22, 23, 24, 25, 26, 27, 28, 29, 30, 31, 32, 33, 34, 35, 36, 37, 38, 39, 40, 41, 42) and one longitudinal study (43) that assessed the potential relationship between serum 25(OH)D anthropometric/adiposity and indices children. The studies about the relation between 25(OH)D serum level and anthropometric indices in pediatric were carried out in the following countries: four studies from Turkey, three from Spain, two from Denmark, five from the USA, two from Brazil, four from Italy and eight others from Iran, Korea, China, Thailand, England, Saudi Arabia, Sri Lanka, and Germany. The included studies were conducted between 2008 and 2021. Tgncluded studies had samples ranging from 51 to 12292 in this age group. Participant ages ranged from 3 months to 21 years.

All included studies involved both sexes, except three investigations (26, 39, 40) that were conducted in females (39, 40) and males (26) only. The included studies mostly considered the following potential confounders: sex, age, weight, BMI, cohort characteristics, Tanner stage, fat mass index, body fat mass percentage, BMI Z-score, parental education, pubertal breast stage, physical activity, dietary or supplemental vitamin D and calcium intake, race, ethnicity, season of blood collection, and height measures.

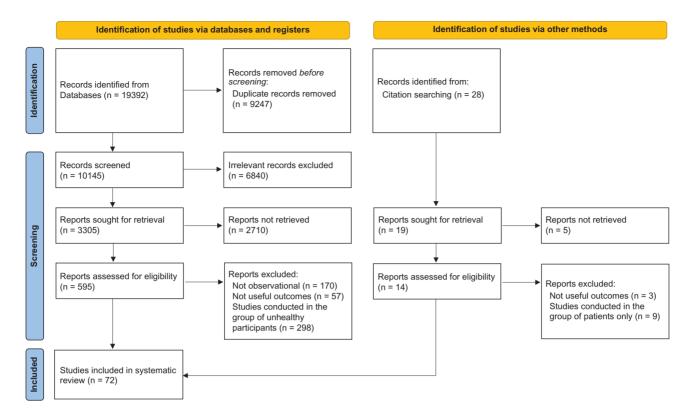


Figure 1PRISMA flow diagram for the selection process of the studies.

The characteristics of the articles are shown in Table 2.

Characteristics of the studies conducted in adult and/or elderly population

Our systematic search identified 42 cross-sectional studies (12, 13, 44, 45, 46, 47, 48, 49, 50, 51, 52, 53, 54, 55, 56, 57, 58, 59, 61, 62, 63, 64, 65, 66, 67, 68, 69, 70, 71, 72, 73, 74, 75, 76, 77, 79, 83, 84, 85) and two cohort investigations (60, 78) (with 33,815 participants) that investigated the potential relationship between 25(OH)D serum concentrations and anthropometric/adiposity indices in adults and/or elderly population. The studies about the link between 25(OH)D serum level and anthropometric indices in adults and/or elderly population were carried out in the following countries: 9 studies from USA, 3 from UK, 3 from Spain, 2 from Turkey, 4 from Saudi Arabia, 2 from Iran, 2 from India, 2 from Italy, 2 from Germany, and 18 others from China, UAE, Pakistan, Australia, Finland, England, Portugal, Malaysia, Bulgaria, Austria, Canada, Denmark, Greece, New Zealand, and the Netherlands. The included studies were conducted between 1981 and 2023. The included studies had samples ranging from 26 to 3113 in this age group. The age of participants was >18 years old. Among the included studies, 27 articles (44, 45, 46, 47, 48, 49, 50, 51, 52, 53, 54, 55, 56, 57, 58, 59, 60, 61, 62, 63, 64, 65, 66, 67, 68, 84, 85) involved both sexes, and 17

investigations conducted in females (12, 13, 72, 73, 74, 75, 76, 77, 78, 79, 80, 81, 82, 83) and males (69, 70, 71) only. The included studies mostly considered the following potential confounders: age, sex, season of blood sampling, smoking, vitamin D status, body fat mess, weight, height, BMI, IGF-1, PTH, UVB, alcohol, tobacco, sports, use of multivitamin supplements, menopausal status/HRT use, physical activity, socioeconomic status, income, job, obesity, education level, lifestyle, sun exposure, lean mass, nutrient intake, residence type, race, ethnicity, serum calcium, magnesium and phosphate, GFR, 25(OH)D, CRP, WC, month of blood collection, marital status, district, area, latitude, phosphorus, serum alanine aminotransferase, creatinine, and IL-6.

The characteristics of the articles are shown in Table 3.

Serum 25(OH)D and anthropometric measurements

Different methods were used to assess serum vitamin concentrations, including chemiluminescent immunoassav (CLIA), electrochemiluminescence immunoassay (ECLIA), radioimmunoassay (RIA), enzyme-linked immune sorbent assay (ELISA), immunoassay (IA), enzyme IA (EIA), protein-binding assay (PBA), chromatography, although three articles did not report the using method.

 Table 2
 Characteristics of the studies investigating the association between vitamin D and/or PTH with anthropometric indices in children/adolescents.

Study quality (NOS)	4	ιΩ	7	4	ιn	rv.	~	4	4	4
Main findings	Obese children had higher level of PTH. Bloavallable and free witamin D were lower in the obese group. There was no difference in terms of total vitamin D between groups.	Suboptimal vitamin D status and high levels of PTH are a common feature in pediatric with severe obesity.	Obese children had lower SDS for 25(OH)D and higher SDS for PTH than the control cohort.	The prevalence of vitamin D deficiency was higher in obese children compared to normal-weight and overweight children.	Subjects with vitamin D deficiency had deficiency had significantly elevated significantly elevated values for BMIZ-score, Wc, waste Z-score, body far percentage, far mess index, and DH than those with normal vitamin D status.	Vitamin D deflicency was significantly high in children with obesity and showed negative correlations with indicators of adposity.	Vitamin D deficiency was common among children and adolescents with obesity was closely was closely was closely was closely was closely with lower serum with lower serum 25(OH)D concentrations.	25(OH)D was lower in overweight and obese adolescents.	hypovitaminosis D and virtum D-deficient groups had higher BMI, far mass, and iPTH, than virtum in 2-sufficient syrup. Far mass was negatively correlated with 25(OH)D (=-0.40, with 25(OH)D (=-0.40, with 25(OH)D), it was positively correlated with IPTH (=0.46, P < 0.0001).	In obese children PTH leet was significantly higher and 25(OHI) was lower than controls. 25(OHI) was negatively correlated with PTH and BMI, PTH was not positively correlated with BMI ppercentile.
Adjustments		1	1	1	1	ı			1	
Anthropometric indices investigated in the study and their cutoff points	Obesity, BMI >95th percentile. Normal: 15th-85th percentiles.	Severe obesity: BMI Z-score >3.0, 99th percentile). Control: Z-score between -1.0 (15th percentile) and +1.0 (85th percentile).	Weight and height.	Normal weight at BMI s85th percentile, overweight at 85th-95th percentile, and obese at ≥95th percentile.	BMI, body composition, WC, WHR.	Above +2 SDS for WC of relevant references and percentage fat mass >28.6 in boys and >33.7% in girls were considered as high.	Obese: BML-5D5 5.2.33 or above the 99th percentile, coveragint BML-5D5 2.1.28 < 2.33), lear: BMI-5D5 2.1.28 or below the 90th percentile.	Weight, WC, hip cricumference were measured. Weight status was determined according to BMT-for-age growth charts. Normal weight: between the 3rd and 58th percentiles; overweight: the 88th and 97th percentiles; obses- above 97th percentiles; obses-	BMI >95th percentile for age, body composition.	Underweight, normal weight and overweight green effined as BMI below the 10th between 10th and 85th and above the 85th percentile, respectively, according to WHO standards.
Cutoffs for vitamin D status	Normal: 25(OH)D > 20 ng/mL and PTH <65 pg/ mL.	Normal: 25(OH)D 230 ng/mL, Insufficiency: 20-29 ng/mL, deficiency: <20 ng/mL Hyperparathyroldism: PTH levels >65 pg/mL.	Electrochemiluminescence assays	Severely Jacking: 25/OHJD <5 ng/mL, <15 ng/ mL as lacking, 15-20 ng/mL as deficiency.	Normal: 25(OH)D 230 ng/mL, Insufficiency: 20-29 ng/mL, deficiency: < 20 ng/mL.	Deficiency: 25(OH)D <20 ng/mL, Insufficiency: 25(OH)D: 20 -29 ng/mL, PTH >65 pg/mL.	Normal: 25(OH)D 550 mmol/L), insufficiency: <50 nmol 2: 30 nmol/L, Deficiency<30 nmol/L.	Deficiency; 25(OHJD < 20 ng/ml. Insufficiency; 25(OHJD: 20 -29 ng/ml	Normal: 25(OH)D 2.75 nMy, insufficiency: 50-74.9 nM, Deficiency<50 nmol/L.	
Method of 25(OHD measurement	25(OH)D: Liquid chromatography-tandem mass spectrometry with a commercial kit. TTH: colorimetric method using ready-made kits	Chemiluminescence immunoassay/two-site chemiluminescent enzyme-labeled immunometric assay in an immulite analyzer	Electrochemiluminescence assays	25(OH)D was measured using commercial kit	25(OH)D was measured by a high-specific themliumisessayland PTH assessed by a highly specific solid-phase, two-site dremiluminescent enzyme-labeled immunometric assay	Immunoassays	Electrochemiuminescence binding assay/	High-performance liquid chromatography	Nichols RJA/Nichols immunochemiluminametric assay	Chemiluminescence immunoussay technology
Study population	84 obese children and 78 healthy children	282 sever obese participants and 348 healthy control children	LIFE child study population	92 obese children and 58 healthy controls without chronic diseases	Adolescents with severe obesity	Children with BMI >2 SDS above the median for age and sex based on WHO	Children and adolescents with overweight/ doesity and 2143 population-based controls	Adolescents who were participated in the Health Survey-Sao Paulo study	Obese adolescents (BMI-95th percentile for age)	Healthy, overweight and obese children
Age of participants (years)	12-18	7.2–15.2	3 months-17 years	5-17	10.2–15.8	5-15	6-18	81-18	13.2-17.8	
Sex	<u>γ</u>	M.	M/A	E/M	M M	E/M	M M	Æ W	Σ Σ	M M
Sample size	162 (84/78)	630 (282/348)	2733 (317/2416)	150 (92/58)	236 adolescents with severe obesity (BMI 2-score > 3.0, 99th percentile)	202	3627 (1484/2143)	98	152	96 (60/36)
Study de sign	Cross sectional	Cross-sectional	Longitudinal	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional
Country, ethnicity	Turkey, NM	Spain; Caucasian	Germany; NM	Turkey; -	Spain; Caucasian	Sri Lanka; -	Denmark; North European White	Brazil; NM	USA; Caucasian, Hispanic, African American	Saudi Arabia; NM
First author (reference number), year	Küçükali (19), 2021	Durá-Travé (18), 2020	Geserick (43), 2020	Gün (17), 2020	Durá-Travé (20), 2020	Adikaram (21), 2019	Plesner (22), 2017	Gludid (23), 2017 Braził: NM	Alemzadeh (24), 2016	Saber (25), 2015

ontinued)

Table 2 Continued.

Study quality (NOS)	4	ហ	4	4	'n	v	ιn	4	
Main findings	tean and overweight groups had similar mean groups had similar mean 25(0Hp) levels and were below the sufficiently first provided the sufficiently freatured by 35(0Hp) insufficiency or deficiency was observed in both normal weight and overweight-obese	Serum ZS(OH)D was negatively associated with BM 2-scores and with BM 1-secondation with nonsignified and adjustment. But the negative association with regards association with adjustment. But the negative association with adjustment, adjustment, adjustment.	Children with hypothaminosis D had a higher mean BMI be percentile than the percentile than the vitamin D-sufficient group. PTH levels in the children with hypothaminosis D were significantly higher than in the children with normal levels of vitamin D.	The two groups were similar for BMI SDS and fat mass SDS, while showed differences for PTH.	Increased risk of hypotraminosis D in overweight-obese adolescents compared to subjects with normal BML, was shown. 25(OHD levels were inversely related to PTH and BMI-SDS.	The level of 23(OH)D was significantly lower in nowned weight group than in normal weight group. The FI Hevels were significantly higher in vitamin D deficient group compared to vidamin D insufficiency and sufficiency group.	Significant increased risk of hypovitaminosis D in overweight and obese subjects compared to individuals with normal BMI was observed. But weight status did not affect PTH levels.	Serum 25(OH)D levels were statistically lower in adolescents with weight excess, abdominal obesity, and higher levels of FTH.	
Adjustments	Weight, cohort characteristics, Tanner stage	FMI, BMI Z-score, parental education		1			1		
Anthropometric indices investigated in the study and their cutoff points	Normal weight, overweight, and seven bossity defined as and seven bossity defined as CDC-8M1-88° percentile and 285° percentile. percentile.	Height, weight, BMI, WC, body.composition.	BMI percentile groups (< 85th vs z 85th percentile).	Weight, height, body composition.	Weight status was categories in normal, overweight and obese according to criteria for subjects ye geers and according to the WHO for subjects aged 18-21 years.	Normal weight defined by BMI 3dt of 8dth percentile overweight was defined by BMI 38dt precentile for age and sex based on Korean standard growth curve.	Weight status was categorized in normal, overweight and obese according to order a for subjects <18 years and according to the WHO for subjects aged 18-21 years.	Weight, height, WC, %BF were measured Northonal stauss was assessed by BMI according the WHO recommendations. Excessive weight was defined as BMI above 1 SD. Addominal above 1 SD. Addominal above 1 SD. Addominal confessional according to the 90th certifies.	
Cutoffs for vitamin D status	Normal: 25(OHJD 230 ng/mL, Insuffidency: 20-29 ng/mL, deficiency: s 20 ng/mL.	Wamin D defidency and insufficiency defined as serum 25(OH)D concentration <25 and ≤50 mmol/L, respectively.	Sufficiency, 25(OHI) Tevel 275 mmol/L, Phyportaminosis D. 25(OHI) tevel 275, mmol/L, insufficiency, 25(OHI) Tevel 50-74, mmol/L, viramin D deficiency; 25(OHI) Tevel 50 mmol/L, viramin D deficiency; 25(OHI) Tevel 50 mmol/L.	25(OH)D <20 ng/mL (group II). 20 ng/mL (group II).	Wamin D defidency < 20; insuffidency 20-30 ng/mL; VH & 65.0 ng/mL; VH & 65.0 ng/L defined as hyperparathyroidism.	Deficiency; 25(OHJD, <20 ng/mL; insufficiency; 25(OHJD, 20-30 ng/mL, and sufficiency 25(OHJD, ≥30 ng/mL.	Wamin D defidency < 20; insuffidency 20-30 ng/mL, PTH 265.0 ng/mL, PTH 265.0 ng/L defined as hyperparathyroidism.	Wiamin D deficiency and insufficiency were s. 70 ng/m Land between 10-20 ng/m, respectively, the Insac PH low between 15 and 65 pg/m, being considered normal.	
Method of ZS(OH)D measurement	Competitive chemiluminescent immunoassay sandwich:type chemiluminescent immunoassay	Automated chemiuminescent immunoassay/ chemiuminescent immunoassay on ADVIA Centaur XP	Electrochemiuminescence immunoassay on Elecsys 2010 analyzers		Radioimnunoassay/immunoradiometric assay	Chemiumines cence immunoassay/ electrochemiumines cence immunoassay	Radioimmunoassay/immunoradionetric assay	RM kit/chemiuminescence immunoassay	
Study population	Boys were recruited in the Operative Unit of Medicine District of Salerno, section of Sapri (SA, Italy)	Date from the Optimal well-bush development and health for Danish children	Healthy children	Obese children with different values of vitamin D (25(OH)D <20 ng/mL (group I) and 25(OH)D >20 ng/mL (group II)	Healthy adolescents	Children who visited Bundang CHA Medical Center for theckup of their health and growth status	Healthy children and adolescents	Healthy adolescents	
Age of participants (years)	11-14		6-12	10.2 ± 2.8	10-21	4-15	Children (2.0–10.9 years) and adolescents (11.0–21.0 years)	15-17	
Sex	Σ	M/A	M.	E/M	M/A	M.	₽/₩	M/A	
Sample size	108	782	159	120	427	1212	652	160 (83/77)	
Study design	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	
Country, ethnicity	Italy; NM	Danish Danish	Reesukumal (28), Thailand: Thai Cross-sectional 2015	Italy; Caucasia, Africans, Asians, others	Italy; Italian	Chung (31), 2014 Korea; Korean Cross-sectional	Italy; Italian	Brazilian Brazilian	
First author (reference number), year	Di Nisio (26), 2015	2015	Reesukumal (28), 2015	Rusconi (29), 2015	Vierucci (30), 2014	Chung (31), 2014	Vierucci (32), 2013	Oliveira (33), 2013	

Table 2 Continued.

Study quality (NOS)	٢	4	4	4	ιν	ιŋ	4	4	4	4
Main findings	Vitamin D deficiency is highly prevalent in overweight and obese children.	Fat mass was negatively correlated with 25(OH)D, but was positively correlated with PTH.	No significant difference was found between the groups with different vitamin D levels in terms of weight SDS, height SDS, BMI SDS, and WC SDS.	Obese children had a significantly lower significantly lower 25(cH)D and a higher liPTH than nonobese children. Insufficient serum lewes of 25(CH)D were detected in 5% of normal children and in 30% of the obese children.	The level of 25(OH)D had a negative correlation with BML-SDS and height-SDS in females, but these correlations were not significant in males.	A significant positive correlation was found between lean body mass and plasma 25(0Hf)D. No significant correlation was found between the %BF and vitamin D status.	It was shown a trend toward a significant negative relationship between 25(OH)D and BMI, PTH. Level of PTH was significantly associated with BMI.	25(OH)D decreased by 0.46 ± 0.22 ng/mL per 1% increment in body fat mass, whereas PTH decreased by 0.78 ± 0.29 pg/mL per 1% increment in visceral adipose tissue.	Hypovtaminosis D and vidamin D-deficient. farmas, and IPH; than vitamin D-sufficient group; far mass was megatively correlated with SciOHD, it was postbively correlated with IPH.	It was shown a negative correlation between serum vitamin D level and BMI in obese and overweight subjects whose vitamin D level was below 20 ng/mL.
Adjustments	Race, age, sex, season, TV use, PA, vitamin D, and milk intake		1	Sex, age, and Tanner stage	1	Pubertal breast stage, physical activity, dietary calcium and vitamin D intake	ВМІ	Age, sex, race, season, vitamin D intake, height measures, Tanner stage	Age, sex, ethnicity, season	1
Anthropometric indices investigated in the study and their cutoff points	Height, weight, BML	Age-adjusted BMI ≥95th percentile.	Height, weight, BMI, WC. Obesity was defined as BMI ≥ 95th.	Weight, Neight, WC, and body composition were measured. Obese defined ad SDS-BMI = 2), extremely obese: SDS-BMI = -4.	Height, weight, BMI.	Height, weight, BMI, body composition.	Height, weight, BML Obesity was defined as BMI ≥95th.	Height, weight, BMI, body composition.	Height, weight, BMI, body composition, Obesty, was defined as BMI 3-95th percentle for age.	Height, weight, BMI.
Cutoffs for vitamin D status	Deficiency: -20 ng/mL, insufficiency: 20-30 ng/mL	1	Normai: 25(OHJD 230 ng/mL, insufficiency: 20-30, deficiency, < 20 ng/mL, PTH >65 pg/ mL was hyperparathyroidism.	Vitamin D sufficient > 20 ng/m) or insufficient < 20 ng/mL.	Severely deficient <12.5 mmol/L; deficient 21.5 and <25 mmol/L; insufficient 22.5 and <50 mmol/L sufficient, 250 mmol/L and s250 mmol/L and toxic >250 mmol/L.		Vitamin D deficiency cutoff; <20 ng/mL	Wamin D deficiency was defined as 25(OH)D concentration <20 ng/mL.	Hypovitaminosis D, 28(OHD <75 mmd/L; sufficiency, 25(OHD o 275 mmol/L; deficiency, 25(OHD loved <60 mmol/L; nsufficiency, 25(OHD of 50-749 mmol/L.	Deficiency 25(OH)D <10 ng/mL; insufficiency as leves of 25(OH)D between 10 and 20 ng/mL, and a normal vitamin D level as >20 ng/mL.
Method of 25(OHD) messurement	Radioimmunoassay	Nichols RIA/Nichols immunochemiluminometric assay	Electrochemiluminescence	Electrochemilumi nes cence im munoassays	Radioimmunoassay/immunoradiometric	Fradioimmunoassay/immunometric assay	Liquid chromatography-tandem mass spectrometry (LC-MS/MS/two-site immunoradiometric assay	In-house competitive protein binding assay/ enzyme immunoassay	Nichols radioimmunoassay.Nichols immunochemiluminometric assay	Competitive protein binding assay/ chemiluminescence with an immulite One analyzer
Study population	Children who were enrolled in the 2003–2006 National Health and Nutrition Examination Survey	Obese adolescents	Children and adolescents with BMI ≥95th percentile	Obese children	Healthy children and adolescents who came to a routine growth monitoring clinic of the university hospital	Apparently healthy adolescent girls	Obese adolescent girls	Healthy obese adoles cents	Obese children and adolescents	Secondary and high school children
Age of participants (years)	6-18	13.1–17.9	8-16	41-7	∞ 0 + 0 + 0 + 0 + 0 + 0 + 0 + 0 + 0 + 0 +	15.0 ± 0.4	14±2	13.0-17.9	6.0-17.9	11-19
Sex	M	Ā	A M	Ā	E. M	ш	ш	E M	M/4	M M
Sample size	12292	133	106	105 (66/39)	ε ε	323	13	88	127	301
Study de sign	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional
Country, ethnicity	USA; White, African American, Latino, others	USA; Caucasian, Hispanic, African American	Turkey, Turkish	Spain; Caucasian	Iran; NM	China; Chinese	England; African- American	USA; Black or African American, Hispanic, Summer, North of Atlanta	usa; Caucasian, Hispanic, African American	Turkey, Turkish
First author (reference number), year	Turer(37), 2013	Alemzadeh (35), 2012	Buyukinan (36), 2012	Codoñer-Franch (34), 2012	Razzaghy-Azar (38), 2010	Foo (39), 2009	Ashraf (40), 2009	Lenders (41), 2009	Alemzadeh (42), 2008	Çizmecioğlu (16), 2008

BMI, body mass index; F/M, female/male; FMI, fat mass index; iPTH, intact parathyroid hormone; NOS, Newcastle-Ottawa Scale; PA, physical activity; PTH, parathyroid hormone; SDS, standard deviation score; WC, waist circumference; WHO, World Health Organization.

 Table 3
 Characteristics of the studies investigating the association between vitamin D and/or PTH with anthropometric indices in adult and/or elderly population.

First author (reference number), year	Country, ethnicity	Study design	Sample size	ĕ	Age of participants (years)	sendy population	Method of 200H () measurement	Carofs for vizamin D status	Anthropometric indices investigated in the study and their cutoff points	Aljustments	Main findings	Saudy quality (NOS)
Tejada-Romero (12), 2022	Spain; NM	Cross-sectional	679 (493/186)	T.	60.6 ± 13.6	Healthy postmenopausal women	Immunochemical lumi nescence/immunochemical luminescence	1	Weight, height, BMI. Obesitywas defined as a BMI ≥30.	Age	Obese women had lower levels of 25(OH)D and higher PTH than nonobese.	9
Shan (13), 2022	China; Chinese	Cross-sectional	1505	ш	30.04 (23.98-37.81)	Women of childbearing age	Liquid chromatography-tandem mass spectroscopy/ electrochemiluminescence immunoassay		Height, weight, WC, BMI.	Season, district, area, latitude, age, BMI, PTH, P, ALT, CRE, IL-6, hs-CRP	Insufficient 25(OH)D was significantly related to the risk of elevated WC after adjusting for confounders.	7
Gariballa (48), 2022	UAE; NM	Cross-sectional	648	F/M	81 %	Community free living adults	Electrochemiluminescence	Deficiency:< 20 ng/ml.; insufficiency: 20-32 ng/ml; sufficient: > 32 ng/ml.	Height, weight, BMI.		There was no difference in BMI between groups, but, it was higher in watmin D deficient subjects aged 250 and females <50 years.	9
Saleem (72), 2021	Pakistan; NM	Cross sectional	397 (264/133)	ш	20-50	Healthy women with normal fasting glucose	Radioimmunoassay/spectrophotometer	Deficient: serum vitamin D<12 ng/ml.; sufficient: vitamin D>30 ng/ml.	Weight, height, BMI, WC, and HC. Obesity was defined as BMI 330 kg/m?.		PTH levels were negatively correlated, though nonsignitively with visini. D. The near BML WC, HC WHR, and PTH were not significantly different in vitamin Defeticient as compared to nondeficient deese women.	4
Djafari (49), 2021	Iran; NM	Cross-sectional	178	F/M	60-83	Elderfy individuals	Enzymatic method using commercial kits		Weight, height, BMI, WC, HC, and fat mass.	Age, sex, income, PA, job, smoking, vitamin D and calcium supplement use	28(OHID was higher in the highest category of BAI compared to the lowest one. An inverse association between BAI with 23(OHID was reported. No significant association between the WBE with PTH and 25(OHID was observed.	4
Menon (69), 2021	India; Indian	Cross sectional	224	Σ	20.74 ± 1.43	Healthy military training	Chemilumin escence an abzer	Sufficient: vitamin D.≥30 ng/ml; insufficient: 20–30 ng/ml; deficient: ≤20 ng/ml.	Height, weight, BMI, body composition.		There was no correlation of 25(OH) cholecalciferol with BMI or FMI.	4
Yaylali (73), 2021	Turkey; NM	Cross sectional	126 (73/53)	ш	17-55	Healthy premenopausal women	High performance liquid chromato graphy/ electrochemiluminescence immunoassay		Height, weight, WC, BMI, body composition. Obesity was defined as BMI>30 kg/m².		A positive association was observed between PTH and visceral fat. Vitamin D levels were inversely associated with visceral fat.	4
Sharma (74), 2020	Australia; NM	Cross-sectional	76 (50/26)	ш	61.9-68.5	Healthy normal and overweight postmenopausal women	Chemilumin escent immunoassays		Weight, height, BMI, WC, HC, and body composition. Normal:BMI <25 kg/m², and overweight; BMI >27 kg/m².		Women in the highest visceral adipose tissue quartile had significantly lower 25(OHID.	4
Albassam (75), 2019	Saudi Arabia; NM	Cross-sectional	265 (179/86)	ш	18-70	Middle-aged Saudi women	Electrochemiluminescence immunoassay/Milliplex MAP Human Bone Magnetic Bead Panel		Weight, height, BMI, WC, HC, NC, body composition.	Age and menopausal status	NC was inversely associated with 25/0HjD and PTH. In the nonobese, WHR was inversely associated with PTH.	so.
Saarnio (50), 2018	Finland; Caucasian	Cross-sectional	595	F/M	37-47	Healthy men and women	1D5 enzyme immun obszay kit/immunol umin escence- based method by immulite 1000		Weight, height BMI.	Age, physical activity, smoking	Inwomen, 25(0HD) keets dd not differ among the BMI normals. Jak (OHD) was weer in doese men than in normal-weight. Alogethet, obbes subjects had fowner and analyweight. Alogethet obbes subjects had fowner sightly than normal-weight. In women FTH was sightlicantly higher in obese compared to normal weight. Alogether, the difference was significant.	so.
Kord-Varkaneh (51), 2018	Iran; NM	Cross-sectional	178	F/M	60-83	Elderly subjects	Enzymatic method	1	Weight, height, BMI, WC.	Age, sex, PA, smoking, marital, supplement use	No significant correlation between 25(OH)D with BMI and W.C.	4
Raposo (52), 2017	Portugal; NM	Cross-sectional	200	F/M	53 (41-67)	Adults registered in primary health-care centers	Chemiluminescent immunoassay electrochemiluminescent immunoassay	Deficiency <12 ng/ml; inad equacy ≥12 and <20 ng/ml; sufficiency ≥20 ng/ml.	Weight, height, WC, HC. Participants were classified into: underweight (BMI - e18.5 kgm/n, normal ange (±8.5 to -2.2), pre-obese (±25 to <30 kg/m²) and obese (±30 kg/m²) ard obese (±30 kg/m²) categories.	Age and sex	The serum 25(OHD) levids were negatively associated with BMI. Positive associations between PTH with BMI and WC were found.	so.
Trevisan (76), 2017	Italy, Caucasian	Cross-sectional	218	L.	565	Healthy older women	Radioi mmun oassay/two-s tep im muno radio metric assay	25(OH)D level <50 nmol/L was considered inadequate; PTH value >60 pg/mL defined typerparatryroidsm	Weight, height, BMI, body composition.	Age, 25(OHJD, PTH, month of blood collection	Fat mass showed a significant negative association with 25(OH)D. Binary logistic analysis revealed a protective effect of adiposity on secondary hyperparathyroidism.	4
Walsh (53), 2016	UK, Caucasian	Cross-sectional	233	F/M	25-40 and 55-75	Healthy men and women in different weight groups	Immunoassay/direct measurement by immunoassay	NM	Weight, height, BMI.	Age, sex	Serum 25(OH)D was inversely correlated with BMI. PTH did not differ by BMI group.	4
Shafinaz (54), 2016	Malaysia; Malays, Chinese, Indians, others	Cross-sectional	858	F/M	<30 to >50	Permanent teach ers who worked in the selected government secondary schools	Electro chemilumin is cence i mmuno assay	Serum ZS(OHID -220 ng/mL vas considered as vitamin D deficient	Weight, height, BMI, WC, body composition, Underweight: BMI <17.5 kgm²; normal weight: 17.5–22.9 kg m²; overweight: BMI 13.5–27.9 kgm²; obese: BMI ≥ 28.0 kg/m²;	Age, sex, ethnicity, sun avoidance score, SBP, BMI, WC, %8F	Asignificant negative association between serum 25(0H) Dwith BM I and MgF was observed. Higher BMI and larger WCwere significantly associated with lower serum 25(0H) D level.	φ
Al-Daghri (55), 2015	Saudi Arabia; NM	Cross-sectional	830	F/M	18-50	Apparently healthy individuals	ELISA	Sufficient vitamin D>50 nmoVL, insufficient: 25-50 nmoVL and deficient: <25 nmoVL	Weight, height, BMI, WC, HC, arm circumference.	Age, BMI, glucose, cholesterol, TG, LDL, HDL	Vitamin D insufficiency was significantly associated with abdominal obesity in males.	9
Wright (56), 2015	India; Caucasian, Black, Asian, Hispanic	Cross-sectional	336	F/M	35-65	Middle-aged overweight and obese healthy adults	Radioimmunoassays	ı	Weight, height, BMI, WC, HC, body composition.	Age, sex, race, season, supplement use, PTH	Total and central adiposity but not peripheral adiposity predicted fow plasma 25(OHJD.	4
George (57), 2015	USA; Caucaslan	Cross-sectional	714	F/M	18-65	Healthy adults	High-performance liquid drromatography (HPLC) kit/ chemiluminescent assay		Weight, height, BMI, WC, body composition.	Age, sex, height, ethnicity, serum calcium, magnesium and phosphate, GFR, smoking	25(OH)D was not associated significantly with BMI and WC.	9
Tos unbayraktar (58), 2015	Turkey, NM	Cross-sectional	8	F/M	18-63	Healthy individuals with various BMIs		Deficiency: 25(OH)D <20 ng/mL; and ≥20 ng/mL as sufficiency	Weight, height, BMI, WC, HC, body composition.		Overweight and obese groups had higher PTH and lower 22(OHD levels than normal weight group. The overweight group had higher 25(OHD and lower PTH than obese group.	4
Sorkin (77), 2014	USA; white and black	Cross-sectional	239	ш	46-78	Sedentary postmenopausal women without diabetes	RJA/immunoradiometric assay		1	Race, age, and time	25(OH)D was inversely related to visceral abdominal fat, %BF, and PTH.	4
											(Continued)	(pan

Table 3 Continued.

Table 3 Continued

Agreement (SON)	4	4	25	7
Main findings	PTH was higher and 25(OHID was lower in the obese than in the nonobese individuals.	The mean plasma 25(OHD) level was significantly lower in the obese group than in age-matched controls.	Compared with normal weight, obese individuals had lower 25(OH)D.	Low levels of 25(OH)D are linked to higher BMI.
y Adjustments	WN	N Z	Age, sex	WW
Anthropome tric indices investigated in the study and their cotoff points	Weight, height, BMI.	Weight, height, BMI.	Weight, height, BMI, body composition.	BMI, WC, HC, WHR, %BF.
Ontoff for vitamin D status	NM	NM	NM	Deficiency: 25(OH)D <20 ng/mL
Method of 200H (i) measurement	Competitive protein binding assay/ radioimmunoassay	Competitive protein-binding assay	Autoanalyzers	Adult population of LIFE Adult-Study Chemiluminescent enzyme immunometric assay
S midy population	Normal white subjects	Obese patients before intestinal bypass Competitive protein-binding assay rgastric partitioning versus healthy normal controls	Community-dwelling men and women Autoanalyzers from South Yorkshire	Adult population of LIFE-Adult-Study
Age of participants (years)	20-35	42 (22/20) F/M Obese: 22-49; nonobese: 19-52	25-40 or 55-75	40-79
S S		F/M	F/M	F/M
Sample size	26 (12/14) F/M	42 (22/20)	200	1032 (533/499)
Study design	Cross-sectional	Cross-sectional	Cross-sectional	Cross-sectional
Country, ethnicity	USA; white	England, NM	UK, NM	Germany, Caucasian Cross-sectional
First author ('ord revoce number), year Country, ethnicity	Bell(46), 1985	Compston (47), 1981	VII aca (84), 2022	Avila Castillo (85), 2023

glomerular filtration rate; hs-CRP, high-sensitivity C-reactive protein; IGF-1, insulin-like growth factor 1; iPTH, intact parathyroid hormone; NC, neck circumference; NM, not mentioned; NOS, female/male; FMI, fat mass index; HC, hip circumference; GFR, Newcastle-Ottawa Scale,P, phosphorus; PA, physical activity; PTH, Parathyroid hormone; SDS, standard deviation score; WC, waist circumference; WHR, waist-to-hip ratio. creatinine; F/M, adiposity index; BMI, body mass index; CRE, %BF, percentage of body fat; ALT, alanine aminotransferase; BAI, body

Anthropometric status was assessed based on weight, BMI, WC, WHR, and body composition.

The relationship between vitamin D with anthropometric and adiposity indicators in children/adolescents population

In a study by Durá-Travé (18), among 282 participants with severe obesity and 348 healthy control children group, vitamin D deficiency was more frequent (P < 0.05) in the obesity group (44.5 vs 11.5% and 22.4 vs 3.9%, respectively). In Geserick et al.'s study (43) obese children had significantly lower 25(OH)D SDS (standard deviation score) levels (-0.43) than the reference group. According to the Codoñer-Franch et al. (34) obese children had a significantly lower 25(OH)D level (P=0.002) compared with nonobese children. The obese group had significantly higher vitamin D deficiency rates compared to the control group in another study (P < 0.001) (17). Furthermore, a study (30) demonstrated an increased risk of hypovitaminosis D in overweight/ obese adolescents (OR 3.89) compared to those with a normal BMI. Serum 25(OH)D levels were inversely related to BMI-SDS (r = -0.141, P = 0.007).

According to a cross-sectional study by Saber *et al.* (25) obese children had significantly lower 25(OH)D levels (21.02 vs 29.45 ng/mL) than controls. There was a negative correlation between serum 25(OH)D with PTH, weight, and BMI (r=-0.45, -0.55, -0.47, P< 0.01, respectively).

A cross-sectional research by Lenders *et al.* on 58 healthy obese adolescents (41) found 25(OH)D decreased by 0.46 ± 0.22 ng/mL per 1% increment in body fat mass (P = 0.05).

The cross-sectional study by Alemzadeh *et al.* (42) among 127 subjects aged 13.0 \pm 3.0 years, found that hypovitaminosis D and vitamin D-deficient subjects had higher BMI, fat mass, and iPTH compared to vitamin D-sufficient subjects (P < 0.01). Also, fat mass was negatively correlated with 25(OH)D (r = -0.40, P < 0.0001) regardless of seasonal and racial/ethnic factors. In other works, by the same research group (24, 35) between obese adolescents, a negative correlation was found between fat mass and 25(OH)D (P < 0.001).

As reported by Chung *et al.* (31), the level of 25(OH)D in overweight children was significantly lower than that of normal weight children (17.1 \pm 5.1 ng/mL vs 19.1 \pm 6.1 ng/mL, P < 0.001). There was an independent association between overweight and vitamin D deficiency (OR 2.21; 95% CI 1.62–3.01).

According to Ashraf *et al.* (40), there is a trend toward a significant relationship between 25(OH)D with BMI (r=-0.24; P=0.087) and PTH (r=-0.24; P=0.088).

According to Buyukinan *et al.*'s study (36), children and adolescents with a BMI ≥95th percentile did not show any significant differences in terms of weight, height,

BMI, and waist circumference between the three groups with deficient, insufficient, and normal vitamin D levels.

There was also a cross-sectional study, by Durá-Travé *et al.* (20), showing that adolescents with severe obesity who had vitamin D deficiency had significantly (P < 0.05) elevated BMI *Z*-scores, waist *Z*-scores, body fat percentages, fat mass indexes, and PTH values than those with normal vitamin D state. The serum 25(OH)D levels were negatively correlated (P < 0.05) with body fat percentage, fat mass index, and PTH.

Lean and overweight boys (based on CDC-BMI percentiles) had similar mean 25(OH)D levels (P=0.160) and were below sufficiency thresholds according to an analysis by Di Nisio *et al.* (2015) (26). However, there was a large proportion of subjects with insufficient or deficient 25(OH)D in both groups (normal weight: 45/59, 76%; overweight/obese: 45/49, 92%, P=0.03).

In the study by Foo *et al.* (39), in adolescent girls, lean body mass and plasma 25(OH)D levels were significantly correlated (r=0.446; P=0.001). However, the correlation between body fat percentage and vitamin D status was not significant (r=0.104; P>0.05). According to Giudici *et al.* (23), compared to normal-weight participants, those with overweight had a lower 25(OH)D. All measures of BMI, weight, and WC were negatively associated with 25(OH)D (P<0.05). Other researchers (16) found that serum vitamin D levels were negatively correlated with BMI in overweight/obese subjects with vitamin D levels <20 ng/mL (r=-0.186, P<0.01).

Bioavailable and free vitamin D were lower in the obese group. However, the total vitamin D level between the two groups did not differ, according to Küçükali *et al.* (19). In another study by Turer *et al.*, vitamin D deficiency is highly prevalent in overweight and obese children. (37). In a study by Oliveira *et al.* (33) serum 25(OH)D levels were statistically lower in adolescents with excess weight, abdominal obesity, and a high level of PTH (P < 0.05).

Another cross-sectional study by Petersen *et al.* (27) showed a negative relationship between serum 25(OH)D and BMI *Z*-scores and fat mass index (P=0.001). However, the association with BMI *Z*-scores became nonsignificant when the model was adjusted for parental education (-0.03, 95% CI -0.07, -0.001, P=0.14).

Plesner *et al.* (22) found that 16.5% of obese children and adolescents showed vitamin D deficiency, with an OR 3.41 (CI 2.27–5.71; P < 0.0001) in comparison with peers with normal weight. An increase in risk of hypovitaminosis D was observed in overweight (OR 5.02) and obese (OR 5.36) subjects compared to normal weight subjects (32).

There was a difference in BMI percentile between children with hypovitaminosis D and children who have sufficient vitamin D (56.7 \pm 33.9 vs 42.6 \pm 36.0; P=0.04) according to a cross-sectional study (28).

According to a multivariate analysis, high BMI percentile and high PTH levels were the parameters related to 25(OH)D concentration <75 nmol/L.

Another evaluation (21) among children with obesity could not observe any significant associations between vitamin D deficiency and the anthropometric or metabolic derangements.

In the study by Rusconi *et al.* (29), 59 had 25(OH)D <20 ng/mL (group I) and 61 had 25(OH)D > 20 ng/mL (group II) were recruited. The two groups were similar for BMI SDS and fat mass SDS.

Razzaghy-azar *et al.* (38) reported that 25(OH)D level had a negative correlation with BMI-SDS and height-SDS in girls (P=0.01 and 0.039, respectively), but these correlations were not significant in boys.

The relationship between vitamin D with anthropometric and adiposity indicators in adult/elderly population

Albassam *et al.* (75) found a negative correlation (R=-0.17; P<0.05) between neck circumference, a surrogate for upper subcutaneous fat, and 25(OH)D in both obese and nonobese participants.

Gonzalez-Molero *et al.* (60) found that neither obesity at baseline (OR 0.98, 95% CI 0.69–1.40, P=0.93) nor obesity after undergoing a second evaluation (OR 0.80, 95% CI 0.48–1.33, P=0.39) was significantly related to vitamin D status. Yaylali *et al.* (73) demonstrated that vitamin D levels were inversely associated with visceral fat (P=0.002, r=-0.366). It has also been shown (12) that obese women have lower 25(OH)D values than nonobese women.

In another analysis by Jungert *et al.* (61), BMI, hip circumference, and body fat were negatively correlated with 25(OH)D, but not in men. Using multiple regression analyses, total body fat was shown to be a negative predictor of 25(OH)D concentrations in women even after adjusting for confounders (β =-0.247; P=0.016), whereas after adjusting for iPTH, there was no statistically significant association between BMI, hip circumference, and 25(OH)D. In men, 25(OH)D was not influenced by anthropometric or body composition variables.

The overweight and obese groups in Tosunbayraktar *et al.*'s study (58) had lower 25(OH)D levels when compared to the normal BMI group (P=0.01). Participants in the overweight group had higher levels of 25(OH)D than those in the obese group (P < 0.05).

A study by Ardawi *et al.* (80) showed that serum 25(OH)D was lower (P=0.001) in upper quintiles of BMI and WHR. Another cross-sectional study (70) found vitamin D deficiency was prevalent among older and obese men with no education and sedentary lifestyles. Among obese, euglycemic women, 90 (40.7%) were deficient in vitamin

D, according to a study (72). There were no significant differences in mean age, BMI, WC, hip circumference, WHR, and PTH between vitamin D-deficient and nondeficient obese women.

Based on Bolland *et al.*'s findings (83) 25(OH)D was negatively correlated with total fat mass, trunk fat, and pelvic fat. On multivariate analysis, 25(OH)D was negatively related to pelvic fat mass (P=0.014; partial r²=0.05). According to Compston *et al.* (1981) (47) obese individuals had significantly lower plasma 25(OH)D levels than age-matched controls.

In another study by Djafari *et al.* (49), among elderly persons, 25(OH)D (P=0.030) was higher in the highest category of body adiposity index (BAI) compared to the lowest one. Additionally, linear regression demonstrated a negative association between BAI with 25(OH)D ($\beta=-0.039$, P=0.029).

Healthier WC was also associated with better 25(OH)D concentrations among adults (63). Other studies found no significant correlation between 25(OH)D with WC (45, 51) and BMI (51). According to Macdonald *et al.* (82), women in the top quintile of BMI had lower 25(OH)D (P < 0.01). There was a significant correlation between 25(OH)D and BMI (r = 0.58; P = 0.01). In addition, BMI was highly predictive of 25(OH)D level (r = -0.52; P < 0.01).

According to another study (69), 25(OH)D is not correlated with BMI or fat mass index in healthy male adults. No significant association was found between anthropometric indices and serum 25(OH)D levels in Moschonis *et al.*'s study (2009) (81) on nonosteoporotic, postmenopausal women.

Among women with different BMI levels, Saarnio $et\ al.\ (50)$ found no difference in 25(OH)D levels. It was found that 25(OH)D was lower in obese man as compared to normal weight man (48.0 \pm 2.4 nmol/L vs 56.4 \pm 2.0 nmol/L, P=0.003), but there was no difference between normal weight and overweight group or overweight and obese group. When obesity was examined in both sexes, obese subjects had lower 25(OH)D than normal-weight subjects (50.7 \pm 1.6 vs 57.0 \pm 1.0 nmol/L, respectively; P=0.003). A significant difference was also found between overweight and obese groups (P=0.023). It was found by Walsh $et\ al.\ (53)$ that serum 25(OH)D was inversely related to BMI.

A study among male participants, by Frost *et al.* (71) showed that 25(OH)D levels were lower in those with a high BMI; those with a BMI over 25 kg/m² had a lower 25(OH)D (61.4 (27.8) nm vs 66.7 (27.5) nm, P=0.015). In participants with a BMI >25 kg/m², an increase in BMI of 1 kg/m² led to a decrease in 25(OH)D of 1.7 nm (95% CI: -2.8; -0.6, P=0.002), whereas in participants with a BMI <25 kg/m², BMI and 25(OH)D were unrelated (coef: 0.7 (95% CI: -0.6; -0.2)). Body fat mass was inversely related to PTH only in individuals with vitamin D insufficiency. Further, 25(OH)D were associated with lean body mass in adjusted analyses and in participants with low vitamin D levels.

An analysis by George *et al.* (57) showed 25(OH)D was not correlated with BMI (P=0.38) and WC (P=0.99). In the Sharma *et al.* study (74), women with a higher visceral adipose tissue quartile had significantly lower 25(OH)D levels (P=0.05).

A negative correlation was found between serum 25(OH)D with BMI (r=-0.4; P<0.0001) and body fat mass (r=-0.41; P<0.0001) among healthy adults (68).

Analysis of the subjects aged \geq 65 years, in the Longitudinal Aging Study Amsterdam (67), revealed that after adjusting for potential confounders, higher BMI, WC, and sum of skin folds were statistically significantly related to lower 25(OH)D (standardized β values were -0.136, -0.137, and -0.140, respectively; all P < 0.05) and with higher PTH (0.166, 0.113, and 0.114, respectively; all P < 0.05). In comparison to anthropometric measurements, total body fat percentage had a stronger relationship with 25(OH)D (-0.261).

A secondary analysis of data from middle-aged healthy adults with excess weight (56) indicated that total and abdominal adiposity, but not peripheral adiposity, predicted low plasma 25(OH)D total fat mass index (FMI): P=0.018; android FMI: P=0.052; gynoid FMI: P=0.15; appendicular FMI: P=0.07). An additional retrospective analysis (78) on women found lower levels of 25(OH)D among obese individuals (P<0.001).

Bell *et al.* (46), in their study among 12 obese and 14 nonobese White subjects, demonstrated that mean serum 25(OH)D (8 \pm 1 vs 20 \pm 2 ng/mL, P < 0.001) was significantly lower in the obese than in the nonobese subjects.

There was a significant inverse association between abdominal obesity and the quintiles of 25(OH)D levels in another cross-sectional study (44) involving both

Valiña-Tóth *et al.* (64) showed in a study of healthy overweight adults that 25(OH)D inversely related to truncal fat mass (P=0.02). Another study (79), among women with BMI 18–57 kg/m², showed that higher BMI was related with lower 25(OH)D levels (r > -0.27, P < 0.001).

Serum 25(OH)D was found to be inversely related to visceral abdominal fat and percentage fat in a cross-sectional study of overweight and obesity (77).

After adjusting for the confounders, Shan *et al.* (13) found that low 25(OH)D levels were significantly related to elevated WC among women (OR=1.612 (1.014–2.561). According to Puntus *et al.* (62) increasing BMI significantly reduced 25(OH)D levels in preand postmenopausal women (P < 0.001 and P < 0.05, respectively).

Further research (76) revealed a significant negative relationship between fat mass and 25(OH)D among fit older women (β = -3.76, P < 0.001).

Researchers examined a total of 830 healthy adults (55) and found that vitamin D insufficiency was

significantly linked to abdominal obesity in males (OR 2.75 (CI: 1.1-7.1); P < 0.05).

Shinkov *et al.* (59) reported that 25(OH)D levels were significantly lower in obese females than in the normal weight females (34.6 \pm 16.2 vs 38.2 \pm 17.8 nmol/L, P=0.014), but, in the males, the 25(OH)D levels did not differ among the BMI groups.

A significant negative correlation was reported by Shafinaz *et al.* (54) between serum 25(OH)D level and body fat percentage (β =-0.14). Multivariate linear regression analysis found that higher BMI and larger WC were significantly associated with lower serum 25(OH)D levels (P 0.05).

A study by Gariballa *et al.* (48) found that, although BMI did not differ statistically significantly between groups, it was higher among vitamin D deficient older subjects and women <50 years, respectively, compared to individuals with adequate vitamin D or optimal concentrations (P=0.05).

Another study (52) found a negative correlation between serum 25(OH)D levels and BMI (β : -0.150; 95% CI: -2.262, -0.037). A study (66) in severely obese subjects found that insufficient 25(OH)D corresponded to higher BMI (insufficient: 47.2 ± 5.6 vs not insufficient: 45.9 ± 4.7 kg/m²; P=0.047).

Another cross-sectional investigation (84) among community-dwelling men and women reported that compared with normal weight, obese individuals had lower 25(OH)D levels (P < 0.05).

In 2023, Avila Castillo *et al.* (85), in their study on the 1032 adult population of the LIFE-Adult-Study, concluded that low levels of 25(OH)D were linked to higher BMI, while fat mass areas showed a negative correlation with 25(OH)D concentrations only in women.

Discussion

The present systematic review investigated the correlation between serum 25(OH)D levels and anthropometric and adiposity measurements in healthy individuals of various ages. Some previous meta-analyses included studies that reported BMI (86) and WC (87) as an index of weight status in adults only, while our systematic review has considered different adiposity measures in both adults and children. The results of most of the included studies, but not all papers, showed lower 25(OH)D levels and a higher prevalence of vitamin D insufficiency and deficiency in subjects with higher weight, BMI, WC, WHR, and fat mass percentage, in all age groups. In fact, 25(OH)D levels were inversely associated with BMI, WC, HC, WHR, and body fat mass percentage, specifically in female subjects.

Adiposity indicators were inversely associated with vitamin D status in numerous studies but not in all studies. The association between low vitamin D levels

and obesity may be attributed to several factors. First, individuals with obesity might face challenges in acquiring sufficient sun exposure owing to limited mobility or specific clothing choices (88, 89, 90). Second, vitamin D is stored in fat compartments and adipose tissues, particularly in the abdominal area, making its release less accessible in obese individuals (88, 91). Third, individuals with obesity often have an increased demand for vitamin D to support their body weight, but the bioavailability of 25(OH)D is reduced, making it challenging to meet these elevated requirements (88, 91). Fourth, as the concentrations of active vitamin D metabolites increase, they initiate a negative feedback control over hepatic 25(OH)D synthesis. This feedback mechanism led to a reduction in serum 25(OH)D levels, providing an additional explanation for the observed association (91). It is important to note that the full acceptance of these mechanisms is still under investigation (92). Conversely, reducing fat mass has been shown to elevate 25(OH)D levels by releasing stored vitamin D into circulation. In a systematic review, Mallard et al. (93) concluded that weight reduction slightly elevated the 25(OH)D level by 1.5 ng/mL. They proposed that the release of vitamin D from fat and fat-free mass after weight loss was responsible for 25(OH)D elevation. Therefore, there is contradictory evidence regarding this issue, and the accurate relationship between vitamin D and adiposity indicators remains unclear.

According to the results of the studies included in the systematic review, overweight and obese individuals of different ages have similar chances of becoming vitamin D-deficient. Hence, age does not appear to have a significant impact on this association.

As a result of the heterogeneity in study characteristics, findings on vitamin D status and adiposity were inconsistent, with the inverse association being more prominent in females. It has been speculated that ethnicity, sex, and age may have a mediating effect on the relationship between 25(OH)D levels and anthropometric measures. It is likely that the differences in associations between females and males stem from the fact that women have a higher percentage of body fat and a different body composition than men. With the same BMI, men have less body fat than women. Thus, men store less vitamin D in adipose tissue and more remains in the blood. Furthermore, serum 25(OH)D levels are not stable throughout the year due to inadequate levels of 25(OH)D in the adipose tissue. In addition, vitamin D-binding protein could also contribute to sex differences in vitamin D status (94). It has been demonstrated that vitamin D-binding protein and adiposity are negatively correlated in men and positively correlated in women (94).

In reviewing the studies included in the systematic review, critical issues were raised, which could contribute to bias and confounding. There are several limitations, including heterogeneity in participant

characteristics, the diversity of methods used to determine vitamin D levels and the analytical challenges involved, the use of variable definitions of hypovitaminosis D, the absence of adjustment for various confounding factors that influence vitamin D levels, and the reliability of various adiposity measures for describing obesity. The results of these studies may also be affected by a number of other factors related to the population studied, such as cultural and religious factors, dressing codes that mandate covering the majority of the body surface, and behavioral and lifestyle differences (95). It is also possible that variations in socioeconomic and developmental status could cause heterogeneity among studies that influence nutrition and lifestyle. Various socioeconomic and developmental factors influence the prevalence of obesity and vitamin D level (96, 97). Moreover, the amount of air pollution may affect vitamin D status, especially in urban areas where UVB wavelengths of sunlight are mostly blocked by pollutants (98). As a result of differences in health policies regarding the fortification of food with vitamin D in Europe and the USA and in national recommendations regarding vitamin D supplement use, vitamin D intake may differ significantly between countries (99). Vitamin D levels are also affected by genetic factors; however, without population-based genetic analyses, it is difficult to quantify their impact (100).

On the other hand, it has been suggested that low serum 25(OH)D concentrations reduce calcium absorption. PTH is secreted in response to low serum calcium concentrations, which stimulates the production of 1,25(OH)₂D. The (nearly) normal serum levels of 1,25(OH)₂D are maintained at the expense of high serum PTH concentrations, known as 'secondary hyperparathyroidism'. Considering that serum 25(OH)D is the substrate for serum 1,25(OH)₂D, serum 25(OH)D levels tend to decrease when serum 1,25(OH)₂D increases. PTH may contribute to fat accumulation by increasing insulin resistance and inhibiting lipolysis (101). Additionally, vitamin D may regulate uncoupling proteins, which may play a role in energy metabolism (102). Despite this, Walsh et al. (53) found that PTH was not affected by BMI or sex and was not correlated with BMI. In obese individuals, there may be alterations in the relationship between 25(OH)D and PTH levels (53). To optimally determine the vitamin D status, the 25(OH)D threshold for maximum suppression of PTH has been suggested (103). A previous study showed that patients with a BMI ≥30 kg/m² had a lower threshold for suppressing PTH levels (5 ng/mL) than patients with a BMI <30 kg/m² (10 ng/mL) (104). This suggests that a very low 25(OH)D level is required to activate the PTH axis, leading to secondary hyperparathyroidism and bone loss (104).

The BMI has been used as an indicator of obesity in the majority of studies included in this systematic review.

However, although BMI is the most widely accepted method for defining obesity, it is not an accurate measure of fat mass and distribution of body fat. Compared with subcutaneous fat, excess visceral fat confers a greater risk of metabolic and cardiovascular diseases for the same BMI value (105, 106). Several more meaningful measures of adiposity have been developed to resolve this methodological issue, including fat mass, WC, and WHR.

The present systematic review investigated the correlation between serum 25(OH)D and anthropometric and adiposity measurements in healthy individuals of various ages. However, some limitations must be considered. As not all included studies separately reported the relationship between serum vitamin D and adiposity measures in men and women, accurate estimates for men and women could not be provided. Different cutoff points were used for defining obesity and vitamin D deficiency in the included studies. This systematic review also examined the relationship between obesity and vitamin D deficiency using cross-sectional studies, which made the causality findings more difficult. Finally, we did not have access to the complete data of all related papers.

Conclusion

Our systematic review highlights the prevalence of vitamin D deficiency among overweight/obese individuals and its inverse correlation with adiposity measures. Despite this association, it is essential to acknowledge the influence of various confounding factors, including dietary intake, physical activity, educational level, and seasonal variations, which could impact the serum 25(OH)D levels. Further prospective investigations are warranted to establish a causal relationship between vitamin D levels and obesity, shedding light on the underlying mechanisms. Additionally, the findings underscore the importance of monitoring serum vitamin D levels in individuals with excess weight. Considering the variability in climate and dietary patterns across different regions, standardizing the 25(OH)D cut-off points would benefit from additional research. Increased awareness of the interplay between obesity and vitamin D levels can prompt adjustments in clinical approaches among nutritionists and health-care professionals. Because the field of vitamin D research is dynamic and continues to evolve, and new studies may indeed provide further insights into the relationship between serum 25(OH)D levels and adiposity. In addition, the scarcity of cohort studies underscores the need for further longitudinal investigations to elucidate the causative mechanisms linking vitamin D with adiposity. So, we recommend that future researchers consider conducting updated systematic reviews to integrate the latest evidence on this important topic.

Declaration of interest

The authors declare that there is no conflict of interest that could be perceived as prejudicing the impartiality of the study reported.

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Data availability

The datasets used and/or analyzed during the current study are available from the corresponding author on reasonable request.

Consent for publication

All authors have given consent for the paper to be published by the corresponding author.

Author contribution statement

B.A. and F.H. designed and wrote the manuscript. S.H. and M.M. performed interpretation of results and critical revision of the manuscript. F.A. and M.V. critically revised the manuscript. All authors read and approved the final version.

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