

The annual meeting of the Irish Section of the Nutrition Society was held at University College Cork, Cork, Republic of Ireland on 18–20 June 2003

Symposium on ‘Optimal nutrition for osteoporosis prevention’

Geographical differences in vitamin D status, with particular reference to European countries

Lars Ovesen*, Rikke Andersen and Jette Jakobsen

Institute of Food Safety and Nutrition, The Danish Veterinary and Food Administration, 30A Sydmarken, 2860 Søborg, Denmark

Vitamin D is produced endogenously when the skin is exposed to sunlight and can be obtained exogenously from a few natural food sources, from food fortification and from supplements. Generally, vitamin D intake is low $\leq 2\text{--}3\ \mu\text{g/d}$ in Europe. Casual exposure to sunlight is thought to provide most of the vitamin D requirement of the human population. However, skin synthesis of vitamin D may not compensate for the low nutritional intake in Europe, even in countries with high supplies from food fortification and supplements. For assessment of vitamin D nutritional status the concentration of 25-hydroxyvitamin D (25(OH)D) in serum is considered to be an accurate integrative measure reflecting an individual's dietary intake and cutaneous production. A substantial percentage of the elderly and adolescents in Europe have a low concentration of 25(OH)D; in the elderly this percentage ranges from approximately 10 in the Nordic countries to approximately 40 in France. Low vitamin D status seems to be aggravated by disease and immobility, and by a low frequency of supplement use.

Vitamin D intake: Vitamin D recommendations: 25-Hydroxyvitamin D: Elderly: Adolescents: Europe

Vitamin D is produced endogenously when the skin is exposed to sunlight and is obtained exogenously from natural food sources, from food fortification and supplements. Vitamin D requires activation by two successive hydroxylation steps: first in the liver to form 25-hydroxyvitamin D (25(OH)D); then in the kidney to form 1,25-dihydroxyvitamin D (Parfitt *et al.* 1982). The 25-hydroxylation is very fast and almost unregulated in contrast to the 1-hydroxylation that is strictly regulated by parathyroid hormone (PTH; Shepard & DeLuca, 1980). Vitamin D status is commonly defined by the level of 25(OH)D in circulating blood. Inadequate vitamin D intake and/or insufficient sunlight exposure reduces serum 25(OH)D concentration causing secondary hyperparathyroidism, which increases the risk of osteoporosis (Aloia *et al.* 1985; Lips *et al.* 1987) and fractures in the elderly (von Knorring *et al.* 1982; Morris *et al.* 1984; Lips *et al.* 1985; Meller *et al.* 1985).

International comparison studies have shown that among adults serum concentrations of 25(OH)D vary between

countries, but are often higher in the USA and Canada and in Scandinavia, compared with the rest of Europe (McKenna, 1992; Lips, 2001). Contrary to expectations the European Euronut SENECA Study found a positive association between latitude and mean serum 25(OH)D, with lower concentrations in Greece and Spain than in Norway (van der Wielen *et al.* 1995). These differences in vitamin D status must be accounted for by international differences, still largely unexplained, in contributions to the vitamin D supply from food, including the supply from fortification and supplements, and the extent of solar skin exposure.

As individuals age the concentration of 25(OH)D decreases, mainly because of a declining efficiency of skin to produce vitamin D (MacLaughlin & Holick, 1985). Also, the elderly may spend less time outdoors, especially if institutionalised, and their food intake (including fish intake) decreases, adding to the risk of vitamin D deficiency and consequently osteoporosis (Lips *et al.* 1987).

The present short review focuses on comparisons of vitamin D intake and frequency of low vitamin D status

Abbreviations: PTH, parathyroid hormone; 25(OH)D, 25-hydroxyvitamin D.

***Corresponding author:** Dr Lars Ovesen, fax +45 33 95 11 19, email lov@fdir.dk

among healthy populations in Europe, with special emphasis on the vitamin D nutritional status in the elderly who, for the reasons given earlier, are at an increased risk of an insufficient vitamin D supply.

Vitamin D status

For assessment of vitamin D nutritional status the concentration of 25(OH)D in serum is considered as an accurate integrative measure reflecting an individual's dietary intake and cutaneous production (Parfitt, 1998).

Many studies have suggested that there is a value of 25(OH)D above which there is little further decrease in PTH (Ooms *et al.* 1995b; Chapuy *et al.* 1997; Reginster *et al.* 1998; Melin *et al.* 1999; Souberbielle *et al.* 2001). However, the threshold concentration of 25(OH)D that delimits deficiency from sufficiency, i.e. the lowest concentration defining the plateau level of PTH, differs between studies and ranges from 20 nmol/l to >100 nmol/l (Gloth *et al.* 1995; Ooms *et al.* 1995b; Chapuy *et al.* 1997; Dawson-Hughes *et al.* 1997; Holick, 1998; Thomas *et al.* 1998). Some studies, however, have shown a continuous decline in PTH with increasing 25(OH)D and no plateau (Bates *et al.* 2003; Kudlacek *et al.* 2003). Other studies have included bone resorption markers (hydroxyproline, pyridinoline and deoxypyridinoline) to derive a 25(OH)D concentration that distinguishes sufficiency from deficiency (Sahota *et al.* 1999; Jesudason *et al.* 2002). The concentration of 1,25-dihydroxyvitamin D, the biologically-active form of vitamin D, will usually be normal or even slightly elevated in vitamin D deficiency, and therefore provides no information on nutritional status (Hollis, 1996). The blood native vitamin D concentration of an individual reflects recent intake of vitamin D and/or exposure to sunlight, and therefore may vary greatly over a short time period.

Presently, there is no clear definition of an absolute value for 25(OH)D above which an individual is considered to be vitamin D replete. Alternatively, a gradual scale has been proposed in which hypovitaminosis D is defined as a 25(OH)D concentration of <100 nmol/l, vitamin D insufficiency as a 25(OH)D concentration of <50 nmol/l and vitamin D deficiency as a 25(OH)D concentration of <25 nmol/l (McKenna & Freaney, 1998).

The classical vitamin D-deficiency diseases, rickets in children and osteomalacia in adults, are caused by severe lack of vitamin D (Hutchison & Bell, 1992), usually with serum 25(OH)D concentrations <10 nmol/l (and hypocalcaemia, hypophosphataemia and increased serum alkaline phosphatase). Rickets and osteomalacia are a major problem in some population groups in Europe, whether as a result of immobility or infirmity (Lips, 2001), or social or religious customs practised by certain immigrant groups (Meulmeester *et al.* 1990; Iqbal *et al.* 1994; Solanki *et al.* 1995; Glerup, 2000), especially those who are living in north western Europe where sunlight exposure is limited.

Solar exposure

The relationship between lack of sunlight and vitamin D deficiency has been recognised for many years. During exposure to sunlight u.v. B photons (wavelength 290 and

315 nm) photolyse epidermal stores of 7-dehydrocholesterol (provitamin D) to form previtamin D, which is unstable and subsequently undergoes isomerisation to the stable vitamin D (Holick *et al.* 1980). After its formation vitamin D slowly moves into the circulation.

Casual exposure to sunlight is thought to provide most of the vitamin D requirement of the human population (Holick, 1996). However, skin synthesis of vitamin D may not compensate for the low nutritional intake in Europe, because Europe is located at high latitude, from approximately 40°N (Madrid, Spain) to 64°N (Reykjavik, Iceland). Webb *et al.* (1988) in the USA demonstrated that in Boston, MA (latitude 42°N) photosynthesis of previtamin D is nearly impossible during the winter months (November–February) and in Edmonton, Canada (latitude 52°N) vitamin D synthesis is impaired from October to March. Model calculations also demonstrate the considerable contrast in skin production of vitamin D between sites of different latitude and that low vitamin D status is more likely to develop in locations where solar u.v. levels are low for much of the year (Kimlin *et al.* 2003).

A number of studies from many European countries have shown that vitamin D status exhibits a seasonal variation. Both serum concentrations of 25(OH)D (Juttman *et al.* 1981; Tjellesen & Christiansen, 1983; Hegarty *et al.* 1994; Scharla *et al.* 1996; Brot *et al.* 2001) and biomarkers of bone turnover (Woitge *et al.* 1998) are elevated in late summer and decrease during winter, whereas PTH concentrations tend to increase during winter (Krall *et al.* 1989; Hegarty *et al.* 1994; Chapuy *et al.* 1996). The influence of sun exposure on vitamin D status is underlined in a large Danish study of 2016 post-menopausal females, which showed that those who sunbathed regularly had higher year-round serum 25(OH)D concentrations compared with those who reported that they usually avoided direct sun exposure (Brot *et al.* 2001).

However, the relationship between u.v. radiation and cutaneous vitamin D production is complex and, apart from time of day, latitude, altitude and season, depends on a large number of other factors, such as the amount of air pollution, cloud cover, concentration of O₃ in the atmosphere and stratosphere, surface albedo, age of the individual and the amount of clothing worn, sunscreen usage and skin pigmentation (Holick, 1995). Some of these factors specifically add to the increased risk of an insufficient vitamin D supply in the elderly across Europe, e.g. decreased outdoor activities because of immobility, decreased capacity of their skin to produce vitamin D because of age-related changes and use of clothing that diminishes the cutaneous production of vitamin D.

Dietary intake

Vitamin D is found naturally in only a few foods, mainly in fish and in lesser amounts in eggs (yolk) and meat products. Animal foods contributing to dietary vitamin D also contain 25(OH)D. Although this vitamin D metabolite has a higher activity than native vitamin D, its precise potency is not known (Ovesen *et al.* 2003).

Table 1 shows the intake of vitamin D in healthy elderly subjects based on results from representative dietary intake

Table 1. Mean dietary vitamin D intake in representative samples of independent elderly subjects and dietary vitamin D recommendations in some European countries (intake from supplements are not included)

Country	Survey	Method	Intake ($\mu\text{g}/\text{d}$)				Recommendation†	
			Age (years)	Men	<i>n</i>	Women		<i>n</i>
Denmark	The Danish Dietary Survey, 1995 (Danish Food Agency, 1996)	7 d record	65–74 75–80	3.3 3.2	103 44	4.1 3.7	122 64	Recommended dietary intake: ≥ 61 years $10 \mu\text{g}/\text{d}$ In DK subjects > 65 years are recommended $10 \mu\text{g}$ as supplement
Norway	Norkost 1997 (Johansen & Solvoll, 1999)	Food-frequency questionnaire	60–69 70–79	5.6 6.0	131 106	4.0 4.0	137 109	
Sweden	Riksmaten 1997–8 (Becker & Pearson, 2002)	7 d record	> 65	7.1	65	4.9	58	
France	INCA 1999 (Volatier, 2000)	7 d record	> 65		2.5	245*		Recommended dietary intake: Elderly $10 \mu\text{g}/\text{d}$
The Netherlands	The Third Dutch National Food Consumption Survey 1997–8 (Hulshof <i>et al.</i> 1998)	2 d record	> 65	4.8	185	3.6	236	Adequate intake: 61–70 years $7.5 \mu\text{g}$ ≥ 71 years $12.5 \mu\text{g}$ Higher recommendations with limited exposure to sunlight and with dark skin colour
Germany	Ernährungsbericht 2000 (Deutsche Gesellschaft für Ernährung, 2000)	3 d record	65–74 75–84 > 65	3.7 3.5 4.1	361 126 23	3.1 3.0 2.6	503 285 73	Population reference intake: ≥ 66 years $10 \mu\text{g}$
UK	National Diet and Nutrition Survey. People aged 65 years and over (Finch <i>et al.</i> 1998)	4 d record	65–74 75–84 > 85	4.3 3.8 3.2	271 265 96	3.0 3.0 2.3	256 217 170	Reference nutrient intake: > 65 years $10 \mu\text{g}$ $10 \mu\text{g}$ for those confined indoors, irrespective of age

DK, Denmark.

*Intake and no. of subjects for men and women combined.

† Different terms are used, e.g. recommended dietary allowance (RDA), recommended dietary intake, recommended nutrient intake or population reference intake. The recommended intake is defined as the intake of an essential nutrient considered as being adequate to meet known nutritional needs of practically all healthy individuals. The recommended intake is normally calculated as the average requirement + 2 SD. An adequate intake is set instead of an RDA if sufficient scientific evidence is not judged to be available to calculate an estimated average requirement. The average intake is based on observed or experimentally-determined estimates of average nutrient intake by a group of healthy individuals.

studies. Generally, vitamin D intake is low, approximately 2–3 $\mu\text{g}/\text{d}$, in Europe (except in Norway and Sweden where fish intake and the contribution from fortified foods are high). McKenna (1992) reported that the mean vitamin D intake is significantly ($P < 0.0001$) lower in Central Europe (2.5 $\mu\text{g}/\text{d}$) than in North America (6.2 $\mu\text{g}/\text{d}$) or Scandinavia (5.2 $\mu\text{g}/\text{d}$).

Direct comparison between countries is difficult, however, because different methods are used to estimate food intake. Furthermore, food tables differ in the way vitamin D content is expressed. Some food tables use values derived by imprecise bioassay techniques, while other tables use analytical data derived by more modern chemical methods for vitamin D determination that often do not include 25(OH)D (Deharveng *et al.* 1999). The imprecision is further augmented by the extreme paucity of systematic analytical data, and the uncertainty of the biological activity of 25(OH)D (Ovesen *et al.* 2003).

Dietary recommendations

There is some variation in dietary recommendations for vitamin D among European countries (Table 1). Evidently, the difficulty in setting daily recommendations for vitamin D arises from the dual nature of its supply, and as the amount of vitamin D that originate from endogenous

production varies, a recommendation cannot be determined accurately (Prentice, 2002). In most countries the recommended daily intake is 5–10 $\mu\text{g}/\text{d}$, often at the higher intake levels in the elderly (and in infants) with less opportunity to produce vitamin D cutaneously. Some national committees recommend a higher intake for the elderly because they have a high risk of vitamin D deficiency. It should be noted that there is controversy about the daily intake of vitamin D required to meet or sustain 'normal' concentrations of 25(OH)D in the blood, and that some research groups advocate a much higher intake of vitamin D (50–100 $\mu\text{g}/\text{d}$) in populations that do not have substantial body stores of vitamin D (Malabanan *et al.* 1988; Vieth *et al.* 2001; Heaney *et al.* 2003).

Food fortification

While food fortification continues to be a widely-used mechanism for increasing vitamin D intake in many industrialised countries, prevailing attitudes, and relevant legislation, differ, and there is no general consensus on the extent to which vitamin D fortification should be practised (Nordic Council of Ministers, 1995). In some countries the addition of vitamin D is completely unregulated (e.g. UK, although it is mandatory in the case of margarine), while in the countries where there is most restriction (e.g. Scandi-

navia) fortification is permitted only after authorisation based on a scientifically-documented public health need. Some countries have instituted mandatory fortification of certain foodstuffs (the addition of vitamin D to margarine in the UK and The Netherlands), while other countries have a voluntary fortification programme (e.g. Finland allows addition of vitamin D to milk and margarine). The maximum amounts of vitamin D that can be added to foods differ widely between countries (e.g. in margarine from approximately 20 µg/kg to >100 µg/kg); however, in most countries the level is approximately 70–80 µg/kg. The level of fortification and the selection and extension of the range of foods fortified, as well as the actual intake of the target group, will determine the effect on vitamin D status, and will obviously differ between countries.

However, the effect of inclusion of fortified foods on vitamin D status of the most important target group, the frail elderly, has been modest. Studies of elderly subjects conducted in the UK have found small increases in 25(OH)D concentrations in those consuming fortified margarine (Scragg *et al.* 1995) and milk (Keane *et al.* 1992, 1998). Significant (compared with the control group; $P < 0.001$) increases in 25(OH)D were observed in frail elderly Dutch subjects who every day consumed fortified dairy products that brought their intake of vitamin D up to recommended levels (from 3.2 µg/d to 11.6 µg/d; de Jong *et al.* 1999). In contrast, no increases in 25(OH)D were found in a Scottish study of long-stay residents of geriatric wards who received fortified foods (margarine, butter or milk) as part of their daily diet for a period of 0.5–1 year (Dunnigan *et al.* 1986).

Supplementation

The effect of vitamin D supplementation on the serum concentration of 25(OH)D has been demonstrated in several studies worldwide, but the wide variation among European countries in the use of vitamin D-containing supplements will affect vitamin D status. The use of supplements is generally much higher in the Scandinavian countries than in the rest of Europe. For example, studies of elderly subjects from Norway (Sem *et al.* 1987) and Denmark (Knudsen *et al.* 2002) showed that more than half used vitamin D supplements, while a study from Iceland reported that 83 % of the elderly took cod-liver oil or vitamin D supplements (Sigurdsson *et al.* 2000). In contrast in the UK about 16 % of the free-living elderly and only 3 % of the institutionalised elderly received vitamin D supplements (Bates *et al.* 2003).

Relative contributions of intake and solar exposure

During the past 20 years the measurement of serum 25(OH)D has been routinely available and many reports have been published that provide evidence for an association (within country) between vitamin D intake and the extent of solar exposure, and serum 25(OH)D. The British National Diet and Nutrition Survey of subjects aged >65 years found that in the non-institutionalised subjects there was a strong association (linear regression; $P < 0.0001$) between 25(OH)D concentrations and vitamin D intake in autumn, winter and spring, but not in the summer season

(Bates *et al.* 2003). This pattern was similar to that observed in preschool children for whom vitamin D intake was significantly associated (linear regression; $P < 0.006$) with 25(OH)D in the winter but not in the summer months (Davies *et al.* 1999), suggesting a higher dependency on dietary vitamin D during seasons with low solar exposure.

Most European populations require regular sun exposure during the summer to build up sufficient stores to ensure adequate vitamin D status during winter and spring, even in younger individuals. An increased dependence on dietary vitamin D occurs when there is restricted skin exposure to sunlight (housebound or institutionalised subjects, individuals wearing protective clothing) and reduced capacity for endogenous synthesis (dark skin, habitual use of sunscreen).

Vitamin D status in Europe

McKenna (1992) reviewed vitamin D status based on serum concentrations of 25(OH)D in 117 studies from twenty-seven regions and found large regional variations in young adults and the elderly. When results were subsequently grouped according to three geographical regions (North America, Scandinavia and Central and Western Europe) serum concentrations of 25(OH)D were found to be markedly higher in the USA and Canada and in Scandinavia compared with the rest of Europe, a difference that could be explained, at least partly, by the higher intakes of vitamin D from fortification and supplements in North America and in Scandinavia.

However, conclusions of regional differences based on comparisons of 25(OH)D concentrations are questionable because of the lack of standardisation between methods. Until the early 1990s analytical methods were mainly competitive protein-binding assays with or without a chromatographic step to isolate 25(OH)D that were developed in-house. For the last 10 years commercial radio-immunoassays have been used, but the large discrepancies found in international studies between 25(OH)D measurements derived with different competitive protein-binding assays and radioimmunoassays prevent comparison between countries (Lips *et al.* 1999). An additional problem is that some immunoassays do not measure 25-hydroxyergocalciferol (25(OH)D₂), and therefore underestimate total 25(OH)D (subjects may have major amounts of the plant-derived 25(OH)D₂ in their blood, e.g. from pharmaceutical vitamin D preparations; Hollis, 2000).

Elderly

A large number of independent studies worldwide have shown that the elderly often have low serum concentrations of 25(OH)D coupled with high levels of PTH (Lips, 2001). Concentrations of 25(OH)D have been repeatedly found to be higher in independent healthy subjects compared with patients in hospitals and residents of nursing homes, indicating the increased reliance of the institutionalised elderly on vitamin D in food.

Within Europe mean 25(OH)D concentrations vary widely (Table 2). The lowest serum 25(OH)D concentrations are found in the Republic of Ireland and The Netherlands, with the highest concentrations in the Nordic countries and intermediate concentrations in other European

Table 2. Studies on vitamin D status in independent elderly populations from several European countries arranged according to latitude (north to south) (Mean values and standard deviations)

Country	Study	Age (years)		Gender	n	Serum 25(OH)D (nmol/l)		Lower reference limit (nmol/l)	Vitamin D deficiency (%)	Season
		Mean	SD			Mean	SD			
Iceland	Sigurdsson <i>et al.</i> (2000)	70		F	308	53	20	30	13	Winter
Sweden	Melin <i>et al.</i> (1999)	≥80		M	23	70	23	25	4	Winter
				F	81	65	30			
Denmark	Brot <i>et al.</i> (2001)	45–58*		F	2016	63		25	7	All the seasons
								50	40	
Republic of Ireland	McKenna <i>et al.</i> (1985)	69	5	M, F	30	21		25	21	Winter
The Netherlands	Lips <i>et al.</i> (1987)	76	4	M, F	74	33	14	20	16	All the seasons
Belgium	Boonen <i>et al.</i> (1997)	72	5	M	40	47	18	30	18	All the seasons
UK	Bates <i>et al.</i> (1999)	> 65		M, F	approx 800	56	27	12.5	2	All the seasons
								25	8	
Austria	Kudlacek <i>et al.</i> (2003)	> 60		M	40	57	31	25	26	Winter
				F	65	43	27			
Germany	Scharla <i>et al.</i> (1996)	50–80*		M, F	415	43	23	25	24	Winter
France	Chapuy <i>et al.</i> (1996)	80	3	F	440	43	25	30	39	Winter
Italy	Bettica <i>et al.</i> (1999)	59	8	F	570	45	20	30	28	All the seasons

25(OH)D, 25-hydroxyvitamin D.

* Range.

countries. The frequency of vitamin D deficiency obviously depends on the lower reference limit and the season. However, a substantial proportion of the independent elderly population of Europe is vitamin D deficient, and this proportion seems to be lower in the Nordic countries and UK compared with the rest of Europe.

Studies from The Netherlands (Ooms *et al.* 1995a) and Sweden (Toss *et al.* 1980) have shown that about one-third of the elderly in nursing homes and in homes for the elderly are vitamin D deficient (serum 25(OH)D < 20 nmol/l), and this number may be even higher in the Republic of Ireland where mean 25(OH)D concentrations of 9 nmol/l have been reported for nursing home residents (McKenna *et al.* 1985). In the UK 2.7 and 37 % of the institutionalised participants in the 1994–5 National Diet and Nutrition Survey of individuals ≥ 65 years had 25(OH)D concentrations < 12.5 and < 25 nmol/l respectively (Bates *et al.* 1999). Very low serum 25(OH)D concentrations were found for geriatric patients in the UK (Corless *et al.* 1975; Nayal *et al.* 1978; Vir & Love 1978), Spain (Quesada *et al.* 1989) and France (Fardellone *et al.* 1995); mean concentrations ranged from 3 nmol/l to 12 nmol/l and > 50 % had severe vitamin D deficiency (serum 25(OH)D < 12.5 nmol/l). In Denmark 44 % of geriatric patients had severe vitamin D deficiency (Egsmose *et al.* 1987). In Sweden vitamin D deficiency, defined as serum 25(OH)D < 30 nmol/l, was found in approximately 50 % of hospitalised women and 35 % of hospitalised men and in 20 % of the home-living elderly (Mowé *et al.* 1996). Severe vitamin D deficiency (serum 25(OH)D < 20 nmol/l) was found in 20 % of hospitalised elderly men and 26 % of hospitalised elderly women in Finland; for a comparable group of outpatients the respective prevalences were much lower, 6 % for men and 2 % for women (Kauppinen-Mäkelin *et al.* 2001).

The European Euronut SENECA Study was conducted in nineteen centres in eleven countries using a central laboratory facility (van der Wielen *et al.* 1995). Lowest mean winter serum 25(OH)D concentrations were found in study centres in Greece (21 nmol/l in women and 25 nmol/l in men) and the highest in Norway (48 nmol/l in women and 45 nmol/l in men). Serum 25(OH)D concentrations were positively associated with latitude. Low 25(OH)D concentrations could be explained by reduced sunlight exposure (time spent outdoors, clothing worn when exposed to sunlight), low intake of fish and low physical health status. The low concentrations of 25(OH)D in southern Europe might also be explained by limited fortification of foods with vitamin D and low frequency of supplement use.

Other adult age-groups

Low vitamin D status has also been demonstrated in the healthy adult population in Europe. In 328 Finnish adults aged 31–43 years low wintertime 25(OH)D concentrations (< 25 nmol/l) were found in 26.2 % of the women and 28.6 % of the men (Lamberg-Allardt *et al.* 2001). In a French urban population comprising 765 men aged 45–65 years and 804 women aged 35–60 years selected from twenty cities 14 % of the subjects had 25(OH)D concentrations < 30 nmol/l (Chapuy *et al.* 1997). In a representative sample (*n* 3276) of the general adult Swiss population 6 % were found to have 25(OH)D concentrations of 20 nmol/l (Burnand *et al.* 1992). Even in southern Italy a relatively high prevalence of vitamin D deficiency can be found among healthy young women (Carnevale *et al.* 2001). The prevalence of hypovitaminosis D, defined by concentrations of 25(OH)D < 30 nmol/l, was 27.8 % in winter and 3.4 % in summer.

Adolescents

Maximising the peak bone mass during adolescence and early adulthood is considered to be the best protection against age-related bone loss and later risk of osteoporosis (Heaney *et al.* 2000). Thus, adolescence becomes a critical period in skeletal development. However, in these stages of life the concentration of 25(OH)D considered necessary to achieve optimal bone mass is ill defined, partly because adolescence may be a time when physiological mechanisms other than the concentration of 25(OH)D play a role in regulating the secretion of PTH (Guillemant *et al.* 1995; Outila *et al.* 2001).

About 18 % of Icelandic 16–20-year-old girls had winter serum 25(OH)D concentrations of <25 nmol/l (Kristinsson *et al.* 1998) and 14 % of Finnish females had winter serum 25(OH)D concentrations of <20 nmol/l (Lehtonen-Veromaa *et al.* 1999). Other studies from Finland have shown a similar high frequency of low concentrations of 25(OH)D in female adolescents. Furthermore, low serum 25(OH)D concentration was associated with lower bone mineral density (Outila *et al.* 2001), and also higher bone resorption and lower gain in bone mass over 3 years (Lehtonen-Veromaa *et al.* 2002). A study from France demonstrated large differences in 25(OH)D values, ranging from high concentrations at the end of summer (58.5 (SD 18.0) nmol/l) to low concentrations at the end of winter (20.6 (SD 6.0) nmol/l), when almost all adolescents had a 25(OH)D concentration <30 nmol/l (Guillemant *et al.* 1999). Finally, data from Spain have shown that in winter 31 % of adolescents had 25(OH)D concentrations below this cut-off level (Docio *et al.* 1998).

Conclusion

Very few of the population in Europe receive anywhere near the recommended dietary intake of vitamin D. A low vitamin D status, defined by a low 25(OH)D concentration in the blood, is found in a large proportion of the elderly in Europe, and may hasten the development of osteoporosis and increase fracture risk. Low vitamin D status is more frequent in southern Europe than in the Scandinavian countries, probably as a result of a higher intake of vitamin D from supplements in Scandinavia. Vitamin D deficiency is exacerbated by disease and immobility in the elderly. Low 25(OH)D concentrations are also found in the adult and adolescent population in Europe. However, the importance of vitamin D deficiency in these age-groups in relation to peak bone mass and the later development of osteoporosis is not known.

References

Aloia JF, Cohn SH, Vaswani A, Yeh JK, Yuen K & Ellis K (1985) Risk factors for postmenopausal osteoporosis. *American Journal of Medicine* **78**, 95–100.

Bates CJ, Carter GD, Mishra GD, O'Shea D, Jones J & Prentice A (2003) In a population study, can parathyroid hormone aid the definition of adequate vitamin D status? A study of people aged 65 years and over from the British National Diet and Nutrition Survey. *Osteoporosis International* **4**, 152–159.

Bates CJ, Prentice A, Cole TJ, van der Pols JC, Doyle W, Finch S, Smithers G & Clarke PC (1999) Micronutrients: highlights and research challenges from the 1994–5 National Diet and Nutrition Survey of people aged 65 years and over. *British Journal of Nutrition* **82**, 7–15.

Becker W & Pearson M (2002) *Kostvanor och näringsintag i Sverige*. (Dietary Habits and Nutrient Intake in Sweden). Uppsala, Sweden: Statens Livsmedelsverket.

Bettica P, Bevilacqua M, Vago T & Norbiato G (1999) High prevalence of hypovitaminosis D among free-living postmenopausal women referred to an osteoporosis outpatient clinic in northern Italy for initial screening. *Osteoporosis International* **9**, 226–229.

Boonen S, Vanderschueren D, Cheng XG, Verbeke G, Dequeker J, Geusens P, Broos P & Bouillon R (1997) Age-related (type II) femoral neck osteoporosis in men: biochemical evidence for both hypovitaminosis D- and androgen deficiency-induced bone resorption. *Journal of Bone and Mineral Research* **12**, 2119–2126.

Brot C, Vestergaard P, Kolthoff N, Gram J, Hermann AP & Sørensen OH (2001) Vitamin D status and its adequacy in healthy Danish perimenopausal women: relationships to dietary intake, sun exposure and serum parathyroid hormone. *British Journal of Nutrition* **86**, Suppl. 1, S97–S103.

Burnand B, Sloutskis D, Gianoli F, Cornuz J, Rickenbach M, Paccaud F & Burckhardt P (1992) Serum 25-hydroxyvitamin D: distribution and determinants in the Swiss population. *American Journal of Clinical Nutrition* **56**, 537–542.

Carnevale V, Modoni S, Pileri M, Di Giorgio A, Chiodini I, Minisola S, Vieth R & Scillitani A (2001) Longitudinal evaluation of vitamin D status in healthy subjects from southern Italy: seasonal and gender differences. *Osteoporosis International* **12**, 1026–1030.

Chapuy MC, Preziosi P, Maamer M, Arnaud S, Galan P, Hercberg S & Meunier PJ (1997) Prevalence of vitamin D insufficiency in an adult normal population. *Osteoporosis International* **7**, 439–443.

Chapuy MC, Schott AM, Garnero P, Hans D, Delmas PD & Meunier PJ (1996) Healthy elderly French women living at home have secondary hyperparathyroidism and high bone turnover in winter. EPIDOS Study Group. *Journal of Clinical Endocrinology and Metabolism* **81**, 1129–1133.

Corless D, Beer M, Boucher BJ & Gupta SP (1975) Vitamin-D status in long-stay geriatric patients. *Lancet* **i**, 1404–1406.

Danish Food Agency (1996) *Danskernes Kostvaner 1995 (The Danish Dietary Survey 1995)*. Danish Food Agency Publication no. 235. Copenhagen, Denmark: Danish Food Agency.

Davies PSW, Bates CJ, Cole TJ, Prentice A & Clarke PC (1999) Vitamin D: seasonal and regional differences in preschool children in Great Britain. *European Journal of Clinical Nutrition* **53**, 195–198.

Dawson-Hughes B, Harris SS & Dallal GE (1997) Plasma calcidiol, season, and serum parathyroid hormone concentrations in healthy elderly men and women. *American Journal of Clinical Nutrition* **65**, 67–71.

Deharveng G, Charrondiere UR, Slimani N, Southgate DAT & Riboli E (1999) Comparison of nutrients in the food composition tables available in nine European countries participating in EPIC. *European Journal of Clinical Nutrition* **42**, 2449–2453.

de Jong N, Adam SGM, de Groot LCPGM, de Graaf C & Executive Group for Development of Nutrient Dense Foods for Frail Elderly (2000) Variability of micronutrient content in enriched dairy and fruit products. *International Journal of Food Science and Technology* **51**, 247–257.

Deutsche Gesellschaft für Ernährung (2000) *Ernährungsbericht 2000 (Nutrition Report 2000)*. Frankfurt am Mein: Druckerei Heinrich.

- Docio S, Riancho JA, Perez A, Olmos JM, Amado JA & Gonzalez-Macias J (1998) Seasonal deficiency of vitamin D in children: a potential target for osteoporosis-preventing strategies? *Journal of Bone and Mineral Research* **13**, 544–548.
- Dunnigan MG, Fraser SA, McIntosh WB, Moseley H & Sumner DJ (1986) The prevention of vitamin D deficiency in the elderly. *Scottish Medical Journal* **31**, 144–149.
- Egsmose C, Lund B, McNair P, Lund B, Storm T & Sørensen OH (1987) Low serum levels of 25-hydroxyvitamin D and 1,25-dihydroxyvitamin D in institutionalized old people: influence of solar exposure and vitamin D supplementation. *Age and Ageing* **16**, 35–40.
- Fardellone P, Sebert JL, Garabedian M, Bellony R, Maamer M, Agbomson F & Brazier M (1995) Prevalence and biological consequences of vitamin D deficiency in elderly institutionalized subjects. *Revue du Rhumatisme* **62**, 576–581.
- Finch S, Doyle W, Lowe C, Bates CJ, Prentice A, Smithers G & Clarke PC (1998) *National Diet and Nutrition Survey: People Aged 65 Years and Over*. vol. 1, *Report of the Diet and Nutrition Survey*. London: H.M. Stationery Office.
- Glerup H (2000) Vitamin D mangel blandt immigranter (Vitamin D deficiency among immigrants). *Ugeskrift for Læger* **162**, 6196–6199.
- Gloth FM, Gundberg CM, Hollis BW, Haddad JG & Tobin JD (1995) Vitamin D deficiency in homebound elderly persons. *Journal of the American Medical Association* **274**, 1683–1686.
- Guillemant J, Cabrol S, Allemandou A, Peres G & Guillemant S (1995) Vitamin D-dependent seasonal variation in growing male adolescents. *Bone* **17**, 513–516.
- Guillemant J, Taupin P, Le HT, Taright N, Allemandou A, Peres G & Guillemant S (1999) Vitamin D status during puberty in French healthy male adolescents. *Osteoporosis International* **10**, 222–225.
- Heaney RP, Abrams S, Dawson-Hughes B, Looker A, Marcus R, Matkovic V & Weaver C (2000) Peak bone mass. *Osteoporosis International* **11**, 985–1009.
- Heaney RP, Davies KM, Chen TC, Holick MF & Barger-Lux MJ (2003) Human serum 25-hydroxycholecalciferol response to extended oral dosing with cholecalciferol. *American Journal of Clinical Nutrition* **77**, 204–210.
- Hegarty V, Woodhouse P & Khaw KT (1994) Seasonal variation in 25-hydroxyvitamin D and parathyroid hormone concentrations in healthy elderly people. *Age and Ageing* **23**, 478–482.
- Holick MF (1995) Environmental factors that influence the cutaneous production of vitamin D. *American Journal of Clinical Nutrition* **61**, Suppl., 638S–645S.
- Holick MF (1996) Environmental factors that influence the cutaneous production of vitamin D. *Journal of Nutrition* **126**, 1159S–1164S.
- Holick MF (1998) Redefining vitamin D insufficiency. *Lancet* **351**, 805–806.
- Holick MF, MacLaughlin JA, Clark MB, Holick SA, Pots JT, Anderson RR, Blank IH, Åarrish JA & Elias P (1980) Photosynthesis of previtamin D₃ in human skin and the physiologic consequences. *Science* **210**, 203–205.
- Hollis BW (1996) Assessment of vitamin D nutritional and hormonal status: what to measure and how to do it. *Calcified Tissue International* **58**, 4–5.
- Hollis BW (2000) Comparison of commercially available ¹²⁵I-based RIA methods for the determination of circulating 25-hydroxyvitamin D. *Clinical Chemistry* **46**, 1657–1661.
- Hulshof KFAM, Kistemaker C & Bouman M (1998) *De inname van energie en voedingsstoffen door Nederlandse bevolkingsgroepen – Voedselconsumptiepeiling 1997–1998 (The Intake of Energy and Nutrients in the Dutch Population – Food Consumption Survey 1997–1998)*. TNO Report V98-805. Zeist, The Netherlands: TNO.
- Hutchison FN & Bell NH (1992) Osteomalacia and rickets. *Seminars in Nephrology* **12**, 127–145.
- Iqbal SJ, Kaddam I, Wassif W, Nichol F & Walls J (1994) Continuing clinically severe vitamin D deficiency in Asians in the UK (Leicester). *Postgraduate Medical Journal* **70**, 708–714.
- Jesudason D, Need AG, Horowitz M, O'Loughlin PD, Morris HA & Nordin BE (2002) Relationship between serum 25-hydroxyvitamin D and bone resorption markers in vitamin D insufficiency. *Bone* **31**, 626–630.
- Johansson L & Solvoll K (1999) *Norkost 1997, Statens Råd for Ernoring og Fysisk Aktivitet. Rapport no. 2/1999*. Oslo, Norway: Statens Ernæringsråd.
- Juttman JR, Visser TJ, Buurman C, de Kam E & Birkenhager JC (1981) Seasonal fluctuations in serum concentrations of vitamin D metabolites in normal subjects. *British Medical Journal* **282**, 1349–1352.
- Kauppinen-Mäkelin R, Tähtelä R, Löyttyniemi E, Kärkkäinen J & Välimäki MJ (2001) A high prevalence of hypovitaminosis D in Finnish medical in- and outpatients. *Journal of Internal Medicine* **249**, 559–563.
- Keane EM, Healy M, O'Moore RO & Walsh JB (1998) Vitamin D-fortified liquid milk: benefits for the elderly community-based population. *Calcified Tissue International* **62**, 300–302.
- Keane EM, Rochfort A, Cox J, McGovern D, Coakley D & Walsh JB (1992) Vitamin-D-fortified liquid milk – a highly effective method of vitamin D administration for house-bound and institutionalised elderly. *Gerontology* **38**, 280–284.
- Kimlin MG, Downs NJ & Parisi AV (2003) Comparison of human facial UV exposure at high and low latitudes and the potential impact on dermal vitamin D production. *Photochemical and Photobiological Sciences* **2**, 370–375.
- Knudsen VK, Rasmussen LB, Haraldsdottir J, Ovesen L, Bulow I, Knudsen N, Jørgensen T, Laurberg P & Perrild H (2002) Use of dietary supplements in Denmark is associated with health and former smoking. *Public Health Nutrition* **5**, 463–468.
- Krall EA, Sahyoun N, Tannenbaum S, Dallal GE & Dawson-Hughes B (1989) Effect of vitamin D intake on seasonal variations in parathyroid hormone secretion in postmenopausal women. *New England Journal of Medicine* **321**, 1777–1783.
- Kristinsson JÖ, Valdimarsson Ö, Sigurdsson G, Franzson L, Olafsson I & Steingrimsdottir L (1998) Serum 25-hydroxyvitamin D levels and bone mineral density in 16–20 years-old girls: lack of association. *Journal of Internal Medicine* **243**, 381–388.
- Kudlacek S, Schneider B, Peterlik M, Leb G, Klaushofer K, Weber K, Woloszczuk W & Willvonseder R (2003) Assessment of vitamin D and calcium status in healthy adult Austrians. *European Journal of Clinical Investigation* **33**, 323–331.
- Lamberg-Allardt CJ, Outila TA, Karkkainen MU, Rita HJ & Valsta LM (2001) Vitamin D deficiency and bone health in healthy adults in Finland: could this be a concern in other parts of Europe? *Journal of Bone and Mineral Research* **16**, 2066–2073.
- Lehtonen-Veromaa M, Mottonen T, Irjala K, Karkkainen M, Lamberg-Allardt C, Hakola P & Viikari J (1999) Vitamin D intake is low and hypovitaminosis D common in healthy 9- to 15-year-old Finnish girls. *European Journal of Clinical Nutrition* **53**, 746–751.
- Lehtonen-Veromaa MK, Mottonen TT, Nuotio IO, Irjala KM, Leino AE & Viikari JS (2002) Vitamin D and attainment of peak bone mass among peripubertal Finnish girls: a 3-y prospective study. *American Journal of Clinical Nutrition* **76**, 1446–1453.
- Lips P (2001) Vitamin D deficiency and secondary hyperparathyroidism in the elderly: consequences for bone loss and fractures and therapeutic implications. *Endocrine Reviews* **22**, 477–501.
- Lips P, Bouillon R, Jongen MJ, van Ginkel FC, van der Vijgh WJ & Netelenbos JC (1985) The effect of trauma on serum

- concentrations of vitamin D metabolites in patients with hip fracture. *Bone* **6**, 63–67.
- Lips P, Chapuy MC, Dawson-Hughes B, Pols HAP & Holick MF (1999) An international comparison of serum 25-hydroxyvitamin D measurements. *Osteoporosis International* **9**, 394–397.
- Lips P, van Ginkel FC, Jongen MJ, Rubertus F, van der Vijgh WJ & Netelenbos JC (1987) Determinants of vitamin D status in patients with hip fracture and in elderly control subjects. *American Journal of Clinical Nutrition* **46**, 1005–1010.
- McKenna MJ (1992) Differences in vitamin D status between countries in young adults and the elderly. *American Journal of Medicine* **93**, 69–77.
- McKenna MJ & Freaney R (1998) Secondary hyperparathyroidism in the elderly: means to defining hypovitaminosis D. *Osteoporosis International* **8**, Suppl., S3–S6.
- McKenna MJ, Freaney R, Meade A & Muldowney FP (1985) Hypovitaminosis D and elevated serum alkaline phosphatase in elderly Irish people. *American Journal of Clinical Nutrition* **41**, 101–109.
- MacLaughlin J & Holick MF (1985) Aging decreases the capacity of human skin to produce vitamin D₃. *Journal of Clinical Investigation* **76**, 1536–1538.
- Malabanan A, Veronikis IE & Holick MF (1988) Redefining vitamin D insufficiency. *Lancet* **351**, 805–806.
- Melin AL, Wilske J, Ringertz H & Saaf M (1999) Vitamin D status, parathyroid function and femoral bone density in an elderly Swedish population living at home. *Ageing* **11**, 200–207.
- Meller Y, Kestenbaum RS, Shany S, Galinsky D, Zuili I, Yankovitch N, Giat J, Conforti A & Torok G (1985) Parathormone, calcitonin, and vitamin D metabolites during normal fracture healing in geriatric patients. *Clinical Orthopaedics* **199**, 272–279.
- Meulmeester JF, van den Berg H, Wedel M, Boshuis PG, Hulshof KF & Luyken R (1990) Vitamin D status, parathyroid hormone and sunlight in Turkish, Moroccan and Caucasian children in The Netherlands. *European Journal of Clinical Nutrition* **44**, 461–470.
- Morris HA, Morrison GW, Burr M, Thomas DW & Nordin BE (1984) Vitamin D and femoral neck fractures in elderly South Australian women. *Medical Journal of Australia* **140**, 519–521.
- Mowé M, Böhmer T & Haug E (1996) Serum calcidiol and calcitriol concentrations in elderly people: variations with age, sex, season and disease. *Clinical Nutrition* **15**, 201–206.
- Nayal AS, MacLennan WJ, Hamilton JC, Rose P & Kong M (1978) 25-Hydroxyvitamin D, diet and sunlight exposure in patients admitted to a geriatric unit. *Gerontology* **24**, 117–122.
- Nordic Council of Ministers (1995) *Addition of Nutrients to Food – Principles and Practices*. TemaNord, p. 643. Oslo, Norway: The Nordic Council of Ministers.
- Ooms ME, Lips P, Roos JC, van der Vijgh WJF, Popp-Snijders C, Bezemer PD & Bouter LM (1995a) Vitamin D status and sex hormone binding globulin: determinants of bone turnover and bone mineral density in elderly women. *Journal of Bone and Mineral Research* **10**, 1177–1184.
- Ooms ME, Roos JC, Bezemer PD, van der Vijgh WJF, Bouter LM & Lips P (1995b) Prevention of bone loss by vitamin D supplementation in elderly women: a randomised double blind trial. *Journal of Clinical Endocrinology and Metabolism* **80**, 1052–1058.
- Outila TA, Karkkainen MU & Lamberg-Allardt CJ (2001) Vitamin D status affects serum parathyroid hormone concentrations during winter in female adolescents: associations with forearm bone mineral density. *American Journal of Clinical Nutrition* **74**, 206–210.
- Ovesen L, Brot C & Jakobsen J (2003) Food contents and biological activity of 25-hydroxyvitamin D: a vitamin D metabolite to be reckoned with? *Annals of Nutrition and Metabolism* **47**, 107–113.
- Parfitt AM (1998). Osteomalacia and related disorders. In *Metabolic Bone Disease and Clinically Related Disorders*, pp. 327–386 [LV Avioli and SM Krane, editors]. San Diego, CA: Academic Press.
- Parfitt AM, Gallagher JC, Heaney RP, Johnston CC, Neer R & Whedon GD (1982) Vitamin D and bone health in the elderly. *American Journal of Clinical Nutrition* **36**, 1014–1031.
- Prentice A (2002) What are dietary requirements for calcium and vitamin D? *Calcified Tissue International* **70**, 83–88.
- Quesada JM, Jans I, Benito P, Jimenez A & Boillon R (1989) Vitamin D status of elderly people in Spain. *Age and Ageing* **18**, 392–397.
- Reginster JY, Frederick I, Deroisy R, Dewe W, Taquet AN, Albert A, Collette J, Pirenne H, Zheng SX & Gosset C (1998) Parathyroid hormone plasma concentrations in response to low 25-OH vitamin D circulating levels increases with age in elderly women. *Osteoporosis International* **8**, 390–392.
- Sahota O, Masud T, San P & Hosking DJ (1999) Vitamin D insufficiency increases bone turnover markers and enhances bone loss at the hip in patients with established vertebral osteoporosis. *Clinical Endocrinology* **51**, 217–221.
- Scharla SH, Scheidt-Nave C, Leidig G, Woitige H, Wuster C, Seibel MJ & Ziegler R (1996) Lower serum 25-hydroxyvitamin D is associated with increased bone resorption markers and lower bone density at the proximal femur in normal females: a population-based study. *Experimental and Clinical Endocrinology and Diabetes* **104**, 289–292.
- Scragg R, Khaw K-T & Murphy S (1995) Life-style factors associated with winter serum 25-hydroxyvitamin D levels in elderly adults. *Age and Ageing* **24**, 271–275.
- Sem SW, Sjoen RJ, Trygg K & Pedersen JI (1987) Vitamin D status of two groups of elderly in Oslo: living in old people's homes and living in own homes. *Comprehensive Gerontology* **1A**, 126–130.
- Shepard RM & DeLuca HF (1980) Plasma concentrations of vitamin D₃ and its metabolites in the rat as influenced by vitamin D₃ or 25-hydroxyvitamin D₃ intakes. *Archives of Biochemistry and Biophysics* **202**, 43–53.
- Sigurdsson G, Franzson L, Steingrimsdottir L & Sigvaldason H (2000) The association between parathyroid hormone, vitamin D and bone mineral density in 70-year-old Icelandic women. *Osteoporosis International* **11**, 1031–1035.
- Solanki T, Hyatt RH, Kemm JR, Hughes EA & Cowan RA (1995) Are elderly Asians in Britain at a high risk of vitamin D deficiency and osteomalacia? *Age and Ageing* **24**, 103–107.
- Souberbielle JC, Cormier C, Kindermans C, Gao P, Cantor T, Forette F & Baulieu EE (2001) Vitamin D status and redefining serum parathyroid hormone reference range in the elderly. *Journal of Clinical Endocrinology and Metabolism* **86**, 3086–3090.
- Thomas MK, Lloyd-Jones DM, Thadhani RI, Shaw AC, Deraska DJ, Kitch BT, Vamvakas EC, Dick IM, Prince RL & Finkelstein JS (1998) Hypovitaminosis D in medical inpatients. *New England Journal of Medicine* **338**, 777–783.
- Tjellessen L & Christiansen C (1983) Vitamin D metabolites in normal subjects during one year. A longitudinal study. *Scandinavian Journal of Clinical Laboratory and Investigation* **43**, 85–89.
- Toss G, Almqvist S, Larsson L & Zetterqvist H (1980) Vitamin D deficiency in welfare institutions for the aged. *Acta Medica Scandinavica* **208**, 87–89.
- van der Wielen RPJ, Löwik MRH, van den Berg H, de Groot LCPGM, Haller J, Moreiras O & van Staveren WA (1995) Serum vitamin D concentrations among elderly people in Europe. *Lancet* **346**, 207–210.
- Vieth R, Chan P-CR & MacFarlane GD (2001) Efficacy and safety of vitamin D₃ intake exceeding the lowest observed adverse effect level. *American Journal of Clinical Nutrition* **73**, 288–294.

- Vir SC & Love AHG (1978) Vitamin D status of elderly at home and institutionalised in hospital. *International Journal of Vitamin and Nutrition Research* **48**, 123–130.
- Volatier J-L (2000), *Enquête Individuelle et Nationale sur les consommations Alimentaires (Individual and National Study of Food Consumption)*. Paris: Editions TEC et DOC Lavoisier.
- von Knorring J, Slatis P, Weber TH & Helenius T (1982) Serum levels of 25-hydroxyvitamin D, 24,25-dihydroxyvitamin D and parathyroid hormone in patients with femoral neck fracture in southern Finland. *Clinical Endocrinology* **17**, 189–194.
- Webb AR, Kline L & Holick MF (1988) Influence of season and latitude on the cutaneous synthesis of vitamin D₃: exposure to winter sunlight in Boston and Edmonton will not promote vitamin D₃ synthesis in human skin. *Journal of Clinical Endocrinology and Metabolism* **67**, 373–378.
- Woitge HW, Scheidt-Nave C, Kissling C, Leidig-Bruckner G, Meyer K, Grauer A, Scharla SH, Ziegler R & Seibel MJ (1998) Seasonal variation of biochemical indexes of bone turnover: results of a population-based study. *Clinical Endocrinology and Metabolism* **83**, 68–75.