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Quality and use of unlicensed vitamin D preparations in primary care in England: retrospective review of national prescription data and laboratory analysis
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What is already known about this subject

- The supply of unlicensed vitamin D preparations (e.g. food supplements) against medical vitamin D prescriptions was common practice in England in 2014.
- Little is known about the quality difference between licensed and unlicensed vitamin D preparations marketed in England.

What this study adds

- The supply of licensed vitamin D preparations against prescriptions has increased over time.
- While the number of licensed vitamin D preparations has increased, 40% of all vitamin D prescriptions dispensed in 2018 were still of unlicensed preparations.
- Vitamin D content of unlicensed vitamin D preparations vary widely from labelled claim.

Abstract

**Aim** To evaluate the type (licensed vs. unlicensed) and cost of preparations used to fulfil vitamin D prescriptions in England over time, and to compare measured vitamin D content of selected vitamin D preparations against labelled claim.


**Results** Alongside a rise in the number of oral licensed colecalciferol preparations from 0 to 27 between 2012 and 2018, the proportion of vitamin D prescriptions in which licensed vitamin D preparations were supplied increased from 11.8 to 54.2%. However, the use of unlicensed food supplements (dose strength: 400-50,000 IU) remained high accounting for 39.7% of vitamin D prescriptions in 2018. The two licensed preparations showed mean (±
standard deviation) vitamin D content of 90.9 ± 0.7% and 90.5 ± 3.9% of the labelled claimed amount, meeting the British Pharmacopeia specification for licensed medicines (90-125% of labelled claim). The 11 food supplements showed vitamin D content ranging from 41.2 ± 10.6% to 165.3 ± 17.8% of the labelled claim, with 8 of the preparations failing to comply with the food supplement specification (80-150% of labelled claim).

Conclusions Despite the increasing availability of quality assured licensed preparations, food supplements continued to be used interchangeably with licensed preparations to fulfil vitamin D prescriptions. Food supplements, manufactured under less stringent quality standards, showed wide variations between measured and declared vitamin D content, which could lead to the risk of under- and over-dosing.

Introduction Vitamin D has attracted considerable interest in recent years, not least because of increased awareness of global prevalence of vitamin D deficiency.[1] Since 2016, the advice from the UK government is that individuals should consider taking daily vitamin D supplement of 400 international units (IU) during the winter months (from October to March), and all year round for anyone who spend limited time outdoors.[2] Vitamin D has also received wide media coverage arising from research linking vitamin D deficiency to an increased risk of a myriad of adverse health problems including diabetes, autoimmune conditions, cancers and infections, among others.[3] This heightened interest has, in turn, been accompanied by a significant rise in healthcare and consumer spending on vitamin D preparations. In England, medical prescriptions for vitamin D monotherapy preparations in primary care increased by more than 8,000% between 2008 and 2013, with the latest annual spending reported at £38 million.[4, 5] Likewise, consumer demand for vitamin D supplements has soared; 2018 retail data from the United Kingdom (UK) reported that vitamin D was the most popular vitamin monotherapy in this multi-million-pound industry.[6] These figures are indicative of a growing trend and reflect widespread interest, although not substantiated by robust clinical trial data, in the role of vitamin D in health.[3]

Vitamin D is produced by cutaneous synthesis, with very small amounts obtained from the diet, unless foods are fortified with vitamin D.[7, 8] Individuals who want to improve their dietary intake of vitamin D can do so through supplements. Vitamin D preparations with the intended purpose of supplementing dietary sources would generally fall within the legal
definition of food supplements and are regulated under the provisions of general food law.[9, 10] However, vitamin D is also used clinically to treat or prevent vitamin D deficiency, and products making a medicinal claim would be subjected to the legislation governing medicinal products.[11–13] These legal distinctions have consequently led to the availability of two types of vitamin D preparations which are regulated by two different quality standards. Notably, the regulatory requirements for manufacturing, quality control, and labelling which apply to medicinal products are far more stringent than those relating to food supplements. While medicinal products are subjected to pre-market licensing assessment and hence are referred as licensed medicinal products, food supplements are outside of the licensing process (referred as unlicensed preparations), and are readily available to the general public without medical supervision through retail outlets including supermarkets, pharmacies, health food stores, and online retailers.

Deciding whether a product is a medicinal product or a food supplement is dependent on a number of factors and the amount of an ingredient is not the only criterion.[11] In practice, the classification is sometimes ambiguous in that two apparently similar preparations are classified differently, but to the consumer can appear to be virtually indistinguishable. The extent to which these situations arise with vitamin D preparations is not known and is further complicated by concerns about product quality with reports of poor quality food grade vitamin D preparations on sale in New Zealand, United State and India.[14–16] Questions over safety and legal issues associated with the use of food supplements to fulfil medical prescriptions of vitamin D in England has also been highlighted.[5] However, neither the quality of food grade vitamin D preparations, nor the trends in supply of these preparations, have been investigated in England to-date. Therefore, the aims of this work were to evaluate the type-and costs of preparations used to fulfil vitamin D prescriptions in primary care in England over time, and to compare measured vitamin D content of selected vitamin D preparations marketed in England against their labelled values.

Methods

Licence status of vitamin D preparations

The National Health Service (NHS) dictionary of medicines and devices (dm+d) via the NHS Technology Reference Data Update Distribution (TRUD) platform was used to identify all vitamin D preparations in use across primary care in England;[17][16] both ergocalciferol (vitamin D2) and colecalciferol (vitamin D3) preparations were included. Combination
preparations (e.g. multivitamins, colecalciferol and calcium) were excluded in this study. For each preparation, we extracted data on its availability, dose strength, dosage form, and type of vitamin D preparation, categorised as either licensed prescription only medicines, licensed pharmacy medicines, licensed general sale list medicines, food supplements, “specials”, “imports” and “undetermined” (the legal definitions of these categories are provided in supplementary method 1).

**Prescription data**

All vitamin D prescriptions dispensed in primary care in England between 2008 and 2018 were extracted from “Prescription Cost Analysis” (PCA), a publicly available national dataset of prescription reimbursement data.[4] The PCA dataset complies with all aspects of the Code of Practice for Official Statistics and prescription processing activity has been internally audited to 99% accuracy. For each vitamin D presentation listed (further details on data processing of PCA dataset are provided in supplementary method 2), we extracted data on the number of prescription item, unit quantity (i.e. tablets or millilitres), net ingredient cost (NIC), and NIC per unit quantity (i.e. cost per tablet).[4, 17, 18] The NIC is the basic cost of the preparation at list price, excluding Value Added Tax. Due to commercial confidentiality, contract prices are not released outside of the NHS.

**Vitamin D content analysis**

As ergocalciferol prescriptions accounted for less than 1% of the annual total vitamin D prescriptions since 2015, only colecalciferol preparations were included in the laboratory analysis. Eleven colecalciferol food supplements, covering a range of dose strengths and different dosage forms, were purchased from 5 different UK pharmacies, health food stores, and online retailers; this represents approximately 6% of the vitamin D food supplements listed in the NHS dm+d. The preparations were selected based on a convenience sample reflecting food supplements that were available for retail purchase and used by the general public. We also included two licensed preparations that are routinely used in the NHS; one UK licensed preparation and one imported licensed preparation from Germany. The UK and Germany operate to the same manufacturing quality standards with regards to licensed medicinal products. All preparations were stored at controlled room temperature in cardboard boxes and protected from light following purchase. All products were within their expiry date at the time of analysis.
A reverse phase high performance liquid chromatography (RP-HPLC) assay was developed to quantify colecalciferol content. All analytical work was carried out at King’s College London (A. Patel and S. Jones). The assay was verified as ‘fit-for-purpose’ against the criteria specified in the International Conference of Harmonisation (ICH) harmonised tripartite guideline on validation of analytical procedures.[19] Details of the analytical development of the RP-HPLC method are provided in supplementary method 3.

Chromatographic separation was achieved on a C18 Synergi Hydro-RP column (250 x 4.6 mm, 4μm, Phenomenex, USA) using an injection volume of 25 µL. The mobile phase comprised acetonitrile and methanol (55:45, %v/v), which was delivered as a flow rate of 1.5 mL/min with the column temperature held at room temperature. The ultra-violet detection was assessed at a wavelength of 265 nm. Data acquisition and processing were performed using ChromNAV software (Jasco, Tokyo, Japan).

Three individual samples from the 13 colecalciferol preparations were subject to extraction and HPLC analysis. Each of the 3 samples was injected twice onto the HPLC and an average of the 6 results was reported. Details of the methods used to extract colecalciferol from the preparations are provided in supplementary method 3. The liquid preparations were vortexed for 2 minutes and the solid preparations were sonicated for 45 minutes to ensure the extraction solutions were homogeneous. To account for losses during extraction, ergocalciferol was used an internal standard to improve the accuracy in the quantification of colecalciferol.

**Data analysis**

Characteristics of vitamin D preparations used across primary care in England were summarised using descriptive statistics. For prescription trend analysis, vitamin D prescriptions dispensed each year were calculated and presented as percentage of total prescription items and total NIC by type of vitamin D preparation. Using 2018 data, the NIC reported for each unlicensed preparation (food supplement, “specials”, or “imports”) was compared with the estimated NIC that would have been spent had licensed preparations been supplied instead. To do this, all vitamin D licensed preparations were categorised into mutually exclusive categories based on dosage form (e.g. tablets, capsules, solution) and dose strength. A weighted NIC per unit quantity (i.e. £ per tablet or £ per millilitre) was calculated for each category based on the NIC per unit quantity and the relative number of items with which each preparation was supplied within that category. Subsequently, the NIC had
licensed medicinal grade preparations been supplied was determined by substituting the NIC per unit quantity for each of the unlicensed preparations with the weighted NIC per unit quantity of the licensed equivalent with respect to dosage form and dose strength. In cases where there were no direct licensed equivalent substitutes, therapeutically equivalent substitutes (e.g. 2,000 IU tablet substitutes for 2,200 IU tablet) as reviewed and agreed by a clinician and a pharmacist were employed. In the analysis of prescription data, ergocalciferol and colecalciferol were also considered therapeutically equivalent.

For RP-HPLC analyses, peak areas were used for quantification of colecalciferol concentrations. The mean percentage difference from labelled content and standard deviation (SD) were calculated for all 13 preparations. An acceptable criterion of 90 to 125% of the stated amount was set in line with the British Pharmacopoeia, the official UK pharmaceutical standards for medicinal products.[20] We also compared our results against food supplement standard which has a wider acceptance range of 80 to 150% of the stated label amount.[21] The relationship between preparation dose strength and the magnitude of percentage difference between measured and labelled contents was assessed with Spearman’s correlation coefficient.

Data analysis was conducted using Stata 15 (Stata Corp, College Station, Texas).

**Ethical approval**
As the study used routinely collected data that was non-identifiable and publicly available, ethics approval was not required.

**Results**

**Characteristics of vitamin D preparations**
As of 9th September 2019, there was a total of 328 oral vitamin D preparations listed as available in the NHS dm+d; 35 (10.7%) were categorised as licensed prescription only medicines, 186 (56.7%) as food supplements, 102 (31.1%) as “specials” and 5 (1.5%) as “imports”. A total of 9 injectable formulations were also listed with 3 preparations each under the 3 categories of ‘licensed’, ‘specials’ and ‘imports’. The oral licensed medicinal preparations were available in a range of dose strengths (Table 1); tablets and capsules were available from 400 IU to 50,000 IU, and liquid preparations from 2,400 IU/mL to 50,000 IU/mL.
Food supplements, presented as tablets or capsules, spanned the same range of dose strengths (400 IU to 50,000 IU) as the licensed medicinal preparations (Table 1), but the liquid formulations were available in a narrower range from 400 IU/mL to 10,000 IU/mL. Oral preparations categorised as “specials” covered the broadest range of dose strengths of all the preparations from 20 IU/mL to 600,000 IU/mL.

**Trends in vitamin D prescriptions**

During the study period, the number of different preparations used to fulfil vitamin D prescriptions increased from 42 to 268; the increase was attributed to an increase in both the use of colecalciferol food supplements and licensed oral medicinal grade colecalciferol preparations (0 to 27) (Figure 1A).

Between 2008 and 2018, the percentage increase in vitamin D prescriptions was 29,842%, (23,543 to 7,049,322) in primary care in England. Ergocalciferol prescriptions accounted for 96.8% of the total vitamin D prescriptions in 2008, decreasing to 5.6% by 2012 and 0.2% in 2018 (Figure 1B). Colecalciferol prescriptions showed a reciprocal trend to ergocalciferol.

Analysis by the type of preparation showed that the supply of licensed colecalciferol preparations increased from 8.7% in 2012 to 54.1% in 2018 (Figure 1B). The proportion of prescriptions using colecalciferol food supplements was 61.3% in 2012, and the proportion remained high at 39.7% in 2018. The overall use of “specials” preparations has decreased over time and remained at less than 0.6% since 2015.

**Vitamin D preparation quality**

The two licensed medicinal grade preparations had a mean (±standard deviation) percentage vitamin D content compared to the label claim of 90.91 ± 0.68 % and 90.53 ± 3.93 %, respectively, which correspond to the quality standard of medicinal products (90-125%). The vitamin D content of food grade preparations varied from 41.2 ± 10.6% to 165.3 ± 17.8% of the labelled value (Figure 2). A total of 8 of the 11 food grade preparations tested failed to comply with the food supplement standard of 80-150% of the labelled claim. There was no association between the dose strength of the preparation and the magnitude of percentage difference from labelled values ($P >0.5$). Vitamin D food supplements presented as solution in dropper bottles ($n = 3$) showed high inter-sample variability, with SD of 42, 28.1 and 17.3.
Cost comparison between licensed and unlicensed vitamin D preparations

In 2018, the total cost of vitamin D prescriptions was £38.4 million (Figure 3). Of that, £18.6 million was for tablets/capsules containing ≤800 IU of vitamin D, £15.9 million for tablets/capsules containing >800 IU of vitamin D, £3.7 million for oral solution, and £0.1 million for injections.

A licensed equivalent preparation was identified for all 236 unlicensed preparations (food supplements, “ specials”, and “imports”). The median cost difference between annual spending on an unlicensed preparation and the estimated cost that would have been spent for its licensed equivalent was a saving of £347, with wide variations in cost from a saving of £6,980 to an additional cost of £313 (interquartile range).

If all unlicensed preparations as supplied in 2018 were to be replaced with licensed vitamin D preparations, the annual cost would have totalled £43.5 million, an increase of 13.4% higher than current cost. The increase in estimated cost was largely attributed to two food supplements, that together accounted for 9% of the total vitamin D prescriptions in 2018 and whose unit costs were considerably lower even compared to other food supplements of the same strength (priced at £0.01 vs median cost of £0.1 per 1,000 IU tablet; £0.13 vs median cost of £0.28 per 20,000 IU capsule, respectively).

Discussion

Using national prescribing data, this study showed that patients presenting with a vitamin D prescription in 2018 were almost as likely to be supplied with a food supplement as a licensed medicinal product despite the availability of licensed preparations of equivalent strength. This suggests that food supplements and licensed preparations, both available in a wide range of dose strengths from 400 to 50,000 IU, were being used interchangeably. However, food supplements vary in vitamin D content by up to 60% compared to their labelled claim, with a potential risk of under- or over-dosing the patient. Existing regulations for vitamin D preparations need to be revised in order to minimise risk and support consumers in making informed decisions on the use of vitamin D preparations.

An earlier review highlighted safety and legal concerns associated with a high proportion of vitamin D prescriptions in primary care in England being supplied with food grade vitamin D preparations.[5] Arguably, in 2013, when there were only 6 licensed oral vitamin D
preparations on the market, it may have been too soon to draw firm conclusions about the appropriateness of such practice, but the current study, 5 years on, provides evidence that the initial concerns were valid. Not only has the present study shown that food supplements continue to be used in 39.7% of vitamin D prescriptions in primary care in England despite the availability of licensed preparations of equivalent strength, but these food supplements may be of substandard quality and are not necessarily lower in cost compared to licensed preparations.

The substantial difference between measured and declared vitamin D content in food supplements shown in our study is in keeping with published data from other countries.[14–16, 22, 23] Compared with variations reported by LeBlanc et al. for preparations marketed in the United State (9-140%) and Garg et al. (8-201%) for New Zealand, our results showed a narrower range of variability.[14, 15] This difference could be a reflection of the differences in legislative requirements between countries, but as 8 of the 11 food supplements tested in the present study failed to comply with national food quality standards, this seems unlikely.[21] Rather, the variability in vitamin D content in food supplements probably arise from differences in manufacturing and quality control practices between manufacturers which in turn may reflect the general lack of regulatory oversight of food supplement manufacturers in these countries.[10] Notably, our result showing variations in measured vitamin D contents outside of the acceptable range of the declared values suggests that food supplements manufacturers are likely to be operating with variable and less stringent control in handling product deviations, as well as pointing to challenges faced by regulator in enforcing quality standards in this industry.[10, 24] The general perception that all food supplements are regarded as low risk may also be a contributing factor.

Our findings have implications in clinical practice. From a public health perspective, data generated from dietary intake assessments, which in turn may be used to provide the evidence base for developing public health policy on preventing vitamin D deficiency, could potentially be misleading if estimations of dietary intake rely entirely on label information for vitamin D food supplements. For individuals, our findings would suggest that some may receive up to 60% more or less vitamin D than intended. The clinical significance of this magnitude of variation for low strength preparations may be of little clinical concern given vitamin D has a wide therapeutic window, although it is possible that high-risk individuals may not receive sufficient vitamin D to correct risk of their vitamin D deficiency. For high
strength preparations, there is a risk of inducing vitamin D toxicity, with an associated risk of hypercalciuria and renal damage, particularly in at-risk populations. The rise in the number of reports of vitamin D toxicity, which have been attributed to errantly manufactured food grade preparations, especially in individuals receiving high doses of vitamin D over a long period of time without medical supervision, should be of concern to both clinicians and consumers.[25, 26] Moreover, the mismatch between intended and actual dose received could mislead clinicians in suspecting other diagnoses and carrying out unnecessary investigations.

The findings also call for clinicians to question the continued use of food supplements to fulfil their vitamin D prescriptions. In England, the use of unlicensed preparations is legally permitted when there is no suitable licensed alternatives.[27] Therefore if generic prescriptions are being written by clinicians then it is difficult to understand why almost half of the time unlicensed food supplements are being supplied. The British National Formulary,[28] a reference book that contains highly authoritative information on medicines prescribing in the UK, lists both medicinal and food grade vitamin D preparations, which may inadvertently be misinterpreted by clinicians to mean that the preparations can be used interchangeably. However, without evidence that a particular brand is superior to another and along with 81% of primary care prescriptions in England prescribed generically,[29] it is unlikely that clinicians are deliberately choosing food grade preparations. An alternative explanation is that clinicians are not aware that food supplements are being used by pharmacists to fulfil their generic vitamin D prescriptions. It is possible that pharmacists are intentionally opting to use food supplements as a means to reduce NHS spending; after all, if only licensed preparations were used, this study predicted that the cost of vitamin D prescriptions in 2018 would have been 13.4% higher (£5 million) compared to the current cost.

The regulatory landscape for vitamin D preparations in England does not appear to help clinicians, pharmacists, patients or consumers to distinguish the differences in vitamin D product quality and intended use. There appears to be a regulatory anomaly for vitamin D preparations in that whilst all licensed vitamin D medicinal preparations of dose strength from 400 IU to 50,000 IU are classified by the drug regulator as products requiring medical prescription, i.e. as described in law as “likely to present a danger either directly or indirectly, even when used correctly, if utilized without medical supervision”, equivalent strengths food supplements are readily available on the market for general sale.[30] Interestingly, there is no
distinction between medicinal and food grade vitamin D preparations in terms of their vitamin D content in England, which contrasts countries such as Canada and Australia where all preparations containing more than 1,000 IU of vitamin D are regulated as licensed medicinal products.[12, 13] Similarly, countries including Belgium, Norway and others, have set statutory maximum limits for the amount of vitamin D permitted in food supplements.[31, 32] It is also of interest to note that food supplements, manufactured under less stringent quality standards in England, may have much lower or higher vitamin D contents than equivalent strengths of medicinal preparations due to having a wider tolerance range around the declared value on the label.

Our data show that vitamin D food supplements on general sale are available in a wide range of dose strengths of up to 50,000 IU. Considering the recommended nutrient intake (RNI) in England for most people is only 400 IU of vitamin D per day, and even after taking into account the higher doses recommended by some organisations (up to 2,000 IU per day) and the recommended tolerable upper intake level of 4,000 IU/day,[2, 33, 34] the availability of these high strength food supplements would detract from the public health message. Indeed, these high strength food supplements are not intended for daily administration, but preparations labelled as 50,000 IU weekly, equating to 18 times above the RNI, are freely available for purchase by consumers and can be easily misused, e.g. taken daily, without medical supervision. Such preparations expose consumers to unnecessary risks without evidence of benefit.[2]

From a clinical perspective, it would therefore seem sensible to regulate high strength vitamin D preparations as medicinal products on the UK market. This would ensure that all prescriptions for the clinical treatment of vitamin D deficiency would be supplied with licensed medicinal grade preparations, but at the same time not unnecessarily restrictive to ensure consumers have appropriate access to preparations at recommended supplement doses to improve their dietary intake of vitamin D. In light of the UK government’s recent recommendation to limit vitamin D prescribing to therapeutic treatment of vitamin D deficiency only, the £18.6 million saved in costs (based on 2018 figures for vitamin D tablets or capsules preparations containing ≤800 IU alone) would outweigh the additional estimated cost associated with using only licensed vitamin D preparations to fulfil vitamin D prescriptions.[4, 35] This would allow clinicians, pharmacists, and patients to be assured that
the dose of vitamin D being received by patients requiring treatment for vitamin D deficiency is as intended.

The problem with regulatory oversight of food supplements has been raised for many years, but the supply of food supplements against medical prescriptions when licensed equivalents are available, as shown in this study, further press for improved oversight of vitamin D preparations. Moreover, the prescribing trend for vitamin D preparations may reflect similar prescribing patterns of other supplements, and the implication of these on the clinical management of patients would deserve similar attention.

**Strengths and limitations of the study**

We used the complete prescription reimbursement data set for all vitamin D prescriptions dispensed in primary care in England, not just a sample. Moreover, as our data set was based on prescription reimbursement and thus data interpretation was not affected by generic prescribing, we were able to provide a comprehensive analysis of the different vitamin D preparations used, providing real-world evidence of practice. We note that our analysis was based on data in England, but we expect similar prescription patterns across the UK given the similarities in primary care practice for prescribing and dispensing prescriptions.

We evaluated the vitamin D content of only a small sample of vitamin D preparations which may not be representative of all the preparations marketed in England and further comprehensive analysis is warranted. Nevertheless, our results obtained were consistent with published literature. Our RP-HPLC method was verified as ‘fit-for-purpose’ against the ICH standards in terms of accuracy, precision and linearity and was used rather than the method stated in the British Pharmacopeia due to its ability to assess both the vitamin D content in the food and medicinal grade preparations. We did not investigate the reasons for the variability in vitamin D contents using stability indicating methods because we were unsure of the grade of vitamin D originally employed in the food grade preparations. We also did not investigate batch-to-batch variability in colecalciferol content of the different preparations and the number of samples per preparation tested was less than that recommended by the British Pharmacopeia. However, our study conclusion would remain valid in the case of minimum variability while the presence of batch-to-batch/sample-to-sample variability would further highlight the problem of substandard quality with these food supplement preparations.
Conclusions
The study showed that whilst food and medicinal grade vitamin D preparations were being used interchangeably to fulfil primary care vitamin D prescriptions, food grade preparations, which are manufactured under less stringent quality standards, seldom contained the expected amount of vitamin D. The work provided evidence that the number and variety of licensed medicinal grade vitamin D preparations was sufficient to enable the supply of only licensed medicinal grade preparations against all vitamin D prescriptions. Clinicians and pharmacists must therefore change practice to use licensed vitamin D preparations whenever possible. We call on regulatory bodies to safeguard public health and ensure that only medicinal grade vitamin D preparations are used to treat vitamin D deficiency.

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Contributors
MW, RS and JP were involved in the conception of the study. AP under the supervision of SJ carried out the laboratory analysis. MW had full access to the prescription data set and performed the data extraction and analysis. MW, AP, JP, GR, SJ and RS contributed to the interpretation of data. MW drafted the manuscript which all authors contributed to and revised critically before approval of the final manuscript. MW is the guarantor. The corresponding author attests that all listed authors meet authorship criteria and that no others meeting the criteria have been omitted.
Competing interests
All authors have completed the ICMJE uniform disclosure form at www.icmje.org/coi_disclosure.pdf and declare: no support from any organisation for the submitted work; no financial relationships with any organisations that might have an interest in the submitted worked in the previous three years; no other relationships or activities that could appear to have influenced the submitted work.

Data sharing

Openprescribing.net is also a free, openly accessible tool available at https://OpenPrescribing.net.

References


Figure 1A: Number of vitamin D preparations used to fulfil vitamin D prescriptions in primary care in England between 2008 and 2018 by type of vitamin D preparation. 1B: Percentage of vitamin D prescriptions dispensed in primary care in England between 2008 and 2018 by type of vitamin D preparation.
Figure 2: Percentage difference between measured colecalciferol content and labelled value for various colecalciferol preparations on the UK market. Red * represents licensed medicinal products; L = Liquid preparation; C = soft gelatine capsule; T = Tablet; Grey dotted lines represent the acceptable criterion of 90-125% as required of licensed medicines.
Figure 3: Net ingredient costs of vitamin D prescriptions dispensed in primary care in England between 2008 and 2018 by type of vitamin D preparation.
Table 1: Dose strengths of oral vitamin D preparations listed as available in the National Health Service (NHS) dictionary of medicines and devices (dm+d) (as of 9th September 2019)

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<th>Licensed preparations n = 35</th>
<th>Food supplements n = 186</th>
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