ORIGINAL RESEARCH

Factors associated with vitamin D status of lowincome, hospitalized psychiatric patients: results of a retrospective study

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Background: Recent evidence has indicated a potential role of vitamin D3 in a range of neuropsychiatric outcomes, as well as on cognitive function, but conflicting data have left that role uncertain. Understanding potential associations of vitamin D status with psychiatric illness will allow clinicians to better assess therapeutic options. Few studies have examined vitamin D status among a racially diverse group of psychiatric patients who have been hospitalized, and none has done so in the southern US where socioeconomic inequality is high.

Methods: In this retrospective study, medical records from 113 patients hospitalized for psychiatric illness were retrieved and analyzed. Vitamin D status in this population was estimated, along with any patterns of association between deficiency and risk factors.

Results: The vast majority of patients hospitalized for psychiatric illness in this biracial, lowincome sample had either insufficient or deficient vitamin D levels. African-American patients had lower levels of vitamin D than Caucasian patients.

Discussion: Our findings demonstrate that hospitalized psychiatric patients are at increased risk for vitamin D deficiency and in particular low-income, African-American populations. These results suggest that vitamin D should be assessed and therapy considered at the initiation of psychiatric hospitalizations.

Keywords: vitamin D, psychiatric diagnosis, cognitive function, neuropsychological function

Introduction

The association between low vitamin D3 serum levels and neuropsychiatric conditions has been the subject of a number of recent research studies. Technically a steroid hormone, vitamin D3 and its receptor VDR (vitamin D receptor) have become more widely implicated in various health and disease states beyond osteomalacia, or rickets, as information on their function in the body has emerged over the past three decades.¹ Deficiency of vitamin D3 (also known as cholecalciferol) has arisen as a possible risk factor for a range of mental health conditions.^{2,3} Preclinical studies and epidemiological research support the contention that prenatal deficiency of vitamin D impairs fetal neurodevelopment potentially contributing to developmental disability in childhood or psychiatric conditions such as schizophrenia in adulthood.^{4,5} Similar findings have been identified in animal models.^{6,7} Numerous clinical studies have also shown that low levels of vitamin D are directly or indirectly linked to adult psychiatric conditions including depression and cognitive impairment.^{8–10} Clinical studies^{11,12} have implied a relationship between low vitamin D levels and conditions such as depression,^{13,14} substance abuse,¹⁵ schizophrenia,^{16,17} delirium,¹⁸ and suicidal ideation.¹⁹

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Deficiency of vitamin D is common in North America and in the past decade has been described as a major public health problem.² Data from the National Health and Nutrition examination surveys demonstrate that about one-quarter of the US population could be categorized as having insufficient vitamin D levels, and about 7%, frank deficiency.²⁰ A study of healthy preschool-aged African-American children from low-income families in Atlanta, Georgia, reported 26% had low vitamin D levels.²¹ Some risk factors associated with vitamin D deficiency include insufficient sunlight, dark skin, old age, obesity, diabetes, and medications that alter vitamin D absorption, in particular medications such as anticonvulsants, steroids, or antiretrovirals.18,22-25 Residing at higher latitude also represents a risk factor for insufficient vitamin D, particularly latitudes more northerly than 35°N.26

Prior descriptive studies have provided evidence that vitamin D deficiency is especially prevalent in the psychiatric inpatient setting, but the literature is inconsistent. In a psychiatric hospital in the UK, researchers found that 83% of patients had vitamin D deficiency and 92% had below optimal levels of vitamin D.27 Menkes et al28 identified a link between decreased level of vitamin D in psychiatric patients and poor physical health outcomes. Psychiatric patients may be exposed to additional physical risk factors associated with low vitamin D states such as cardiovascular disease.²⁹⁻³¹ Psychiatric patients may also be at risk for vitamin D deficiency due to their age, less exposure to sunlight, and dietary factors.28 Sufficient levels of vitamin D are protective against infectious diseases, autoimmune disorders, type 1 and type 2 diabetes, musculoskeletal disorders, infertility, and adverse pregnancy and birth outcomes.³²

Among psychiatric illnesses, vitamin D supplementation has been extensively studied in depression, as a potential factor in improvement of mood.^{33,34} Vitamin D supplementation is likely to be only one part of treatment, as depression has myriad causes, yet its use could be important as research has indicated that vitamin D deficiency impairs and extends recovery from depression.³³ Gezen-Ak et al³⁵ reported that vitamin D treatment increased the production of glia-derived neurotrophic factor (GDNF). Neurotrophins like GDNF enhance the survival and growth of neurons. Since other neurotrophins (eg, brain-derived neurotrophic factor) and vascular endothelial growth factor are low in depression, the effects of vitamin D on GDNF could be important to its antidepressant effects.

Many of the characteristics linked with insufficient vitamin D levels are present in the psychiatric patient

population in New Orleans, Louisiana (29.95°N), a southern US city that has undergone two recent, major natural disasters that have exacerbated mental health issues.^{36,37} The state is currently underresourced with regard to health facilities and providers dedicated to mental health; the National Association of State Mental Health Program Directors Research Institute reported in 2009 that Louisiana spent \$60.89 per resident for psychiatric institutes and administrative costs in state mental health spending, representing 60% less than the national average of \$104.10.³⁸ Statewide budget cuts over the past several years, and reduced spending on mental health system.³⁹

However, there is little knowledge regarding the actual vitamin D status of psychiatric inpatients in Louisiana and no studies to date on this population. This study aimed to determine the point prevalence of hypovitaminosis D in a local inpatient psychiatric population and to assess any potential relationships between vitamin D levels and risk factors for vitamin D deficiency such as age, gender, race/ethnicity, psychiatric diagnosis, medical diagnosis, length of stay, severity of illness (measured by number of psychiatric hospitalizations), and routine lab measurements obtained during psychiatric hospitalization. This study presents the results of a retrospective chart review with implications for clinical practice.

Methods

We undertook a retrospective chart review utilizing the medical records of the Louisiana State University Health Sciences Center (LSU-HSC), a hospital with a majority of low-income patients who are either uninsured or underinsured. A list of medical records of adult patients aged 18-65 years, admitted to the LSU-HSC inpatient psychiatric unit, were obtained and reviewed. Eligibility criteria included patients with at least one psychiatric diagnosis, one or more inpatient psychiatric hospitalizations since May 1, 2012, and with at least one of those hospitalizations being at the LSU-HSC inpatient unit. Eligible patient records must have also included a vitamin D hydroxy test taken on admission and dated between May 1, 2012, and November 1, 2014. This test is commonly utilized for diagnostic purposes, as it is thought to be a good indicator of exogenous sources of vitamin D such as diet, sunlight exposure, or supplementation. Measurements are based on total 25(OH)D, with levels <20.0 ng/mL indicative of vitamin D deficiency. Insufficient 25(OH) D is defined by levels between 20.0 and 31.9 ng/mL and normal 25(OH)D by levels >31.9 ng/mL. Exclusion criteria included the following: 1) patients for whom the primary diagnosis was

a medical condition at the time of discharge, 2) patients for whom no psychiatric diagnosis could be established or was reported at the time of discharge, 3) patients taking vitamin D supplements at the time of admission, or patients who had taken vitamin D supplements regularly in the 3 months leading up to admission.

The data collected from the 113 patient records included in this study were anonymized at the time of export from electronic medical records, initially assessed in Microsoft Excel (2013), and then subsequently imported for further analysis into SPSS Statistical Package for Windows (Version 23; IBM Corporation, Armonk, NY, USA). The data set was cleaned by an experienced data analyst. The sample utilized for analysis represented the maximum number of electronic medical records available for the researchers to review according to institutional and logistical constraints.

Variables with missing information were coded as "998" and were accounted for in the results and analysis. Previous literature23,40 has described older age and dark skin as potential risk factors associated with vitamin D deficiency and, therefore, "age" was recoded from a continuous to a dichotomous variable with a cutoff value of 50 years and "race/ethnicity" was recoded from a categorical to a dichotomous variable of Caucasian and non-Caucasian. The continuous variable of "vitamin D" was maintained in its original form as well as recorded into a categorical variable in which values <20 ng/ mL were categorized as vitamin D deficient, values of 20-31.9 ng/mL as vitamin D insufficient, and values >31.9 ng/mL as normal vitamin D status in accordance with the LSU-HSC standard values. Descriptive statistics, or univariate analysis, was run on the sample to obtain frequencies, point prevalence, means, standard deviations, and interquartile ranges.

The Institutional Review Boards of LSU-HSC and Tulane University both reviewed the study and deemed it exempt from ethical approval. Due its to retrospective nature and use of only de-identified data, patient consent was not required for this study.

Results

Sample characteristics

As illustrated in Table 1, the total sample size was 113. The average age of psychiatric inpatients included in the sample was 44 years, where 56.6% were male and 43.4% were female, and 59.3% were African American. A majority of the sample population (65.5%) were unemployed at the time of admission, and for 92.9% of the sample, this was not the first admission into a psychiatric inpatient care facility. The average length of stay at LSU-HSC was about 2 weeks (13.4 days).

Characteristics	Percent (%)	Mean of sample	(n)
Age			
Average age		44 years	
>50 years	42.5		48
<50 years	57.5		65
Gender			
Female	43.4		49
Male	56.6		64
Race/ethnicity			
African American	59.3		67
Caucasian	33.6		38
Hispanic	3.5		4
Other	2.7		3
Did not specify	0.9		Ι
Employment status			
Employed	8		9
Unemployed	65.5		74
Did not specify	26.5		30
Admission history			
Length of stay		13.4 days	
>I Admission	92.9		105
Vitamin D status on adn	nission		
Average level (ng/dL)		22.5	
Normal (30–100 ng/dL)	15.0		16
Insufficient (20–30 ng/dL)	36.3		41
Deficient (<20 ng/dL)	48.7		55
Psychiatric diagnosis			
Schizophrenia	32.7		37
Bipolar I disorder	34.5		39
Major depressive disorder	29.2		33
Depression NOS	30.1		34
Schizoaffective disorder	21.1		24
Psychosis unspecified	18.6		21
Anxiety disorder	13.3		15
Drug abuse disorder			
Alcohol use	60.2		68
Cannabis use	52.5		59
Opiate use	28.2		32
Cocaine use	41.6		47
Medical diagnosis			
Hypertension	38.9		44
Hyperlipidemia	10.6		12
Diabetes	14.2		16

Abbreviation: NOS, not otherwise specified.

Vitamin D levels

Table 2 presents the results for vitamin D status among the sample. Only 15% of patients in the sample had a normal vitamin D level. Mean vitamin D status on admission was insufficient (22.5 ng/mL) and at the lower end of the vitamin D insufficiency range (20–31.9 ng/mL) verging on deficient. Although the mean level of vitamin D on admission was in the insufficiency range, nearly one-half (48.7%) of the sample population was admitted with deficient levels of vitamin D (<20 ng/mL). Vitamin D levels were also

Table 2 Vitamin E) status by selected	l sample characteristics	(N=113)
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	Deficient % (n)	Insufficient % (n)	Normal	Number in category (n)	Mean vitamin D levels
			% (n)		ng/mL (SD)
Sample	48.7 (55)	36.3 (41)	15.0 (17)	113	22.57 (11.09)
Age					
>50 years	50.0 (24)	31.1 (15)	18.8 (9)	48	23.49 (13.47), P=0.586
<50 years	47.7 (31)	40.0 (26)	12.3 (8)	65	21.87 (8.98)
Gender					
Male	37.5 (24)	37.5 (24)	25.0 (16)	64	24.28 (11.34), P=0.058
Female	63.3 (31)	26.5 (13)	10.2 (5)	49	20.30 (10.43)
Race/ethnicity					
Non-Caucasian*	60.6 (43)	29.6 (21)	9.9 (7)	71	20.94 (11.75), P=0.035
Caucasian	29.3 (12)	46.3 (19)	24.4 (10)	41	25.61 (9.19)
Psychiatric disorder					
Schizophrenia	51.4 (19)	35.1 (13)	13.5 (5)	37	21.81 (11.94), <i>P</i> =0.623
Bipolar I	46.2 (18)	33.3 (13)	20.5 (8)	39	24.62 (13.53), P=0.152
Major depressive disorder	42.4 (14)	42.4 (14)	15.2 (5)	33	21.95 (9.37), P=0.713
Depression NOS	44.1 (15)	32.4 (11)	23.5 (8)	34	24.01 (10.14), P=0.361
Schizoaffective disorder	58.3 (14)	37.5 (9)	4.2 (1)	24	19.98 (8.31), <i>P</i> =0.201
Psychosis unspecified	71.4 (15)	14.3 (5)	14.3 (3)	21	19.25 (11.49), <i>P</i> =0.131
Anxiety disorder	33.3 (5)	53.3 (8)	13.3 (2)	15	23.71 (11.22), P=0.668
Substance abuse disorder					
Alcohol	50.0 (34)	29.4 (20)	20.6 (14)	68	23.45 (12.58), P=0.298
Cannabis	49.2 (29)	37.3 (22)	13.6 (8)	59	22.32 (11.15), <i>P</i> =0.754
Opiates	43.8 (14)	34.4 (11)	21.9 (7)	32	23.58 (11.52), P=0.978
Cocaine	42.6 (20)	44.7 (21)	12.8 (6)	47	23.47 (11.85), P=0.639

Notes: *Non-Caucasian includes African American and Hispanic. Medical records indicating race/ethnicity as "Other", or where none was specified were treated as missing values (n=4).

Abbreviation: NOS, not otherwise specified.

analyzed categorically; 36.3% had insufficient vitamin D levels (20–31.9 ng/mL) and 48.7% deficient vitamin D levels (<20 ng/mL).

The lowest mean levels of vitamin D were found among patients diagnosed with schizoaffective disorder (19.98 ng/mL) and unspecified psychosis (19.25 ng/mL); however, the results were not statistically significant (P=0.201 and 0.131, respectively). The most common psychiatric diagnoses were bipolar 1 disorder (34.5%), schizophrenia (32.7%), and depression not otherwise specified (30.1%). Other diagnoses included major depressive disorder, schizoaffective disorder, unspecified psychosis, and anxiety disorder. Alcohol use (60.2%) and cannabis use (52.5%) were the most common substance abuse disorders, followed closely by cocaine use (41.6%). A little over a quarter of the sample (28.2%) had a diagnosis of opiate abuse disorder. Less than half of the sample population had any compounding medical diagnosis. These included cardiovascular diseases such as hypertension (38.9%) and hyperlipidemia (10.6%), and either type 1 or type 2 diabetes (14.2%), where no significant differences were found with regard to vitamin D status.

Age also did not result in any significant differences in this sample. For those over the age of 50 years, half (50%)

were deficient in vitamin D, and for those under the age of 50 years, 47.7% were deficient. The mean vitamin D levels for gender were significantly different (P=0.058), where females were found to have a mean vitamin D level of 20.30 ng/mL and males a mean value of 24.28 ng/mL. Race/ ethnicity was also found to have significantly different mean values of vitamin D (P=0.035). Caucasian patients had a mean vitamin D level of 25.61 ng/mL, while non-Caucasians had a mean value of 20.94 ng/mL. This translates into 60.6% of non-Caucasians being vitamin D deficient compared to Caucasians, among whom 29.3% were deficient.

All patients in this sample had at least one psychiatric diagnosis but may have had more than one. Therefore, it was impossible to compare deficiency within the category of psychiatric diagnosis. Looking at univariate analysis, however, it is evident that 51.4% of schizophrenics were vitamin D deficient and 35.1% were insufficient. Of those with bipolar 1 disorder, 46.2% had deficient vitamin D levels and 33.3% had insufficient levels. It is interesting to note that the prevalence of non-normal vitamin D levels in those with bipolar 1 disorder was not greatly different from those without bipolar 1 disorder, who had values of 50% and 37.8% for deficiency and insufficiency, respectively. Similar results were found in those with depression and schizoaffective disorder, in that

there is almost no difference between deficiency levels of vitamin D in those with the disorder compared to those without the disorder (22.85 and 22.27 ng/mL, respectively, for any depression diagnosis; 19.98 and 28.25 ng/mL, respectively, for schizoaffective disorder).

Unspecified psychosis and anxiety disorder showed a larger difference in deficient levels of vitamin D in those with the disorder compared to those without the disorder (71.4% and 43.5%, respectively, for unspecified psychosis and 33.3% and 51%, respectively, for anxiety disorders). However, their mean vitamin D levels remained insignificant (P=0.131 for unspecified psychosis and P=0.668 for anxiety disorders). As stated, all study subjects had at least one psychiatric diagnosis but many had more than one from a previous hospitalization in a different facility or following evaluation by a non-LSU-HSC health provider. For this reason, we could not ascertain with certainty why depression and schizoaffective disorder exhibited almost no difference in vitamin D deficiency or why bipolar 1 disorder resulted in the reported values. Future investigations, such as a case-control or a chart review, that include patients with non-psychiatric diagnoses may offer better solutions in explaining these possible differences.

Substance abuse did not result in significantly different mean deficiency levels between those with a substance abuse disorder and those without the disorder. Fifty percent of those with an alcohol use disorder were vitamin D deficient but 46.7% of those without an alcohol disorder were also vitamin D deficient. There was almost no difference in cannabis users, with 49.2% of users being deficient in vitamin D and 49.1% of nonusers also being deficient. A total of 43.8% of opiate users and 50.6% of non-users were vitamin D deficient. Similarly, 42.6% of cocaine users and 55% of non-users were vitamin D deficient. As with psychiatric diagnosis, and due to the nature of this study, it was not possible to explore these results beyond univariate analysis.

Discussion

To date, this is the first study to determine the point prevalence of vitamin D deficiency in a psychiatric hospital patient population in Louisiana. Serum 25(OH)D levels in our sample indicate a high prevalence of vitamin D deficiency and insufficiency among hospitalized psychiatric patients. These results are consistent with similar studies conducted by Zeng et al in the US Midwest,⁴¹ McCue et al in New York,²² Berk et al in Australia,¹¹ and Humble et al in northern Europe.⁴² However, in our study, 85% of the sampled patients had lower than normal vitamin D, a much higher level than that identified in similar studies. A 2016 cross-sectional

study in the US determined the prevalence of vitamin D deficiency among psychiatric inpatients and found that 30% of patients had sufficient levels,⁴¹ double that of the present study. Per Centers for Disease Control data, ~20%–25% of North American people are considered to be vitamin D insufficient and 7% had hypovitaminosis D.⁴³

Our mean 25(OH)D value was also lower than the levels found generally in the US population, as illustrated by the 2009–2010 NHANES sample, where the mean was 25.67 ng/mL.⁴³ The mean vitamin D level for this sample was 22.56 ng/mL. The standard deviation at the first stage of analysis was 11.086, which indicates a wide range of variation around the mean. Upon closer inspection it was found that the study sample contained 3 outlying values, and when those values were set to missing and the analysis rerun, the mean vitamin D level decreased slightly to 21.49 ng/mL and the standard deviation decreased to 9.016. Because most nutrients are positively skewed, the skewed nature of this sample is to be expected and, therefore, the outlying values were maintained in the study sample.

Despite being lower than in the general population, the mean serum 25(OH)D level in our study was slightly higher than the levels observed in other similar studies conducted in psychiatric hospital settings: 20.9 ng/mL in long-term psychiatric inpatients in Virginia,⁴⁴ 17.2 ng/mL in psychiatric inpatients and outpatients in Norway,²⁰ and <20 ng/mL among the psychiatric patients in a London hospital.²⁷ The differences are likely due to the latitude of New Orleans being lower than that found in other studies, though they could also represent increased risk factors for vitamin D deficiency among a lower socioeconomic status population. For example, our study indicated lower levels than those found in psychiatric patients in the US Midwest (26.7 ng/mL) where socioeconomic status is higher though latitude is more northerly.⁴⁵

The intensity of ultraviolet (UV) light in the critical 290–315 nm waveband reaching the earth's surface is determined by the zenith angle of the sun, which varies considerably with the month of the year and with latitude⁴⁶ affecting the amount of 25(OH)D level related to sun exposure.² Kimlin et al⁴⁷ reported that increase in latitude dramatically decreases the amount of vitamin D UV, which may inhibit vitamin D synthesis in humans. New Orleans is situated at latitude 29.9°N, so vitamin D synthesis should be higher as compared to other locations mentioned earlier. Generally, people who live at higher than 35°N latitudes are at increased probability of having vitamin D insufficiency, which also explains the differences in serum 25(OH)D levels from our study and other studies conducted in different people living in different latitudes.^{42,47} The highest level is mostly observed 2 months after the summer in the northern hemisphere.⁴² Lucas et al⁴⁸ showed this seasonal variation with a lag of 8 weeks between the maximum monthly UV dose and the peak 25(OH)D level. The time taken to establish the steady state of a compound is ~3–4 times its half-life, which for 25(OH)D is about 22 days. Thus, this lag period may represent the time taken to establish a new steady state.^{48,49} The 25(OH)D level variation in our study may be related to sample collection during this period. It is also possible that some of these differences can be due to the different 25(OH)D assay methods applied by different laboratories.⁵⁰

African American patients had significantly lower vitamin D levels compared to Caucasian patients in our study. According to the April 2010 US census, the racial composition of New Orleans that consisted of 60.2% African American and 33% Caucasian population, respectively,51 was similar to the racial composition of our study, indicating that race or ethnicity may play an important part when determining confounding factors for vitamin D status in psychiatric patients. The sample for this study is representative of the two major racial groups in the New Orleans population, and this largely excludes the possibility that the difference was associated with sample collection but could be due to differences in skin pigmentation.⁵² Anything that can cause interference with the penetration of sunlight into the skin will affect the synthesis of vitamin D3, and melanin is extremely efficient in absorbing sunlight, leading to reduction in vitamin D3 synthesis.53 Some studies have hypothesized that vitamin D insufficiency could account for the increased risk of schizophrenia observed among dark-skinned immigrant people.^{54,55} Based on our data, no sufficient evidence is available to support or refute this hypothesis.

Cardinal and Gregory²⁷ and Humble²⁶ indicated that vitamin D deficiency is related to increased age.^{26,27} In this study, however, 31.1% patients were >50 years of age and 40% were <50 years of age and we did not observe any significant difference in vitamin D levels between those two groups, though gender was borderline significant. Several investigators have shown that low vitamin D levels are significantly associated with cardiovascular risk factors, obesity,²⁴ and diabetes.²⁵ Though it is common to find these medical illnesses coexisting with lower vitamin D levels, this study identified no significant correlation between them. Based on the risk of metabolic syndrome being related to antipsychotic medications use, which might be related to vitamin D deficiency, the effect of antipsychotic medications on vitamin D levels remains elusive.⁴⁰

Limitations

This study had several limitations, most inherent to retrospective studies. First, there were no controls of similar age and sex distribution, so it is not clear whether the results could be generalized to inpatients in other clinical settings. Second, the present study was at risk of misclassification and selection bias. Third, the sample size in our study was small and limited due to logistical and institutional constraints. In addition, the impact of multiple psychiatric diagnoses obscured the relationship of specific individual diagnoses on vitamin D levels.

Conclusion

Monitoring of vitamin D levels on hospital admission for psychiatric illness and supplementation of vitamin D may be an important and beneficial strategy, especially for African Americans. Supplements are generally safe and easy to administer. Other interventions to prevent vitamin D insufficiency among this population could also be considered at the community level, including adequate sunlight exposure and dietary balance. In addition, it could be of important diagnostic value to monitor vitamin D levels as a routine screening test in all psychiatric patients. Further studies are needed to explore the explicit role of vitamin D in the causality of psychiatric illness, but the present data further highlight the need for awareness of this issue among behavioral health providers.

Disclosure

The authors report no conflicts of interest in this work.

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