

Prevalence of Serum Vitamin B₁₂ Levels among Patients Presented to Hebron Charitable Medical Center, Palestine

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Abstract: Background: Vitamin B₁₂ is an essential element for proteins, phospholipids metabolism and neurotransmitters synthesis. It also plays a key role in DNA biosynthesis and cellular energy production. However, vitamin B₁₂ deficiency leads to neurologic and psychiatric disturbances including depression, dementia, and a demyelinating myelopathy. Moreover, it's also associated with other diseases such as pernicious anemia and other hematologic manifestation. **Methods:** After approval from Hebron Charitable Medical Center (HCMC), a retrospective study was carried out among patients at Charitable Medical Center (HCMC) in Hebron, Palestine. It was carried out over a period of one year and half starting from January 2013 till June 2014. **Aim:** This study was conducted to determine the prevalence of vitamin B₁₂ among the patients of HCMC. **Results:** In general, a total of 305 patients screened for serum vitamin B₁₂ levels were enrolled. There were 90 (29.5%) men and 215 (70.5%) women, there were 285 (93.4%) <60 years old and 20 (6.6%) >60 years old. There were 50 patients (16.4%) had vitamin B₁₂ deficiency. Mean \pm SEM levels of B₁₂ in 305 subjects were observed to be 389.73.1 \pm 12.93 (range: 60 - 1432 pg/ml). Men had mean value of 423.11 \pm 30.17 (range: 60 - 1432 pg/ml) and women had 375.75 \pm 13.24 (range: 76 - 1292 pg/ml). **Conclusion:** This study had highlighted a number of patients that reveals Vitamin B₁₂ deficiency according to gender and age. We recommend the necessary implementation of Vitamin B₁₂ supplements for all men and women with any age.

Keywords: Vitamin B₁₂, Deficiency, Gender, Age.

Introduction

Vitamin B₁₂ also called cobalamin is an essential element for proteins, phospholipids metabolism and neurotransmitters synthesis. It also plays a key role in DNA biosynthesis and cellular energy production (O'Leary & Samman). Vitamin B₁₂ sources are mainly from animal origin, since it's synthesized by certain bacteria in the gastrointestinal tract of animals then absorbed by the host animal, however it's concentrated in animal tissues (Heyssel, Bozian, Darby, & Bell, 1966). So insufficient intake or malabsorption of this vitamin leads to vitamin B₁₂ deficiency, which is often seen mainly among vegetarians and people who are suffering from gastrointestinal conditions (Andrès et al., 2004). Hence, this vitamin deficiency leads to neurologic and psychiatric disturbances including depression, dementia, and a demyelinating myelopathy (Bottiglieri, 1996). Moreover, it's also associated with other diseases such as pernicious anemia

and other hematologic manifestation (Clementz & Schade, 1990).

Vitamin B₁₂ deficiency patients may present different signs and symptoms including fatigue, weakness, numbness, decreased memory, irritability, confusion, although initial symptoms might often be vague (Oh & Brown, 2003) In fact vitamin B₁₂ affects all ages (Healton, Savage, Brust, Garrett, & Lindenbaum, 1991) especially older people (Miles, Mills, Clarke, & Dangour, 2015). To get a specific and clear diagnosis of vitamin B₁₂ deficiency should be done by measuring the serum homocysteine and methylmalonic acid levels, since they are high in vitamin B₁₂ deficiency, and considered as reliable indicators of vitamin B₁₂ deficiency than the concentration of B₁₂ in blood (Donaldson, 2000).

The prevalence of vitamin B₁₂ deficiency in people varies widely depending on the population's heterogeneity and lifestyle.

Till now establishing a universally normal range for vitamin B₁₂, as well as, prevalence of vitamin B₁₂ deficiency in the general population has not been well established (Carmel, 2000). Therefore, it is not always easy to decide whether a patient suffers from vitamin B₁₂ deficiency or not. From here, early detection of deficiency is imperative for treatment to be effective, and timely screening and replacement of this vitamin deficiency will help prevent many complications and side effects that may appear in late stages. So treatment can be ensured either by injections or orally doses of vitamin B₁₂ (Hvas & Nexø, 2006). It's clinically preferable to take into account preventative measures to avoid deficiencies, which include improving life style and diet control, especially for vegetarian people, and try to eat plenty food-based sources of vitamin B₁₂, such as fish, poultry, and eggs, also vitamin B₁₂ commercially available supplements could be used (Watanabe, 2007).

Material and Methods

A retrospective study was carried out among patients at Charitable Medical Center (HCMC), it was carried out over a period of one year and half starting from January 2013 till June 2014. Patients were recruited from the laboratory records using convenience sampling. All of the medical records belonging to the patients of interest whose vitamin B₁₂ was assayed were screened to collect data about their vitamin B₁₂ levels, ages, and genders. The targeted patients were from the district of Hebron of the Palestinian territories; therefore, they are from the same culture and religious background.

Based on our laboratory's protocol, patients were asked for overnight fasting, and then blood samples were drawn by a professional phlebotomist for vitamin B₁₂ level test. Serum vitamin B₁₂ level test was performed using commercially available kits (Access 2 vitamin B₁₂ Beckman Coulter). According to our laboratory normal range standards, individuals whose serum vitamin B₁₂ levels were <200 pg/ml are considered as vitamin B₁₂ deficient people, while the vitamin B₁₂ concentrations at 370 pg/ml were roughly

considered as borderline deficiency. Moreover, individuals whose vitamin B₁₂ levels were noticed to be between 200 pg/ml and 900 pg/ml were taken as normal individuals, more than 900 pg/ml were taken up normal level. The sample included 305 men and women who were tested the serum vitamins B₁₂ levels.

Statistical Analysis

After data collection, the data were reviewed, organized, tabulated and statistically analyzed using SPSS version 23 (Statistical Package for Scientific Studies). Descriptive statistics (e.g. Frequencies & descriptive) were used to analyze the data. Inferential statistics were used to given a mean score of Vitamins B₁₂ and also using T-test (one way Anova) to determine if there is a significance difference between groups. All tests were conducted at level of significance $\alpha = 0.05$, results with p-values <0.05 will be considered statistically significant.

Ethical Considerations

We have stressed the anonymity and confidentiality of any collected information, and that only generic outcome data might be disseminated in scientific settings.

Results

A total of 305 patients screened for serum vitamin B₁₂ levels were enrolled. Out of the total 305 patients, there were 90 men (29.5%) and 215 women (70.5%), there were 285 (93.4%) whose are under 60 years and 20 (6.6%) whose are above 60 years old. Out of the total 305 patients. Moreover, there were 50 (16.4%) patients had vitamin B₁₂ deficiency.

Table 1 depicts serum vitamin B₁₂ levels and percentage of male and female studied. Mean±SEM levels of B₁₂ in 305 subjects were observed to be 389.73.1±12.93 (range: 60 - 1432 pg/ml).

Men had mean value of 423.11±30.17 (range: 60 - 1432 pg/ml) and women had 375.75±13.24 (range: 76 - 1292 pg/ml).

Table 1: Vitamin B12 level mean of subjects according to gender.

	Mean	Std. Error of Mean	Std. Deviation	Min	Max
Total results of all participants (n=305)					
n= 305 (ALL)	389.73	12.93	225.8827	60.0	1432.0
Total results of Men participants (n=90)					
n= 90 (Men)	423.11	30.17	286.2859	60.0	1432.0
Total results of Women participants (n=215)					
n= 215 (Women)	375.75	13.24	194.2433	76.0	1292.0

Table 2. If 200 pg/ml was taken as cut off for deficiency state, total 50 (16.4%) out of 305 subjects turned out to be B₁₂ deficient. In this group, 16 (17.8%) were men with mean levels of 133.9±11.6 pg/ml and 34 (15.8%) were women

with mean of 146.6±5.0 pg/ml. However, it was observed that 246/305 (80.7%) subjects had B₁₂ Normal levels. There were 67 males (74.4%) with Mean level 413.2±20.3 and 179 females (83.3%) with mean level 410.8±12.3.

Table 2: vitamin B12 percentage out of total study subject.

	Deficiency level n(%)		Normal level n(%)		Up normal level n(%)		Total n(%)	
	Mean ± SEM		Mean ± SEM		Mean ± SEM		Mean ± SEM	
Male	16 (17.8%)	133.9 ± 11.6	67 (74.4%)	413.2 ±20.3	7 (7.8%)	1178.9 ± 48.2	90 (29.5%)	423.1 ± 30.2
Female	34 (15.8%)	146.6 ± 5.0	179 (83.3%)	410.8 ±12.3	2 (0.9%)	1132.5 ± 159.5	215 (70.5%)	375.8 ± 13.2
Total	50 (16.4%)	142.6 ± 5.0	246 (80.7%)	411.5 ±10.5	9 (3.0%)	1168.6 ± 45.9	305 (100%)	389.7 ± 12.9

Table 3. If 200 pg/ml was taken as cut off for deficiency state, total 50 out of 305 subjects (16.4%) turned out to be B₁₂ deficient. In this group, 46 (16.1%) were <60 with mean

levels of 133.9±11.6 pg/ml and 4(20%) were >60 with mean of 146.6±5.0 pg/ml.

Table 3: Vitamin B12 percentage out of total study subject according to age.

	Deficiency level n(%)		Normal level n(%)		Up normal level n(%)		Total n(%)	
	Mean ± SEM		Mean ± SEM		Mean ± SEM		Mean ± SEM	
<60	46 (16.1%)	133.9 ± 11.6	234 (82.1%)	413.2 ± 20.3	5 (1.8%)	1178.9 ± 48.2	285 (93.4%)	423.1 ± 30.2
>60	4 (20.0%)	146.6 ± 5.0	12 (60.0%)	410.8 ± 12.3	4 (20.0%)	1132.5 ± 159.5	20 (6.6%)	375.8 ± 13.2
Total	50 (16.4%)	142.6 ± 5.0	246 (80.7%)	411.5 ± 10.5	9 (3.0%)	1168.6 ± 45.9	305 (100.0%)	389.7 ± 12.9

Table 4. Analysis of serum Vitamin B12 by socio-demographic characteristics of respondents: In comparing the serum Vitamin B₁₂ level, there is no significant

difference between Male and Female (p = 0.153) and no significant difference between <60 and >60 (p= .143).

Table 4: Differences of vitamin B₁₂ levels according to age and gender

Variable	Group (n)	Mean (SD)	F statistic (df)	P value
Age	<60(285)	381.44 (211.1)	12.35 (19.89)	.143
	>60(20)	507.80 (366.4)		
Gender	Male(90)	423.11 (286.2)	10.20 (124.68)	.153
	Female(215)	375.75 (194.2)		

Discussion

The prevalence of vitamin B₁₂ deficiency was 16.4 % in this representative sample. This is in accordance with the previously reported prevalence rates in other countries (Baik & Russell, 1999; Clarke et al., 2004; Loikas et al., 2006; Wolters, Ströhle, & Hahn, 2004).

Vitamin B₁₂ deficiency is prevalent in different countries in the world, because of insufficient intake from different food sources, and/or some troubles in absorbing of vitamin B₁₂. The state of deficiency may present different types of signs and symptoms like neuropsychiatric problems and other associated maladies so Cobalamin occurs in substantial amounts only in foods derived from animals and is essential for one-carbon metabolism and cell division. Low nutritional intake of vitamin B-12 may lead to negative balance, and ultimately, to functional deficiency when tissue stores of vitamin B₁₂ are depleted. Early diagnosis of vitamin B-12 deficiency seems to be useful because

irreversible neurological damages may be prevented by cobalamin substitution (Herrmann & Geisel, 2002).

In this study, It was found that 20% of older patients have vitamin B₁₂ 200 pg/ml or below. There are some studies that support our results, these studies revealed that vitamin B₁₂ deficiency increases in the elderly, mainly because atrophic gastritis decreases the production of the acid and digestive enzymes needed to cleave protein-bound vitamin B₁₂ from the natural chemical form of vitamin B₁₂ found in meat, poultry, fish, and dairy foods and also depending on the biochemical criterion that is used, 5% to more than 20% of older adults have marginal or frank vitamin B₁₂ deficiency (Park & Johnson, 2006).

Moreover, there are Sixteen percent of geriatric outpatients had serum Cobalamin levels of 200 pg./mL or below(Yao, Yao, Yao, & Lou, 1992).

In addition, our study showed that the male have Vitamin B₁₂ deficiency 17.6% compared with 15.8% in female.

These results are supported by some particular studies which appeared that men and women have Vitamin B₁₂ deficiency 59% and 23.8% respectively (Carmel, Mallidi, Vinarskiy, Brar, & Frouhar, 2002). Moreover, 34(8%) of men and 28 (4.7%) of women have Vitamin B₁₂ deficiency (Loikas et al., 2006).

Conclusion

We conclude that, undiagnosed vitamin B₁₂ deficiency is remarkably common in the community of Hebron. This suggests that, in the current clinical practice, only some overt signs and symptoms the individual may have trigger laboratory testing for vitamin B₁₂ deficiency. Evidently, routine screening would help to have earlier diagnosis for individuals and reduce the related problems. No specific risk groups among the aged can be defined, but aging itself increases the probability of vitamin B₁₂ deficiency, so we recommend the necessary implementation of Vitamin B₁₂ supplements for all men and women with any age.

Acknowledgements

Authors are highly appreciated to the Hebron Charitable Medical Center team for cooperation.

Conflicts of Interest

There are no conflicts of interests

References

- [1] Andrès, E., Loukili, N. H., Noel, E., Kaltenbach, G., Abdelgheni, M. B., Perrin, A. E., Blicklé, J.-F. (2004). Vitamin B₁₂ (cobalamin) deficiency in elderly patients. *Canadian Medical Association Journal*, 171(3), 251-259.
- [2] Baik, H., & Russell, R. (1999). Vitamin B₁₂ deficiency in the elderly. *Annual review of nutrition*, 19(1), 357-377.
- [3] Bottiglieri, T. (1996). Folate, vitamin B₁₂, and neuropsychiatric disorders. *Nutrition reviews*, 54(12), 382-390.
- [4] Carmel, R. (2000). Current concepts in cobalamin deficiency. *Annual review of medicine*, 51(1), 357-375.
- [5] Carmel, R., Mallidi, P. V., Vinarskiy, S., Brar, S., & Frouhar, Z. (2002). Hyperhomocysteinemia and cobalamin deficiency in young Asian Indians in the United States. *American journal of hematology*, 70(2), 107-114.
- [6] Clarke, R., Grimley Evans, J., Schneede, J., Nexo, E., Bates, C., Fletcher, A., Refsum, H. (2004). Vitamin B₁₂ and folate deficiency in later life. *Age and ageing*, 33(1), 34-41.
- [7] Clementz, G., & Schade, S. (1990). The spectrum of vitamin B₁₂ deficiency. *American family physician*, 41(1), 150-162.
- [8] Donaldson, M. S. (2000). Metabolic vitamin B₁₂ status on a mostly raw vegan diet with follow-up using tablets, nutritional yeast, or probiotic supplements. *Annals of nutrition and metabolism*, 44(5-6), 229-234.
- [9] Healton, E. B., Savage, D. G., Brust, J. C., Garrett, T., & Lindenbaum, J. (1991). Neurologic aspects of cobalamin deficiency. *Medicine*, 70(4), 229-245.
- [10] Herrmann, W., & Geisel, J. (2002). Vegetarian lifestyle and monitoring of vitamin B-12 status. *Clinica Chimica Acta*, 326(1), 47-59.
- [11] Heyssel, R., Bozian, R., Darby, W., & Bell, M. (1966). Vitamin B₁₂ turnover in man. The assimilation of vitamin B₁₂ from natural foodstuff by man and estimates of minimal daily dietary requirements. *American Journal of Clinical Nutrition*, 18, 176-184.
- [12] Hvas, A.-M., & Nexo, E. (2006). Diagnosis and treatment of vitamin B₁₂ deficiency--an update. *Haematologica*, 91(11), 1506-1512.
- [13] Loikas, S., Koskinen, P., Irjala, K., Löppönen, M., Isoaho, R., Kivelä, S.-L., & Pelliniemi, T.-T. (2006). Vitamin B₁₂ deficiency in the aged: a population-based study. *Age and ageing*, 36(2), 177-183.
- [14] Miles, L. M., Mills, K., Clarke, R., & Dangour, A. D. (2015). Is there an association of vitamin B₁₂ status with neurological function in older people? A systematic review. *British Journal of Nutrition*, 114(4), 503-508.
- [15] O'Leary, F., & Samman, S. More About Vitamins.
- [16] Oh, R., & Brown, D. L. (2003). Vitamin B₁₂ deficiency. *American family physician*, 67(5), 979-986.
- [17] Park, S., & Johnson, M. A. (2006). What is an adequate dose of oral vitamin B₁₂ in older people with poor vitamin B₁₂ status? *Nutrition reviews*, 64(8), 373-378.
- [18] Watanabe, F. (2007). Vitamin B₁₂ sources and bioavailability. *Experimental Biology and Medicine*, 232(10), 1266-1274.
- [19] Wolters, M., Ströhle, A., & Hahn, A. (2004). Cobalamin: a critical vitamin in the elderly. *Preventive medicine*, 39(6), 1256-1266.
- [20] Yao, Y., Yao, S.-L., Yao, S.-S., Yao, G., & Lou, W. (1992). Prevalence of vitamin B₁₂ deficiency among geriatric outpatients. *Journal of family practice*, 35(5), 524-529.