

RESEARCH ARTICLE

SEROPREVALENCE OF HEPATITIS B VIRUS AMONG MULTI-TRANSFUSED BETA THALASSEMIA PATIENTS ATTENDING AT THE YEMENI SOCIETY FOR THALASSEMIA - SANA'A, YEMEN

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Abstract

Regular blood transfusion is one of the most important and emergency treatment methods for thalassemia patients. This category of patients is considered sensitive to infection because their continued survival is linked to blood transfusion, and therefore the possibility of infection is great, especially in light of the war and the deterioration of health services in Yemen. Among this infection is the hepatitis B virus, which causes serious complications such as acute hepatitis, liver cirrhosis and hepatocellular carcinoma (HCC). The aim of the study was done to determine incidence of hepatitis B virus and risk factors among Multi-Transfused Beta Thalassemia Patients attending at The Yemeni Society for Thalassemia and Genetic Blood Disorder (YSTGBD) Sana'a –Yemen. This study was conducted during the period from January 2021 to December 2021, during which 200 Blood samples were collected from beta thalassemia patients attending at (YSTGBD). All samples were examined using the ECLIA technique, during which the basic information was collected through special questionnaires for this purpose. The rate of incidence by HBsAg in all study samples was determined to be 7(3.5%). The study showed that there was a statistically significant relationship between infection with the hepatitis B virus and the age group of more than 10 years age ($P = 0.042$), in addition to that number of blood transfusions received by HBsAg positive was significantly higher than that of HBsAg negative thalassemia patients ($P = 0.022$).

Keywords: β -thalassemia patients, Hepatitis B virus, Yemeni Society for Thalassemia (YST).

1. Introduction

Thalassemia is one of the most common genetic diseases worldwide [1]. β -thalassemia constitutes a major health problem in Yemen. There are about 632 Beta-thalassemia patients that estimated Yemeni society for Thalassemia And genetic Blood disorders in Sana'a, Yemen. The prevalence of β thalassemia trait is 4.4% with an estimated incidence of 11.3/10,000 of homozygous [2]. Recent data indicate that about 7% of the World's population is a carrier of a hemoglobin disorder and that 3,00,000-5,00,000 children are born each year with the severe homozygous states of these diseases [3] Regular blood transfusion in patients with thalassemia has improved their overall survival and quality of life, but it can increase risk of transmission of blood born viral infections, especially HBV [4].

2. Subjects and Methods

2.1 Study population and Setting:

This cross-sectional study was carried out using patients with β -thalassemia Patients (n= 200) attending at the Yemeni Society for Thalassemia in Sana'a. Ethical approval for the study was obtained from Department of Biological Science, Faculty of Sciences, Sana'a University. Information was collected by questionnaire that included demographical data, including: (age, gender, frequency of blood Transfusions, and risk factors including Hepatomegaly, splenectomy, splenomegaly and Iron overload).

2.2 Samples Collection:

5-ml of Blood samples were collected into sterile tube before the blood transfusion allowed to clot at room temperature for 25 minutes, and centrifuged.

2.3 Sampling processing:

The sera were transported inside cryo box to Molecular Biology Research Center (USTY) Sana'a, at the time of study the determinate of Hemoglobin (Hb) levels, Serum Ferritin Levels, Alanine Transaminase level (ALT) were performed by using Semi-Automatic Clinical Chemistry Analyzer spectrophotometer (Mod-Analyzer BAS-100 plus, Labomed, USA). All sera separated into Eppendorf tubes were labeled and stored in deepfreeze at -20 °C, Serological assay were performed by using the Electrochemiluminescence immunoassay (ECLIA) method on Roche cobas e 411, the ECLIA kits used to serological analysis of HBsAg infection markers amongst the participants in this study were performed by using Roche Cobas reagents [5].

2.4 Statistical analysis:

The collected data was coded and entered in a data base file. After complete entry, data were transferred to the IBM SPSS statistics 21 (Copyright IBM Corporation, 2012). The Chi-square test for categorical variables was used to find significant associations between patient's characteristics and HBV positivity. Statistical tests were conducted at the $P < 0.05$ significance level.

3. Results:

3.1 Patients disposition:

in this study, a total of two hundred multi-transfused beta-thalassemia patients, 111(55.5%) males and 89(44.5%) Females were enrolled during a period of Jun 2021 to Dec. 2021 in Sana'a city in this study, 7(3.5%) were positive for Hepatitis B Virus. All thalassemic patients collected from 12 governorates. The average age of patient was divided in two groups, ≤ 10 years age group was 103(51.5%) and > than 10 years age group was 87(48.5%).

3.2 Distribution of Hepatitis B Virus according the socio demographic condition:

The current study results presented prevalence of the HBV-positive among thalassemia patients were found 6 (3%) among more than 10 years age group, followed 1(0.5%) among ≤ 10 years age group There was significant association between HBsAg positive and > than 10 years of blood transfusion age group, Chi-Square = 4.140, P = 0.042 and OR= 8.8, the high prevalence of HBV was recorded in the group <100 times of blood transfusion 6 (3%), while was 1(0.5%) in >100 times of blood transfusion group., the sero-prevalence of HBV markers (HBsAg) was 2 (1%) in male and 5 (2.5%) in female thalassemia patients **Table. 1.**

Table. 1: Distribution of HBsAg positivity according to their socio demographic condition:

	HBsAg						Total	P. value	X2	OR
	Positive (n=7)		Negative (n=193)							
	N	%	N	%	N	%				
Age										
≤ 10 years	1	(0.5)	102	(51)	103	51.5	0.042*	4.140	8.88	
> than 10 years	6	(3)	91	(45.5)	97	48.5				
Total	7	(3.5)	193	(96.5)	200	100				
Number of blood transfusion time										
≤ 10 years of blood transfusion	1	(0.5)	114	(57)	115	57.5	0.022*	5.717	6.88	
> than 10 years of blood transfusion	6	(3)	79	(38.5)	85	42.5				
Total	7	(3.5)	191	(95.5)	200	100				
Gender										
Male	2	(1)	109	(54.5)	111	55.5	0.144	2.13	-	
Female	5	(2.5)	84	(42)	89	44.5				
Total	7	3.5	193	96.5	200	100				

X2 = Chi-Square, OR= Odd Ratio.

3.3 Distribution of HBsAg positivity according to their Governorates:

Regions wise distribution of the prevalence of HBV-positive among thalassemia patients were found, 1 (14.3%) were from Amran, 2 (28.6%) from Hajjah, 1(14.3%) from Al-Hudaydah, 1 (14.3%) from Rima, 1(14.3%) from Sana'a, 1(14.3%) from Taiz, and sero-negative of HBV were found in Al-Bayda, Ibb, Al-Mahwit, Al- Dhali, Sa'da, and Dhamar **Table. 2.**

Table. 2: Distribution of HBsAg positivity according to their Governorates

Governorates	HBsAg				Total	
	Positive (n=7)		Negative (n=193)			
	N	%	N	%	N	%
Amran	1	14.3	31	16.1	32	16
Al-Bayda	0	0	4	2	4	2
Al-Dali	0	0	1	0.5	1	0.5
Hajjah	2	28.5	38	19.7	40	20
Al-Hudaydah	1	14.3	10	5.2	11	5.5
Ibb	0	0	22	11.4	22	11
Al-Mahwit	0	0	27	14.1	27	13.5
Rima	1	14.3	8	4.1	9	4.5
Sa'da	0	0	1	0.5	1	0.5
Sana'a	1	14.3	24	12.4	25	12.5
Taiz	1	14.3	9	4.7	10	5
Dhamar	0	0	18	9.3	18	9
Total	7	100	193	100	200	100

3.4 Distribution of HBsAg positivity according to Clinical examination:

The rates HBV viral infection in thalassemia patients according to clinical examination were 7 (3.5%) pallor, 2 (1%) splenectomy, 3 (1.5%) jaundice, 1 (0.5%) splenomegaly, 5(2.5%) Iron overload and 2(1%) enlarged Spleen, **Table 3**.

Table 3: Distribution of HBsAg positivity according to Clinical examination

	HBsAg						P. value
	Positive (n=7)		Negative (n=193)		Total		
	N	%	N	%	N	%	
Pallor							
Yes	7	3.5	169	84.5	176	88	X2 = 0.989 P= 0.320 (Non-Sig.)
No	0	0	24	12	24	12	
Total	7	3.5	193	96.5	200	100	
Jaundice							
Yes	3	1.5	18	9	21	10.5	X2 = 8.08 P= 0.004 (Sig.)
No	4	2	175	87.5	179	89.5	
Total	7	3.5	193	96.5	200	100	
Splenomegaly							
Yes	1	0.5	0	0	1	0.5	X2 = 27.7 P= 0.01 (Sig.)
No	6	3	193	96.5	199	99.5	
Total	7	3.5	193	96.5	200	100	
Splenectomy							
Yes	2	1	21	10.5	23	11.5	X2 = 2.07 P= 0.150 (Non-Sig)
No	5	2.5	172	86	177	88.5	
Total	7	3.5	193	96.5	200	100	
Iron overload							
Yes	5	2.5	179	89.5	184	92	X2= 15.10 P= 0.035 (Sig.)
No	2	1	14	7	16	8	
Total	7	3.5	193	96.5	200	100	
Enlarged Spleen							
Yes	5	2.5	97	48.5	102	51	X2= 4.10 P= 0.052 (Non-Sig)
No	2	1	96	48	98	49	
Total	7	3.5	193	96.5	200	100	

3.5 Distribution of HBsAg positivity according to their laboratory tests:

The distribution was not significantly association with Ferritin level, less than 1000 µg/L group was 0 (0%) in positive cases compared to 5 (2.5%) cases in negative cases. While Ferritin level more than 1000 µg/L group was 7 (3.5%) in positive cases compared to 188 (94%) cases in negative cases, (P = 0.666). There was highest significantly (P= 0.001) association with the Alanine transaminase (ALT) level in the Normal range: 0-40 U/L were 0 (0%) in positive cases compared to 121 (60.5%) cases in negative cases. While the high level group (> 45 U/L) were 7 (3.5%) in positive cases compared to 72

(36%) cases in negative cases.

Table 4: Distribution of HBsAg positivity according to their laboratory tests

	HBsAg				Total		P value
	Positive (n=7)		Negative (n=193)		N	%	
	N	%	N	%			
Ferritin level (µg/L)							
(<1000 µg/L)	0	0	5	2.5	5	2.5	X2 = 0.186 P= 0.666
(>1000 µg/L)	7	3.5	188	94	195	97.5	
Total	7	3.5	193	96.5	200	100	
Alanine transaminase (ALT) (U/L)							
< 45 U/L	0	0	121	60.5	121	60.5	X2 = 11.1 P = 0.001
> 45 U/L	7	3.5	72	36	79	39.5	
Total	7	3.5	193	96.5	200	100	

4. Discussion

The results obtained in this study indicated that HBsAg positive were 7(3.5%) of the samples studied (n= 200). Another study in Egypt reported 4.21% were infected with hepatitis B virus [6]. Another studies (29%) and (32.5%) were reported positive for HBsAg among thalassemia patients in Egypt [7, 8]. and Several studies such as in Palestine 23.3% [9], Pakistan 7.4% [10]. Another study was conducted in which total 32 of Thalassemia Patients in West North of Iran observed All of them Patients were seronegative for HBs antigen [4].

Showed similar results for the prevalence of hepatitis virus among thalassemia patients in a global countries such as in Pakistan 3.13% [11]. and 3.69% in West Bengal [12]. Another studies reported 6.4% in Taiwan [13] and 6.31% in Pakistan [14]. Low incidence of HBsAg (1.5%) was found among thalassemia patients in India [15]. In our study, the average age of patient was divided in two groups, ≤ 10 years age group was 103(51.5%) and > than 10 years age group was 87(48.5%). In the present study, it was found that there high significant different between the prevalence of HBV infection in relation to age group. The highest incidence was 6 (3%) observed among > than 10 years age group, followed 1(0.5%) among ≤ 10 years age group, X2 = Chi-Square Value =4.140, P. value = 0.042 and OR= 8.88.Seronegative for HBsAg in 0–5 years age group while 3 (7.7%) in 5–10 years age group were positive for HBsAg and 1 (4.0%) were positive in 10–15 years age group reported by Jaiswal *et al* [16]. Another studies reported 0/27 children aged 1–5 years and 1/59 children aged 6–10 years [17]. In the current study there was significant association between HBsAg positive and more than 100 times of blood transfusion, X² = 5.71. P = 0.022 and OR= 6.8, this study a similar finding Egyptian study was reported by [6]. this is due to non- compliance with international standards and protocols in blood

transfusion procedures. In the present study, there was no significant association between HBsAg positive and gender 1% in male 2.5% in female ($X^2 = 2.13$ P value = 0.144). In the current study there was no significantly association different between HBsAg positive and some clinical examination such as pallor 7 (3.5%) $P = 0.320$ and Splenectomy was 2 (1%) $P = 0.150$, While a significantly associated with jaundice 3 (1.5%) $P = 0.004$, high significantly associated with Splenomegaly, 1 (0.5%) and P value = 0.01. **Ramadan et al**, found that no significant association between HBsAg seropositive and Splenectomy [6]. In the current study there were a significant association between HBsAg seropositive and iron overload were detected 5(2.5%), while there was no significant with enlarged spleen were detected 5(2.5) P value =0.052. There was elevated of serum Alanine transaminase level (ALT) was 7 (100%) in HBsAg positive cases P value = 0.001, this investigation findings conducted by *Ramadan et al* [6].

5. Conclusions

According to present study the following facts can be concluded: low rate for HBsAg Positive sign among beta thalassemia patients that attending at the YST and these seropositive cases were significantly associated with older age, Multi blood transfusion, some clinical examination such as (jaundice, iron overload and Splenomegaly) and elevated of serum Alanine transaminase level (ALT). The risk factors associated with old age and frequency blood transfusion.

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مقالة بحثية

الانتشار المصلي لفيروس الكبد بي بين مرضى النقل المتكرر للدم بيتا ثلاسيميا المترددين على الجمعية اليمنية للتلاسيميا، صنعاء – اليمن

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المُلخَص

يعتبر فيروس الكبد البائي من أبرز الملوثات لمرضى التلاسيميا خاصة أثناء عملية نقل الدم، وتعتبر عملية نقل الدم لمرضى التلاسيميا عملية مستمرة ومنتظمة مدى الحياة، مما يجعلهم عرضة للإصابة بفيروس الكبد البائي (HBV)، خاصة إذا كانت إجراءات النقل تتم بصورة غير آمنة ودقيقة مما قد يؤدي إلى مضاعفات خطيرة قد تؤدي إلى الوفاة، يعتبر بيتا ثلاسيميا (Beta-Thalassemia) أحد أهم أنواع التلاسيميا وأكثرها انتشاراً في دول البحر المتوسط وغرب اسيا وهو مرض وراثي ناتج عن خلل في السلسلة (B) للهيموجلوبين، يؤدي إلى انيميا حادة. تهدف الدراسة إلى دراسة معدل الانتشار المصلي لفيروس الكبد وعوامل الخطورة بين مرضى بيتا ثلاسيميا المترددين على الجمعية اليمنية للتلاسيميا وأمراض الدم الوراثية – صنعاء، وتشمل الدراسة عينات من 12 محافظة يمنية نظراً لارتباط المرضى بالخدمات المقدمة لهم من الجمعية. أجريت هذه الدراسة خلال الفترة من يناير 2021 إلى ديسمبر 2021 تم خلالها جمع 200 عينة من مرضى التلاسيميا المترددين على الجمعية اليمنية للتلاسيميا وأمراض الدم الوراثي في العاصمة صنعاء. وقد تم فحص جميع العينات للبحث عن HBsAg باستخدام تقنية الفحص المناعي التآلفي (ECLIA)، تم خلالها جمع المعلومات الأساسية من خلال استبيان صمم لهذا الغرض. وقد أظهرت نتائج الدراسة إلى تحديد نسبة الانتشار للمستضدات السطحية لفيروس الكبد البائي HBsAg في جميع عينات الدراسات بنسبة 3.5%. كما أظهرت الدراسة أن هناك دلالة إحصائية بين الإصابة بفيروس الكبد والفئة العمرية (أكثر أو يساوي عشر سنوات) $P = 0.042$ ، ودلالة ذات تأثير معنوي بين معدل الإصابة بالفيروس وعدد مرات نقل الدم لمرضى التلاسيميا في اليمن $P\text{-Value} = 0.022$. إضافة إلى ذلك لم تسجل دلالة إحصائية بين نسبة انتشار الفيروس والجنس. أظهرت نتائج هذه الدراسة ارتفاع معدل الإصابة في المرضى المصابون بتضخم في الطحال $p = 0.001$ وكذلك المرضى المصابون باليرقان $P = 0.004$ ، كما أظهرت نتائج الدراسة أن هناك دلالة إحصائية بين معدل الانتشار والأمراض الأخرى التي سجلت في المرضى مثل الإصابة بالسرطان، السكري، وغيرها $P = 0.035$. أظهرت الدراسة ارتفاع نسبة الانتشار في الأشخاص الذين يعانون من ارتفاع في نسبة انزيم الكبد (ALT) بدلالة إحصائية معنوية $P = 0.001$. أظهرت نتائج التحليل الإحصائي عدم وجود علاقة بين الإصابة بالفيروس ونوع الزمرة الدموية وعامل رابيس $P = 0.084$.

الكلمات المفتاحية: مرضى بيتا ثلاسيميا، فيروس الكبد البائي، الجمعية اليمنية للتلاسيميا.

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