# **Pharmacophore**

ISSN-2229-5402

Journal home page: <u>http://www.pharmacophorejournal.com</u>



# THE PREVALENCE OF ANTI HCV INFECTION AND ITS RELATED FACTORS IN PATIENTS WITH BETA-THALASSEMIA IN SHIRAZ-IRAN

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#### ARTICLE INFO

Received: 22th Sep 2017 Received in revised form: 14th Jan 2018 Accepted: 23th Jan 2018 Available online: 28th Feb 2018

Keywords: Beta-thalassemia, HCV infection, Retrospective Cohort.

## ABSTRACT

Purpose: Hepatitis C virus (HCV) is a serious public health problem around the world. Patients with major thalassemia are at high risk of hepatitis C due to the blood transfusion from donors infected by HCV. The aim of this study was detecting the prevalence of HCV and its related factors in Shiraz-Iran.

Material and Methods: This Retrospective Cohort study was conducted in Shiraz City, the south of Iran, from 2004 to 2014 enrolling 839 beta-thalassemic patients. Data were collected from the patient's medical records and they were analyzed using multiple logistic regression. Results: The prevalence of HCV was estimated 4.4% among thalassemic patients. The mean age of patients was 22.87(8.11). 23(5.5%) and 14(3.3%) of beta thalassemic patients which were infected by HCV were male and female respectively. Age, type of thalassemia and heart complications had a significant association with HCV seropositivity in multiple logistic regression models.

Conclusions: Our findings revealed that Age, type of thalassemia and heart complications were the main risk factors for HCV infection among beta-thalassemic patients. Due to the increase of patients survival and since the HCV is related to the other complications, we must use some methods such as weekly cares, fresh blood transfusions to patients and timely and early hemoglobin treatment to reach to decrease and delay the effect of this disease in patients.

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**To Cite This Article:** Mostafa Sadeghi, Maryam Soltani, Koorosh Etemad, Maliheh Abdollahi, Mohammad Sayyadi, Mohyedin Barzegar, et al, (2018), "The prevalence of anti HCV infection and its related factors in patients with Beta-Thalassemia in Shiraz-Iran", *Pharmacophore*, *9(1)*, *80-84*.

#### Introduction

Beta thalassemic patients are exposed to hundred or thousand blood of donors during their lives, because they need blood products repeatedly. So they are at high risk of blood-borne disease such as HIV and viral hepatitis. HCV virus is one of the

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#### Pharmacophore, 9(1) 2018, Pages 80-84

blood-borne pathogen associated with multiple complications and mortality [1-2]. The studies have showen more than 50% of major thalassemic patients would be infected by a type of hepatitis [3-4]. The infectious after blood transfusion is still one of the most principal challenges in human society despite all of the progresses of human knowledge. All the efforts for HCV prevention and treatment lead to decrease transmission of the hepatitis significantly [5-6-7]. The studies have shown the prevalence of the infectious depends on the first time they ever used blood products [8-9]. Iran is situated in the middle of thalassemia belt and has a high frequency rate for thalassemia carriers [10]. Iran has more than 2000 patients with thalassemia already [11]. According to different researches, the prevalence of HCV in beta thalassemic patients is 20%-30% in our country [12-13]. Beta thalassemic patients are at high risk of liver fibrosis because of a high range of hemoglobin in liver, further their livers are more vulnerable than non-thalassemic patients. Moreover, it must be more dangerous for them to be infected by HCV and it will cause serious damages such as Hepatocellular Carsinoma [6]. Considering that no study has been done about the prevalence of HCV in Fars yet, it can examine the effects of factors related to thalasemic patients which are infected by HCV. By recognizing these factors we can improve blood transfusion centers and try to minimize the complications associated with blood and blood products. In this study, the prevalence of HCV and its related factors have been examined in Fars Province.

# **Materials and Methods**

This Retrospective Cohort study was conducted in Shiraz City, the south of Iran, from 2004 to 2014 enrolling 839 betathalassemia patients. The data of all patients based on census method were cross-sectionally extracted from patients' medical records using a predetermined checklist including age (calendar age at the study enrollment time), sex, education level, marital status (single or married), consanguinity, ferritin & hemoglobin level, blood transfusion, type of beta-thalassemia (major or intermedia), and accompanied diseases (Hepatitis C, Diabetes type 2, Spelenectomy, Hypogonadism, Thyroidism). Ferritin and hemoglobin levels are measured every six months and once a month, respectively. The average level of the last four times (except for two patients) was considered as the level of ferritin and hemoglobin in the analysis. Statistical Analysis

We used mean and standard deviation and relative and absolute frequency to analyze quantitative and qualitative variables, respectively. They were used to find significant associations between patient's characteristics and HCV positivity. Multiple logistic regression models were applied to estimate adjusted odds ratio and 95% confidence interval for risk factors of HCV positivity. Statistical tests were conducted at the P < 0.05 significance level. The collected data were coded and entered in a data base file. After complete entry, data were transferred to the SPSS software version 16 (SPSS Inc., Chicago, IL). This study has been approved in ethical committee of Shiraz University of Medical Sciences. All information from medical

# Results

records of patients is kept confidentially.

Eight hundred and thirty nine patients with thalassemia enrolled in this study. 37 of participants (4.4%) were infected by HCV .23(5.5%) and 14(3.3%) of them were male and female respectively. The mean age of these patients were 22.87(8.11). Out of all 37 patients, 30(5.2%) had major thalassemia and 7(2.9%) had intermedia thalassemia. Also 18 patients who had consanguineous marriage were infected by HCV. 18 patients with education level higher than diploma and 16 ones with diploma and lower than diploma [**Table 1**]. In beta thalassemic patients, the frequency of patients with Splenectomy was 386(43.5%), patients with heart complications was 170(20.3%), patients with diabetes was 125(14.9) and in patients with Osteoporosis was 313(37.3) and at last in patients with Parathyroid complications was 232(27.3).

-	Total	HCV			
		positive	negative		
Gender (n=839)					
Female	421(50.2)	14(3.3)	407(96.7)		
Male	418(49.8)	23(5.5)	395(94.5)		
Marital Status (n=835)					
Single	742(88.9)	34(4.4)	708(75.4)		
Married	93(11.1)	2(2.2)	91(97.8)		
Education Level(n=808)					
Diploma and lower than diploma	325(40.2)	16(4.9)	309(95.1)		
Higher than diploma	483(59.8)	18(3.7)	465(96.3)		
Type of Thalassemia(n=826)					
Major	581(70.3)	30(5.2)	551(94.8)		
Intermedia	425(29.7)	7(2.9)	238(97.1)		
Type of Parents Marriage (n=833)					
Consanguineous	420(50.4)	18(4.3)	403(95.7)		
Non-consanguineous	413(49.6)	18(4.4)	395(95.6)		

Pharmacophore, 9(1) 2018, Pages 80-84

The crude analysis of variables related to HCV has been shown in **[Table 2]**. The results showed the risk of HCV increases 9% by aging [OR1.09,95% CI(1.05-1.15)] which is statistically significant. The risk of HCV in patients with some complications is more than other patients who have not these complications. For example, in patients with splenectomy was 4times more and patients with heart disease was 10 times more [OR:3.73,95%CI(1.82-7.66)]. Also in patients with diabetes was 2.5 times more [OR:2.55,95%CI(1.23-5.31)] and 3 times more in patients with thyroid complications[OR:3.73,95%CI(1.82-7.66)]. In patients with Hypogonadism 2.5 times more than the patients who have not this complications [OR:3.73,95%CI(1.82-7.66)]. All of these results were statistically significant as well. The risk of HCV in men was 69% more than women and 2 times more in single compared with married ones. But these were not statistically significant. This risk in high educated people was 34% more than low educated ones. Also, patients with a hemoglobin level of greater than 9.5 g/dl (33%) had a lower risk of HCV positivity in comparison with patients who had a greater hemoglobin level and in patients who had serum ferritin levels less than (2500 ng/ml) were 34% less than those who had higher ferritin levels. But also these were not statistically significant.

	Hepatitis	
	OR(95% CI)	Pvalue
Age(years)		0.001**
Sex		
Female	Reference	Reference
Male	1.69(0.85-3.37)	0.128
	Married status	·
Married	Reference	Reference
Single	2.19(0.52-9.25)	0.288
	Education	·
Higher than diploma	Reference	Reference
Diploma and lower than diploma	1.34(0.67-2.67)	0.408
	Type of thalassemia	
Intermedia	Reference	Reference
Major	1.85(0.80-4.27)	0.149
	Hemoglobin Level(g/dl)	
>9	Reference	Reference
<=9	1.80(0.91-3.52)	0.090
	Ferretin Level(ng/mL)	
More than 2500	Reference	Reference
Less than 2500	0.57(0.29-1.12)	0.103
	SPELENECTOMY	
No	Reference	Reference
Yes	3.73(1.82-7.66)	0.001**
	DIABET	
No	Reference	Reference
Yes	2.55(1.23-5.31)	0.012
	Thyroid complications	
No	Reference Reference	
Yes	3.36(1.27-9.07)	0.018
	HYPOGONADISM	
No	Reference Reference	
Yes	2.44(1.26-4.74)	0.008

Table 2. Crude Odds Ratio (OR) estimating the factors associated with Thalassemia patients with HCV

#### Pharmacophore, 9(1) 2018, Pages 80-84

Adjusted multiple logistic regression about related factors of HCV in patients with thalassemia have been shown in **[Table 3]**. The risk of HCV infection in major thlassemic patients was 3 times more than patient with intermedia thalassemia [OR:3.09,95%CI(1.19-8.03)]. Also it was 10 times more in patients with heart complications in comparison with those who had not such complications.

	Hepatitis			
	OR(95% CI)	Pvalue		
Age(years)	1.09(1.05-1.15)	0.001**		
Type of thalassemia				
Intermedia	Reference	Reference		
Major	3.09(1.19-8.03)	0.021		
Heart Attack				
No	Reference	Reference		
Yes	4.57(2.25-9.29)	0.001**		

Table 3. Adjusted Odds Ratio (OR) estimating the factors associated with Thalassemia patients with HCV

#### Discussion

Among all of the blood-borne infections, HCV is the most important one. The prevalence of HCV all over the world is 3% [9]. The screening tests of HCV were discovered in 1990. After 1992 the screening test on blood bags became mandatory. Nowadays the risk of transmission of HCV decreases to 1:100000 by the help of the third generation kits [14]. In Iran HCV Ab tests were conducted on blood donors by first generation of ELISA (enzyme linked immunosorbant assay) method and by since 1375 and by third generation of ELISA method since 1377 [15].

This Retrospective Cohort study were conducted on 911 beta thalasse, ic patients in Shiraz from 2004 to 2014. In this study the prevalence of HCV in beta thalassemic patients was 4.4% .The former researches reported that the prevalence of HCV was 7-64% in Iran [16].For example, 28.1% in Khuzestan [17], 64% in Rasht [18], 25% in Shiraz [19], 8% in Isfahan [20] and 13% in Sistan and Baluchestan as well [21]. The reason of such this wide range is the difference between type and sensitivity of diagnostic tests. In Arabic countries the prevalence of HCV in beta thalassemic patients was in a range from 33% to 67.3% [22-23]. A study was conducted in Pakistan which includes 180 children with major thalassemia from January 2002 to December 2003. Out of all 180 children, 75 of them were HCV infected [24]. In another study in India, the prevalence of HCV was 23.1% in beta thalassemic patients [25].

This study also evaluated the risk factors of HCV in beta thalssemic patients. The results of multiple logistic regression modeling have shown that age, type of thalassemia and heart complication have a significant correlation with HCV positivity. The findings proved that the risk of HCV will be increased about 9% by aging. Patients who were infected by HCV not only are at risk of Cirrhosis and Hepatocellular Carsinoma but also have other comorbidities disease included renal diseases, hematological, rheumatic and dermatological conditions [26]. It has been reported that HCV infection cab increase the risk of Atherosclerosis and Coronary artery diseases [27-28]. The findings of this study correspond to the findings of Roed study which shows the increase of Cardiovascular disease risk in patients with HCV [29].

#### Conclusions

Our findings revealed that Age, type of thalassemia and heart complications were the main risk factors for HCV infection among beta-thalassemic patients. Due to the increase of patients survival and since the HCV is related to the other complications, we must use some methods such as weekly cares, fresh blood transfusions to patients and timely and early hemoglobin treatment to reach to decrease and delay the effect of this disease in patients.

#### Acknowledgment

We would like to thank the Vic-chancellor of Health of Shiraz University of Medical Sciences for technical support of this study. This study has been approved by Fasa University of Medical Sciences (No. 93188). Also, the authors are thankful to the director and all the employees of the Dastgheyb Hospital, especially Ms. Lotfi who helped us in performing this project.

# **Conflict of interest**

The Authors have no conflict of interest.

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## Mostafa Sadeghi et al, 2018

#### Pharmacophore, 9(1) 2018, Pages 80-84

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