Original Research



Seroprevalence of hepatitis B and C viral infections in the premarital adult population of Al Majmaah, Saudi Arabia

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Abstract

Background

Viral hepatitis is a major global health problem affecting millions of people worldwide. The main objective of the present study was to estimate the seroprevalence of hepatitis C and hepatitis B in Saudi adults undertaking premarital screening voluntarily.

Methodology

This observational retrospective study was conducted at King Khaled General Hospital, Al Majmaah, Saudi Arabia, for a period of 3 years (October 1, 2016 to September 30, 2019). The prevalence of hepatitis B and hepatitis C infections was calculated using Microsoft Excel and the hematological data analysis was performed using SPSS software. A total of 3755 premarital Saudi adults who voluntarily participated in the premarital screening were included in this study. Subjects were screened for hepatitis C virus (HCV) and hepatitis B virus (HBV). The demographic and epidemiological parameters included patient age, sex, nationality, and seropositivity for HBV and/or HCV.

Results

Of the 3755 subjects, eight (0.242%) and 12 (0.364%) subjects were positive for HCV and HBV, respectively. Among the 12 HBV-positive subjects, nine (75%) subjects were men, and three (25%) subjects were women. However, among the eight HCV-positive subjects, five (62.5%) subjects were men, and three (37.5%) subjects were women.

Conclusion

This study concluded that the prevalence of HBV infection was slightly higher than that of HCV infection among the Saudi adult population of Al Mjamaah. In addition, both HBV and HCV were less prevalent in women than in men in the study population.

Keywords: Seroprevalence, Hepatitis, Hepatitis C virus, Hepatitis B virus, Infection

Introduction

Viral hepatitis being a major global health problem affects approximately 400 million people globally and is responsible for about 1.4 million deaths each year, most (95%) of which are due to hepatitis B and hepatitis C1, 2. Viral hepatitis is caused by five different types of viruses, including hepatitis virus A, B, C, D and E. The hepatitis B virus (HBV) and hepatitis C virus (HCV) can cause chronic hepatitis^{3, 4}. HBV and HCV are transmitted through exposure to infected blood and/or semen. More than 95% of the hepatitis B infected adults resolve the infection during the acute phase. However, about 80% of the patients with hepatitis C progress to chronic infection^{1, 2, 4}. Viral hepatitis is mostly diagnosed by testing the virus specific serological markers. The diagnosis of HBV infection requires the detection of few serological markers, including anti-HBs/anti-HBc antibodies, and HBsAg. HBsAg is the serological hallmark of infection as it can be detected within 1-10 weeks postinfection and its persistence for >6 months is indicative of chronic infection. HCV screening is generally performed using anti-HCV antibodies and is occasionally confirmed by HCV RNA test4

Hepatitis B is the most widespread viral hepatitis, with approximately 57 million people infected with HBV worldwide⁵. Its prevalence is higher in the Middle East than in Europe and the United States. Its prevalence ranges

from 0.6% in Iraq to >8% in Sudan⁶. The number of HBVinfected persons in the Eastern Mediterranean region is >21 million (3.3% of the population). Hepatitis C affects approximately 3% of the world population; however, it has a high prevalence of approximately 10% in developing countries³. In 2007, viral hepatitis was ranked as the second most common viral disease in Saudi Arabia⁷. Previously, in 1998, Saudi Arabia witnessed a high prevalence (8.3%) of hepatitis B8. In a community based epidemiological study HBsAg seroprevalence of approximately 7% was reported in Saudi children. These reports of high prevalence of HBV led to state and scientific intervention, and as a result, the universal HBV immunization was started in the country in 19899. The immunization program successfully resulted in a decrease in the prevalence of HBV in the country¹⁰. Data on the prevalence of HCV in Saudi Arabia are insufficient and controversial. The premarital screening program estimates the prevalence of HCV to be 0.33%, while screening centers for blood donors indicate HCV infection rates of 0.4%-1.1%¹¹⁻¹⁵. In addition, HCV prevalence of 1.0 %–1.9% has been reported in a systematic review that included most of the published reports on the subject¹⁶.

The insufficiency of confirmed data leads to diminished planning for the prevention and treatment strategies of the disease¹⁷. Hence, there is a need for considerable community-based epidemiological studies to obtain more accurate incidence rate of HCV and HBV infection in Saudi Arabia.

Therefore, the present study was performed to estimate the prevalence of HBV and HCV infection among adult population of Al Majmaah, Saudi Arabia. To accomplish this study, adults who attended the health service at King Khaled General Hospital (KKGH) for premarital screening were included and examined. The study population included adult men and women of Saudi nationality from Al Majmaah. In addition, the effects of HBV and HCV seropositivity on hematological parameters were also evaluated.

Materials and Methods

Source/study population

The present study was conducted at the premarital screening center of KKGH and involved 3755 adult Saudi people from the Al Majmaah region (1953 men and 1802 women, aged 20-55 years). The 3-year clinical data (October 1, 2016 to

September 30, 2019) were analyzed using a statistical analysis system. The study was approved by Central Institutional Review Board, Ministry of Health, Saudi Arabia (IRB No. 2019-0126E).

Saudi nationals visiting the health service for premarital screening with no recent illness, surgery, transfusion blood or blood components were included in the study. Individuals underwent premarital

7 **o** 3 Oct 2016 to Sep 2017

screening with a recent history of health issues, major surgery, or transfusion of blood and/or blood components in the previous 6 months were excluded from the study.

Blood sampling

Fresh whole blood (5 mL) was collected from all the study subjects and divided into two parts; one part was transferred to a tube containing ethylene diamine tetra acetic acid (EDTA), mixed gently and stored at ambient temperature and analyzed for hematological parameters within 48h of collection. The other part was transferred into a plain tube (without any anticoagulant) and allowed to clot by leaving it undisturbed at room temperature for 45-60 min. The clotted blood samples were centrifuged for 10 min at 1000g in a refrigerated centrifuge and the resulting supernatant (serum) from each sample was collected into separately labelled clean and sterile tubes and stored at -20°C until analysis.

Serological tests for HCV and HBV

HCV and HBV serological markers were identified and evaluated using the ARCHITECT Immunoassay System (Abbott, Abbott Park, Illinois, USA) according to the manufacturer's instructions. The detection of HBV surface antigen (HBsAg) and anti-HCV antibodies was performed using HBsAg ULTRA kit (Bio-Rad, France) and anti-HCV PLUS version 3 kit (Bio-Rad, France), respectively. The initial screening results were confirmed by retesting positive samples18.

Hematological analysis

Hematological tests for healthy normal individuals and HBVand HCV-positive individuals were performed using an automated hematology analyzer, CELL-Dyn Ruby (Abbott Laboratories, Diagnostic Division, Abbott Park, Illinois, USA)

Statistical Analysis

Statistical analysis for estimating the prevalence of HBV and HCV was performed using Microsoft Excel. The hematological data was analyzed by one-way ANOVA test using SPSS software. The results were considered statistically significant if p values were <0.05.

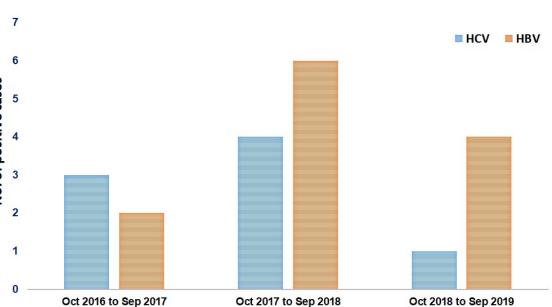


Fig. 1: Yearly distribution of the positive cases of HBV and HCV in the study population

Results

Seroprevalence of HBV and HCV

In total, 3755 participants (1953 [52.01%] men and 1802 [47.99%] women) were recruited from October 1, 2016 to September 30, 2019. All participants were Saudi nationals aged 20-55 years. All participants underwent premarital screening and were examined for HBsAg and anti-HCV serum markers. As shown in Table 1, 0.32% (n = 12) and 0.21% (n = 8) subjects were positive for HBsAg and anti-HCV, respectively, and 99.47% (n = 3735) subjects were negative for both the serum markers tested. The yearly distribution of HBV- and HCV-positive cases in the study population is shown in Figure 1.

On further analysis of our data and results, we observed that among the 1953 men, 99.28% (n = 1939) were seronegative, 0.26% (n = 5) were positive for HCV, and 0.46% (n = 9) were positive for HBV (Table 2). Among the 1802 women, 99.67% (n = 1796) were seronegative, 0.17% (n = 3) were positive for HCV, and 0.17% (n = 3) were positive for HBV (Table 2). Additionally, based on their age, we grouped the HBV and HCV seropositive men and women into two categories—>30 years old and <30 years old—and estimated the frequency of HBV and HCV in both men and women according to age groups.

Table 1: Prevalence of viral hepatitis among 3755 participants undergoing premarital screening in Al Majmaah, Saudi Arabia (from October 1, 2016 to September 30, 2019)

Category	No. of cases (n)	Percentage (%)
HCV positive	8	0.21
HBV positive	12	0.32
HCV and HBV Negative	3735	99.47
Total	3755	100

Table 2: Combined results of HCV and HBV positive male and female cases/groups.

Total study (N=3755)	al study population 3755)		HBV +ive cases (n=12)		HCV +ive cases (n=8)	
Male,	Female,	Male,	Female,	Male,	Female,	
n (%)	n (%)	n (%)	n (%)	n (%)	n (%)	
1 9 5 3 (52.01)	1 8 0 2 (47.99)	9 (0.46)	3 (0.17)	5 (0.26)	3 (0.17)	
		>30 yrs = 7	>30 yrs = 1	>30 yrs = 4	>30 yrs = 1	
		< 30 yrs = 2	< 30 yrs = 2	< 30 yrs = 1	< 30 yrs = 2	

Table 3: Comparison of hematological parameters between healthy controls, HBV positive and HCV positive individuals

Hematologic parameters	Healthy controls	HBV positive	HCV positive
RBC (10 ⁶ /mm ³)	5.121 ± 0.24	4.95 ± 0.45	4.86 ± 0.45
HB (g/dL)	14.912 ± 0.52	14.69 ± 0.88	14.36 ± 1.12
HCT (%)	45.463 ± 4.09	45.98 ± 3.09	42.94 ± 2.8
MCV (fL)	86.556 ± 3.66	89.55 ± 4.31	88.62 ± 4.7
MCH (pg)	29.273 ± 0.75	30.47 ± 0.93 ***	29.56 ± 0.41
MCHC (g/dL)	34.58 ± 0.86	35.26 ± 0.74	34.58 ± 0.43
RDW	12.844 ± 0.86	12.13 ± 0.51 *	12.16 ± 0.56

Values are shown as Mean ± SD. Hb: Hemoglobin, RBC: Red blood cell, HCT: hematocrit, MCH: mean cell of HCV to be 0.1%, 3%, homoglobin MCHO: mean cell of HCV to be 0.1%, 3%, hemoglobin, MCHC: mean cell hemoglobin concentration, MCV: mean cell volume, RBC: red blood cell, RDW: and 0.83% in 2008, 2009, and cell distribution with red cell distribution width.

Among the nine HBV-positive men, two were <30 years old and seven were >30 years old. However, in three HBVpositive women, one was >30 years old, and two were <30 years old (Table 2). Similarly, among the five HCV-positive men, four were >30 years old and one was <30 years old, and among the three HCV-positive women, two were <30 years old and one was >30 years old (Table 2). However, no cases of HBV/HCV co-infection were identified in this study.

Hematological findings

As shown in Table 3, the analyzed hematological parameters, including hemoglobin levels, total RBC count, hematocrit, mean cell volume and mean cell hemoglobin concentration, did not show any statistically significant differences between HBV- or HCV-seropositive and seronegative individuals. However, a statistically significant difference was observed in the values of mean cell hemoglobin (MCH) and red cell distribution width (RDW) of the HBV-seropositive subjects compared to those of the seronegative ones (p < 0.005).

Discussion

Premarital screening is particularly important because it can prevent diseases in the future generations and provide insight into the incidence of several diseases in the adult population¹⁹. Premarital screening is mandatory in Saudi Arabia and provides an excellent platform to estimate the prevalence of several diseases²⁰. In Saudi Arabia, consanguineous marriages are the main factor for births with a high risk of carrying a disease, especially viral hepatitis and hemoglobinopathies¹⁹. As there is a considerable incidence of hepatitis in Saudi Arabia, it is recommended to acquire data on the prevalence of the disease for planning/improving its prevention and treatment strategies. However, the relevant studies reported in the literature are insufficient to acquire complete data. In this regard, we have previously reported the prevalence of various transfusion-transmitted diseases, including HCV and HBV, in blood donors of Al Majmaah, Saudi Arabia²¹. In the present study, we have reported the prevalence of HCV and HBV in adult persons who underwent premarital screening at KKGH, Al Majmaah.

The study results showed that seroprevalence of HBV and HCV was higher in men (0.46% and 0.26%, respectively) than in women (0.17% and 0.17%, respectively). The prevalence of HBV and HCV infection in couples tested through

> the premarital screening program in a previous similar study was 1.31% and 0.33%, respectively, which is slightly higher than that reported in our results15. Furthermore, a study conducted in the Qassim region showed that the positivity for HBV was 0.7%, 1.5%, and 2.04% in 2008, 2009, and 2010, respectively. The same study also reported the prevalence

> and 2010, respectively²².

The differences in the prevalence rates might be due to regional differences, as the studies were performed

in different regions/cities of Saudi Arabia. Furthermore, in these studies, the prevalence of HBV infection was higher than that of HCV infection, which is consistent with the results of our study; however, the prevalence values reported in our study are lower than those reported in previous studies. Several studies have reported a decrease in the incidence and prevalence of viral hepatitis B and C in Saudi Arabia over time^{18,23}. Correspondingly, our results also revealed a lower prevalence of HCV and HBV infection in the adult population. This decline in the prevalence of HBV and HCV may be due to many factors, including timely active vaccination programs, mandatory premarital screening, and complementary immunological and therapeutic approaches that started in Saudi Arabia in 1989.

In agreement with previous studies, our results confirmed that HBV infection was more prevalent than HCV infection in our study population. Further analysis of seropositive results showed that the prevalence of HBV and HCV infection was lower in women than in men, consistent

^{*} Significant difference: * p<0.05 and *** p<0.001

with previous studies conducted in the Riyadh region^{24,25}. Hematological analysis revealed that the slight differences in the hematological parameter between the HBV- or HCVseropositive subjects and the seronegative individuals were statistically insignificant, except for MCH and RDW, that too in case of HBV only (Table 3). Some of our hematological findings are in agreement with those of Ahmad et al (2018) and Sabry et al (2007)^{26,27}, but not with some other findings available in literature^{28,29}. Our main focus was to estimate the prevalence of HBV and HBC, and the small sample size of HBV- and HCV-seropositive groups could be the reason for the some controversial results.

Conclusions

This study reports the prevalence of HCV and HBV infection in the adult population of Al Majmaah, emphasizing a higher prevalence of HBV infection than that of HCV infection and a higher prevalence of both diseases in men than in women.

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Both the authors (SAM and BA) contributed equally to the study design, data analysis, and writing of the manuscript. The authors are very thankful to the staff of the premarital screening center of KKGH, Al Majmaah, particularly Mr. Mohaammad Khaled Alturaiki, for providing the data. We also thank the Deanship of Scientific Research, Majmaah University, for their support in proofreading the manuscript on time.

Ethical Approval

All procedures in this study were performed in accordance with the ethical standards of the Central Institutional Review Board (IRB) of the Ministry of Health, Saudi Arabia (central IRB log number 2019-0126E).

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Conflicts of interest

The authors have no conflicts of interest

References

- 1. WHO. Towards ending viral hepatitis. Global health sector strategy on viral hepatitis 2016-2021. (Accessed on February 2, 2020, at http:// apps.who.int/iris/bitstream/10665/246177/1/WHO-HIV-2016.06-eng. pdf?ua=1)
- 2. Angeleri P, Levite V, Vidiella G, Solari J, Coronel E, Adaszko D, et al. Viral hepatitis and Treponema pallidum prevalence in persons who underwent premarital blood tests in Argentina. Sci Rep 2019;9:9611.
- 3. Rutherford A, Dienstag JL. Viral hepatitis, In: Greenberger NJ, Blumberg RS, Burakoff R editors. Current diagnosis & treatment: gastroenterology, hepatology, & endoscopy. 3rd ed. New York: McGraw-Hill Education; 2016.
- 4. Pondé, RAA. The serological markers of acute infection with hepatitis A, B, C, D, E and G viruses revisited. Arch Virol 2017;162:3587–3602.
- 5. WHO. New hepatitis data highlight need for urgent global response. News room, Geneva: World Health Organization, 2017. (Accessed February 2, 2020, at https://www.who.int/news-room/ detail/21-04-2017-new-hepatitis-data-highlight-need-for-urgentglobal-response)
- 6. Habibzadeh F. Viral hepatitis in the Middle East. Lancet 2014;384:1-

- 7. Ministry of Health of Saudi Arabia. A review of health situations Annual Health Statistics Books Riyadh: Ministry of Health of Saudi Arabia; 2007.
- 8. Al-Faleh FZ. Hepatitis B infection in Saudi Arabia. Ann Saudi Med 1988;8:474-480.
- 9. Al Faleh F, Al Shehri S, Al Ansari S, Jeffri MA, Mazrou YA, Shaffi A, et al. Changing patterns of hepatitis A prevalence within the Saudi population over the last 18 years. World J Gastroenterol 2008;14:7371-
- 10. Abdo AA, Sanai FM, Al Faleh FZ. Epidemiology of viral hepatitis in Saudi Arabia; Are we off the hook? Saudi J Gastroenterol 2012;18:349-
- 11. Mehdi SR, Pophali A, Al-Abdul Rahim KA. Prevalence of hepatitis B and C and blood donors. Saudi Med J 2000;21:942-944.
- 12. Bashawri LAM, Fawaz NA, Ahmad MS, Qadi AA, Almawi WY. Prevalence of seromarkers of HBV and HCV among blood donors in eastern Saudi Arabia, 1998-2001. Clin Lab Haematol 2004;26:225-228.
- 13. El-Hazmi MM. Prevalence of HBV, HCV, HIV-1, 2 and HTLV-I/ II infections among blood donors in a teaching hospital in the Central region of Saudi Arabia. Saudi Med J 2004;25:26-33.
- 13. Madani TA. Hepatitis C virus infections reported in Saudi Arabia over 11 years of surveillance. Ann Saudi Med 2007;27:191-194.
- 15. Alswaidi F, O'Brien S. Is there a need to include HIV HBV and HCV viruses in the Saudi premarital screening program on the basis of their prevalence and transmission risk factors? J Epidemiol Community Health 2010;64:989-997.
- 16. Sievert W, Altraif I, Razavi HA, Abdo A, Ahmed EA, Alomair A, et al. A systematic review of hepatitis C virus epidemiology in Asia, Australia, and Egypt. Liver Int 2011;61–80.
- 17. Alghamdi AS, Alqutub A, Abaalkhail F, Sanai FM, Alghamdi F, Altraif I, et al. SASLT position statement on the direct-acting antiviral agents for the treatment of hepatitis C virus infection. Saudi J Gastroenterol 2015;21:60-63.
- 18. Abdullah SM. Prevalence of hepatitis B and C in donated blood from Jazan region of Saudi Arabia. Malays J Med Sci 2013;20:41-46.
- 19. Memish ZA, Saeedi MY. Six-year outcome of the national premarital screening and genetic counseling program for sickle cell disease and β-thalassemia in Saudi Arabia. Ann Saudi Med 2011;31:229–235.
- 20. Karaosmanoglu HK, Aydin OA, Sandikci S, Yamanlar ER, Nazlican O. Seroprevalence of hepatitis B: do blood donors represent the general population. J Infect Dev Ctries 2012; 6:181-183.
- 21. Alaidarous M, Choudhary RK, Waly MI, Mir S, Dukhyil AB, Banawas SS, et al. The prevalence of transfusion-transmitted infections and nucleic acid testing among blood donors in Majmaah, Saudi Arabia, J Infec and Pub Health 2018;11:702-706.
- 22. Aljarbou AN. Current prevalence of HBV and HCV seropositivity: the initiative for attentiveness and deterrence of viral hepatitis in the Qassim region of Saudi Arabia. J Antivir Antiretrovir 2012;4:75-79.
- 23. Al-Tawfiq JA, Anani A. Profile of viral hepatitis A, B and C in a Saudi Arabian hospital. MedSci Monit 2008;14:52–56.
- 24. Mansoor AA, Salih AI, Al-Jaroudi DH. Screening of hepatitis B and C and human immunodeficiency virus in infertile couples in Saudi Arabia. Saudi Med J 2011;32:260–264.
- 25. Tripodi A, Mannucci PM. The coagulopathy of chronic liver disease. N Engl J Med 2011;365:147-156.
- 26. Ahmad AE, Bakari AG, Musa BOP, Mustapha SK, Nasir AI, Tahir MI, et al. Haematological and Immunological Parameters in Patients with Chronic Hepatitis B Infection in Zaria, Nigeria. Sokoto J Med Lab Sci 2018;3:84-88.
- 27. Sabry A, El-Dahshan K, Mahmoud K, El-Husseini A, Sheashaa H, https://dx.doi.org/10.4314/mmj.v33i3.10

Abo-Zenah H. Effect of hepatitis c virus infection on haematocrit and haemoglobin levels in egyptian hemodialysis patients. Eur J Gen Med 2007;4:9-15

28. Dar MS, Gupta S, Gowhar O. Estimation of hematological parameters in patients with Hepatitis B and C. Int Arch Integr Med 2019; 6: 76-80.

29. Alshebani KE, Jpireal JM. , Shawesh F, Elazomi A. Variation in haematological parameters in patients with chronic hepatitis B and hepatitis C in Libya. Eur J Pharm Med Res 2020;7:158-166.