ARTICLE IN PRESS

Transactions of the Royal Society of Tropical Medicine and Hygiene (2008) xxx, xxx-xxx



Hepatitis C virus infections reported over 11 years of surveillance in Saudi Arabia

Tariq A. Madani^{a,b,*}

⁵ ^a Ministry of Health, Riyadh, Saudi Arabia

6 Q1 ^b Department of Medicine, Faculty of Medicine, King Abdulaziz University Hospital, P.O. Box 80215, Jeddah, Saudi Arabia

7 Received 11 May 2007; received in revised form 1 August 2008; accepted 1 August 2008

8 9 10 11 12 13 14	KEYWORDS Hepatitis C; HCV; Hepatitis C antibodies; Prevalence; Saudi Arabia	Summary This was a case series descriptive study of all subjects reported to the Ministry of Health in Saudi Arabia from January 1995 to December 2005 as having hepatitis C virus (HCV) infection, diagnosed by detection of antibodies to HCV. A total of 24 948 cases were reported, of whom 19 185 (76.9%) cases were Saudis. The number of HCV infections by region ranged from 16 to 322 cases, with a mean of 124 cases per 100 000 population, or 0.124%. The number of cases reported among children <15 years was 998 cases (12 cases per 100 000 pediatric population, or 0.012%), and that among adults was 23 950 cases (202 cases per 100 000 adult population,
11 12 13 14 15 16 17 18 19 20 21	Hepatitis C antibodies; Prevalence; Saudi Arabia	whom 19185 (76.9%) cases were Saudis. The number of HCV infections by region ranged from 16 to 322 cases, with a mean of 124 cases per 100 000 population, or 0.124%. The number of cases reported among children <15 years was 998 cases (12 cases per 100 000 pediatric population, or 0.012%), and that among adults was 23 950 cases (202 cases per 100 000 adult population, or 0.202%). There was a slight steady increase in the annually reported infections from 1995 to 2002, followed by a plateau. The lower number of HCV infections reported in children compared with those reported in adults suggested that perinatal and childhood transmission was not a major mode of infection and that other modes of transmission, such as unscreened blood transfusion before 1990 and intravenous drug use, were likely to be the main modes of infection. The study was limited by being a passive reporting of cases and not a cross-sectional survey. © 2008 Published by Elsevier Ltd on behalf of Royal Society of Tropical Medicine and Hygiene.

1. Introduction

Hepatitis C virus (HCV) infection is an important pub-23 lic health problem worldwide.¹ As of June 1999, WHO 24 estimated that 169.7 million people (3% of the world's pop-25 ulation) were chronically infected with HCV globally, and 26 that three to four million people are newly infected each 27 year.² The prevalence rate was estimated to be 5.3% in Africa 28 (31.9 million cases), 4.6% in the Eastern Mediterranean 29 region (21.3 million cases), 3.9% in the West Pacific region 30

* Tel.: +966 5 55683107/+966 2 6408243 (office); fax: +966 2 6408315 (office). *E-mail address*: taamadani@yahoo.com (62.2 million cases), 2.15% in Southeast Asia (32.3 million cases), 1.7% in the Americas (13.1 million cases) and 1.03% in Europe (8.9 million cases).¹ Data on the prevalence of HCV infection in Saudi Arabia is limited. The objective of this study was to describe the number of HCV infections reported in Saudi Arabia during 11 years of surveillance, from January 1995 to December 2005.

31

32

33

34

35

40

41

2. Materials and methods

2.1. Saudi Arabia

Saudi Arabia occupies most of the Arabian Peninsula, with an area of about 2 240 000 km². It comprises 13 administrative

0035-9203/\$ — see front matter © 2008 Published by Elsevier Ltd on behalf of Royal Society of Tropical Medicine and Hygiene. doi:10.1016/j.trstmh.2008.08.001

Please cite this article in press as: Madani TA. Hepatitis C virus infections reported over 11 years of surveillance in Saudi Arabia. *Trans R Soc Trop Med Hyg* (2008), doi:10.1016/j.trstmh.2008.08.001

2

74

75

76

77

78

79

80

81

82

83

84

85

86

87

88

89

90

91

92

93

94

95

96

97

98

99

100

101

102

Province	No. reported cases	Mean population during the surveillance period	Cases per 100 000 population
Baha	1268	393 327	322
Jeddah	9186	2 866 113	321
Najran	734	356 250	206
Eastern	3522	1 824 952	193
Qunfoda	102	55 725	183
Makkah	2022	1 483 258	136
Riyadh	4159	4 538 346	92
Qassim	812	890 625	91
Bisha	218	288 321	76
Tabook	389	575 000	68
Madina	808	1 283 251	63
Jouf	112	205 882	54
Asir	568	1 297 311	44
Ahsa	351	940 217	37
Qerayyat	33	113 393	29
Hail	143	487 778	29
North borders	78	270 808	29
Tayef	221	883 186	25
Hafr Albaten	62	280 727	22
Jizan	160	1 030 159	16
Total	24948	20 064 629	2036

provinces: namely, Makkah province (which includes the holy 42 city of Makkah, Jeddah and Tayef); Madinah province (which 43 includes the holy city of Madinah); Riyadh province (which 44 includes the capital city, Riyadh); the Eastern province 45 (which includes Dammam, Ahsa and Hafr Albaten); Asir 46 47 province (which includes Abha and Bisha); Jouf province 48 (which includes Jouf and Qerayyat); Hudud Shamaliyah (North Borders) province (which includes Arar); and Baha, 49 Jizan, Nairan, Hail, Oassim and Tabook provinces. The latest 50 census conducted in Saudi Arabia in 2004 indicated that the 51 total population was 22 673 538, of whom 16 529 302 (72.9%) 52 subjects were Saudis. Approximately 40.8% of the popula-53 tion is below 15 years, 56.1% is 15 to 64 years, and 3.1% is 54 above 64 years of age. The population annual growth rate is 55 3.3%. The infant mortality rate is 19.1 per 1000 live births, 56 and the maternal mortality rate is 1.8 per 10 000 live births. 57 The total life expectancy at birth is 71.4 years. 58

59 2.2. Data collection

HCV infection and other causes of acute or chronic viral hep-60 atitis have been notifiable in Saudi Arabia since 1990. The 61 Ministry of Health officials rely on health-care providers, lab-62 oratories and other public health personnel to report the 63 occurrence of these infections to the Department of Pre-64 ventive Medicine in the Central Ministry of Health office 65 in Riyadh, where all surveillance data are compiled. HCV 66 infection during the study period was identified by detec-67 tion of antibodies to HCV (anti-HCV) by ELISA. Specimens 68 with a reactive ELISA result were retested in duplicate. If 69 the result of either duplicate test was reactive, the speci-70 men was defined as repeatedly reactive and was interpreted 71 as screening-test-positive. Screening-test-positive results 72 73 were confirmed using the recombinant immunoblot analyses (RIBA; Chiron Corporation, Emeryville, CA, USA). Only RIBA-positive patients were included in the study. Indications for testing for anti-HCV included: clinical suspicion and routine screening of blood and organ donors; contacts of HCV-infected patients; prisoners; intravenous drug users; patients with other sexually transmitted infections; and expatriates pre-employment. Obtaining informed consent for testing for HCV or other blood-borne infections is not required in Saudi Arabia.

3. Results

From January 1995 to December 2005, 24948 people with HCV infection were reported to the Ministry of Health, and of these 19185 (76.9%) were Saudis. The number of cases per 100000 population by regions is shown in Table 1. The number of HCV infections by region ranged from 0.016% to 0.322% (16–322 cases per 100000 population), with a mean of 0.124% (124 cases per 100 000 population). Figure 1 shows the number of HCV cases by year. Figure 2 shows the number of cases by age group. The mean pediatric (children <15 years of age) and adult (subjects \geq 15 years of age) populations during the study period were 8 186 369 and 11 878 260 individuals, respectively. The total number of HCV cases reported among children was 998 cases, or 0.012% (12 cases per 100 000 pediatric population). The total number of HCV cases reported among adults was 23950 cases, or 0.202% (202 cases per 100 000 adult population).

4. Discussion

In 1989, HCV was first identified and found to be responsible for most transfusion-associated non-A non-B viral

Please cite this article in press as: Madani TA. Hepatitis C virus infections reported over 11 years of surveillance in Saudi Arabia. *Trans R Soc Trop Med Hyg* (2008), doi:10.1016/j.trstmh.2008.08.001

ARTICLE IN PRESS

Hepatitis C virus infection in Saudi Arabia



Figure 1 Annually reported subjects with antibodies to hepatitis C virus in Saudi Arabia with a moving average trend line (1995–2005).

hepatitis.^{3,4} Before HCV identification, the major causes 103 of HCV infection worldwide were the use of unscreened 104 blood transfusions, and re-use of needles and syringes that 105 were not adequately sterilized. Screening of blood and organ 106 donors for HCV since 1990 has virtually eliminated the spread 107 of HCV by these routes. Consequently, sharing contaminated 108 needles has become the most common mode of transmission 109 of this infection.¹ Sexual and perinatal transmission may also 110 occur, although less frequently.⁵⁻⁷ Infection via other modes 111 of transmission, such as ear and body piercing, circumcision, 112 tattooing and cupping (Hijama), can occur if inadequately 113 sterilized equipment is used.² 114

The prevalence of anti-HCV in some neighboring and 115 other Islamic countries was reported to be as follows: Alge-116 ria 0.2%; Egypt 18.1%; Indonesia 2.1%; Iraq 0.5%; Jordan 117 2.1%; Kuwait 3.3%; Libya 7.9%; Malaysia 3.0%; Mauritania 118 1.1%; Morocco 1.1%; Oman 0.9%; Pakistan 2.4%; Palestine 119 2.2%; Qatar 2.8%; Somalia 0.9%; Sudan 3.2%; Tunisia 0.7%; 120 Turkey 1.5%; United Arab Emirates 0.8%; and Yemen 2.6%.¹ 121 The prevalence of HCV in Saudi Arabia and the aforemen-122 tioned countries is similar to the prevalence reported in 123 industrialized countries.¹ One notable exception is Egypt, 124 in which the prevalence is known to be exceptionally high 125 (up to 40% in some parts of Egypt), probably because of the 126 use of non-disposable, non-sterilized syringes to adminis-127 ter tartar emetic in mass treatment campaigns to control 128 schistosomiasis from the 1960s to the 1980s.8-12 129





Previous studies in Saudi Arabia indicated that the anti-HCV prevalence was 0.4 to 1.7% for adults and 0.1% for children.^{13–16} For instance, a study in Jeddah among 528 blood donors showed an anti-HCV prevalence of 1.7%.¹³ In a large study among 557813 Saudi subjects of all ages in Riyadh province, the anti-HCV prevalence was 1.1% for adults and 0.1% for children.¹⁶ A study among 24173 blood donors in Riyadh province over a three-year period from January 2000 to December 2002 showed an anti-HCV prevalence of 0.4%.¹⁵ A recent study among 13 443 blood donors in the Eastern province of Saudi Arabia showed a decline in the prevalence of anti-HCV from 1.04% in 1998 to 0.59% in 2001.¹⁴ In this nationwide study, the number of HCV infections reported in Saudi Arabia ranged from 0.016 to 0.322% in various regions, with a mean of 0.124%. The number of cases reported among adults was on average 0.202% and that among children was 0.012%. The relatively low number of reported cases in this study is likely to be due to underreporting, as the majority of subjects infected with this virus are usually asymptomatic. The prevalence of anti-HCV slightly and steadily rose from 1998 to 2002, perhaps owing to improved reporting and/or population growth, estimated to be 3.3% annually. From 2003 to 2005, the number of annually reported cases seemed to have plateaued (Figure 1).

In this study, the number of HCV infections reported 154 among children (0.012%) was much lower than that among 155 adults (0.202%). This suggests that perinatal and childhood 156 transmission is not a major mode of infection. However, pre-157 dominance of this infection in adults suggests that other 158 modes of transmission, such as unscreened blood transfu-159 sion before 1990 and intravenous drug use, are the main 160 modes of infection in Saudi Arabia. Substance abuse is an 161 increasing problem in Saudi Arabia, as it is in the rest of 162 the world.¹⁷ Substances abused include injectable drugs 163 such as heroin and cocaine and non-injectable drugs such as 164 cannabis and amphetamine-type stimulants. The estimated 165 annual prevalence of substance abuse in Saudi Arabia in 166 2000 as percentage of the population aged 15 and above 167 was 0.01% for heroin and 0.002% for amphetamine.¹⁷ The 168 number of drug abusers annually admitted to detoxifica-169 tion centers in Riyadh, Jeddah, Dammam and Qassim from 170 1996 to 2001 ranged from 4740 to 6650 patients, with an 171 average annual increment of 5.1% (unpublished data). Sev-172 eral studies were conducted in Saudi Arabia to describe 173 the socio-demographic characteristics, pattern of substance 174 abuse, and prevalence of blood-borne infections among drug 175 abusers. For instance, 799 drug abusers from a voluntary 176 detoxification unit in Jeddah were studied.¹⁸ Sixty-eight per-177 cent of subjects were under 35 years of age, and 64% starting 178 taking drugs before age 25. Eighty-seven percent used heroin 179 or alcohol, and 14% were dependent on more than one drug. 180 Among heroin users, 91% injected the drug. The prevalence 181 of HCV infection among these patients was 69%.¹⁸ In another 182 study of 349 drug abusers in Jeddah, 281 (80.5%) subjects 183 were intravenous drug users. The prevalence of hepatitis B 184 surface antigen (HBsAg), antibodies to HBsAg and antibod-185 ies to hepatitis B core antigen was 12.6, 49.0 and 53.6%, 186 respectively, suggesting that sharing of needles was a com-187 mon practice.¹⁹ In a more recent study in Jeddah including 188 1321 drug abusers, 1038 (78.6%) subjects were intravenous 189 drug users, and the prevalence of HBsAg and anti-hepatitis 190 D virus was 6.1 and 0.8%, respectively.²⁰ Among 116 drug 191

Please cite this article in press as: Madani TA. Hepatitis C virus infections reported over 11 years of surveillance in Saudi Arabia. *Trans R Soc Trop Med Hyg* (2008), doi:10.1016/j.trstmh.2008.08.001

130

131

132

133

134

135

136

137

138

139

140

141

142

143

144

145

146

147

148

149

150

151

152

153

4

192

193

194

195

196

197

198

199

201

204

252

253

278

279

280

281

282

283

284

285

286

287

288

289

290

291

292

293

294

295

296

297

298

299

300

301

302

303

304

305

306

307

308

309

310

311

312

313

users in the Eastern province, 83% of subjects were below 32 years of age, 52.6% were unemployed, and the majority was of intermediate education.²¹ Eighty-four percent of the patients abused heroin either alone or in combination with other drugs, 31% used alcohol, 26% used cannabis and 10% used stimulants. The use of other drugs was rare.²¹ Thus, substance abuse seems to be a potential major risk factor for the spread of HCV among adults in Saudi Arabia.

The prevalence of HCV infection is lower than the preva-200 lence of hepatitis B virus (HBV) in Saudi Arabia. A recent study showed that the prevalence of HBV infection in Saudi 202 Arabia was on average 0.15%, with wide variations between 203 various regions, ranging from 0.03 to 0.72%.²² In that study, the prevalence of HBV among children was 0.05% and that 205 among adults was 0.22%. The prevalence of HIV infection 206 in Saudi Arabia has also been recently described. It ranged 207 from 0.002 to 0.074%, with an average of 0.039%.²³ 208

An important limitation of this study is that it was not a 200 cross-sectional survey but rather a passive reporting of anti-210 HCV positive cases to the Ministry of Health, thus carrying a 211 212 risk of under- or perhaps over-estimation of the actual magnitude of HCV infection in Saudi Arabia. On the one hand, 213 routine testing of low-risk groups (such as blood or organ 214 donors and expatriates pre-employment) may underesti-215 mate the prevalence. On the other hand, testing high-risk 216 groups (such as drug abusers, prisoners, contacts of HCV-217 infected patients, patients with other sexually transmitted 218 infections) may overestimate the actual prevalence. Given 219 such a limitation, comparing the anti-HCV prevalence in this 220 study to cross-sectional survey prevalence in other countries 221 may not be sound. However, comparing anti-HCV prevalence 222 between regions within Saudi Arabia may be acceptable, 223 as all regions followed the same system of diagnosis and 224 notification. 225

As there is as yet no vaccine to prevent HCV infection, 226 the strategy to prevent this infection in Saudi Arabia focuses 227 mainly on health education, routine screening of blood 228 and organ donors and high-risk subjects such as household 229 and sexual contacts of HCV patients, hemodialysis patients, 230 patients requiring recurrent blood transfusion, intravenous 231 drug users and patients with other sexually transmitted 232 infections. Additionally, implementation and maintenance 233 of proper infection control practice in health-care settings, 234 including standard precautions and proper sterilization of 235 surgical and dental equipment, and good hygienic practice 236 in barbershops and traditional therapy settings such as wet 237 cupping (Hijama), are emphasized in Saudi Arabia. 238

In conclusion, the prevalence of HCV infection in Saudi 239 240 Arabia is low. In the absence of an HCV vaccine, efforts to prevent the spread of this infection should focus on: ensuring 241 safe blood, blood products and organs for transplantation; 242 ensuring safe use of syringes, needles, sharps and other 243 equipment used for medical or traditional percutaneous 244 interventions; implementation and maintenance of proper 245 infection control practice in health-care settings; and health 246 education, particularly targeting high-risk groups. 247

Author's contribution: TAM is responsible for all aspects of 248 this study and is guarantor of the paper. 249

Funding: None. 250

Conflicts of interest: None declared.

Ethical approval: Not required.

References

- 1. WHO. Hepatitis C global prevalence (update). Wkly Epidemiol Rec 1999:74:421-8.
- 2. Thomson BJ, Finch RG. Hepatitis C virus infection. Clin Microbiol Infect 2005;11:86-94.
- 3. Choo QL, Kuo G, Weiner AJ, Overby LR, Bradley DW, Houghton M. Isolation of a cDNA clone derived from a blood-borne non-A, non-B viral hepatitis genome. Science 1989;244: 359-62.
- 4. Houghton M. Hepatitis C viruses. In: Fields BN, Knipe DM, Howley PM, Chanock RM, Melnick JL, Monath TP, Roizman B, Straus SE, editors. Fields Virology. Philadelphia, PA: Lippincott; 1996. p. 1035-58.
- 5. Magder LS, Fix AD, Mikhail NN, Mohamed MK, Abdel-Hamid M, Abdel-Aziz F, et al. Estimation of the risk of transmission of hepatitis C between spouses in Egypt based on seroprevalence data. Int J Epidemiol 2005;34:60-165.
- 6. Ohto H, Terazawa S, Sasaki N, Sasaki N, Hino K, Ishiwata C, et al. Transmission of hepatitis C virus from mothers to infants. The Vertical Transmission of Hepatitis C Virus Collaborative Study Group. N Engl J Med 1994;330:744-50.
- 7. Tahan V, Karaca C, Yildirim B, Bozbas A, Ozaras R, Demir K, et al. Sexual transmission of HCV between spouses. Am J Gastroenterol 2005:100:821-4.
- 8. Darwish MA, Raouf TA, Rushdy P, Constantine NT, Rao MR, Edelman R. Risk factors associated with a high seroprevalence of hepatitis C virus infection in Egyptian blood donors. Am J Trop *Med Hyg* 1993;**49**:440–7.
- 9. Darwish MA, Faris R, Clemens JD, Rao MR, Edelman R. High seroprevalence of hepatitis A, B, C, and E viruses in residents in an Egyptian village in The Nile Delta: a pilot study. Am J Trop *Med Hyg* 1996;**54**:554–8.
- 10. Darwish MA, Faris R, Darwish N, Shouman A, Gadallah M, El-Sharkawy MS, et al. Hepatitis C and cirrhotic liver disease in the Nile delta of Egypt: a community-based study. Am J Trop Med Hyg 2001;64:147-53.
- 11. Frank C, Mohamed MK, Strickland GT, Lavanchy D, Arthur RR, Magder LS, et al. The role of parenteral antischistosomal therapy in the spread of hepatitis C virus in Egypt. Lancet 2000;355:887-91.
- 12. Rao MR, Naficy AB, Darwish MA, Darwish NM, Schisterman E, Clemens JD, et al. Further evidence for association of hepatitis C infection with parenteral schistosomiasis treatment in Egypt. BMC Infect Dis 2002;2:29.
- 13. Abdelaal M, Rowbottom D, Zawawi T, Scott T, Gilpin C. Epidemiology of hepatitis C virus: a study of male blood donors in Saudi Arabia. Transfusion 1994;34:135-7.
- 14. Bashawri LA, 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;26:225-8.
- 15. 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.
- 16. Shobokshi OA, Serebour FE, Al-Drees AZ, Mitwalli AH, Qahtani A, Skakni LI. Hepatitis C virus seroprevalence rate among Saudis. Saudi Med J 2003;24:S81–86.
- 17. United Nations Office on Drugs and Crime. Global illicit drug trends 2003. New York, NY: United Nations Publications; 2003.
- 18. Iqbal N. Substance dependence. A hospital-based survey. Saudi Med J 2000;21:51-7.

Please cite this article in press as: Madani TA. Hepatitis C virus infections reported over 11 years of surveillance in Saudi Arabia. Trans R Soc Trop Med Hyg (2008), doi:10.1016/j.trstmh.2008.08.001

314

315

316

ARTICLE IN PRESS

Hepatitis C virus infection in Saudi Arabia

330

 Njoh J. Prevalence of hepatitis B virus markers among drugdependent patients in Jeddah Saudi Arabia. *East Afr Med J* 1995;**72**:490–1.

 20. Njoh J, Zimmo S. Prevalence of antibody to hepatitis D virus among HBsAg-positive drug-dependent patients in Jeddah, Saudi Arabia. *East Afr Med J* 1998;75:327–8.

- 21. Hafeiz HB. Socio-demographic correlates and pattern of drug abuse in eastern Saudi Arabia. *Drug Alcohol Depend* 1995;38:255–9.
- 22. Madani TA. Trend in incidence of hepatitis B virus infection during a decade of universal childhood hepatitis B vaccination in Saudi Arabia. *Trans R Soc Trop Med Hyg* 2007;101: 278–83.
- 23. Madani TA, Al-Mazrou YY, Al-Jeffri MH, Al Huzaim NS. Epidemiology of the human immunodeficiency virus in Saudi Arabia; 18-year surveillance results and prevention from an Islamic perspective. *BMC Infect Dis* 2004;4:25.

Please cite this article in press as: Madani TA. Hepatitis C virus infections reported over 11 years of surveillance in Saudi Arabia. *Trans R Soc Trop Med Hyg* (2008), doi:10.1016/j.trstmh.2008.08.001