

Original Article

Prevalence and Factors Associated with Hepatitis C Virus Infection among Myanmar Blood Donors

Myo-Khin^a, San-San-Oo^a, Khin May Oo^a, Kunio Shimono^b,
Norio Koide^c, and Shigeru Okada^{c*}

^aDepartment of Medical Research-Lower Myanmar, No. 5 Ziwaka Road, Yangon, Myanmar,

^bShimono Naika-Geka Clinic, Bizen, Okayama 705-0035, Japan, and

^cDepartment of General Medicine, Okayama University Hospital, Okayama 700-8558, Japan

We studied the prevalence of hepatitis C virus infection among blood donors from 3 hospitals of Central Myanmar and 7 hospitals of Lower Myanmar in the Yangon area, and analyzed the factors associated with the infection. The study period was from November, 2005 to June, 2007. A pre-tested questionnaire was used to obtain information on age, ethnic group, marital status, tattooing, body piercing, history of receiving transfusions, and liver diseases in self and in sexual partners. Data on seropositivity to hepatitis C, hepatitis B and human immunodeficiency virus infections were recorded. A total of 65,240 blood donors participated in the study. Their ages ranged from 18 years to 60 years (mean \pm SD = 29.5 \pm 9.3). The male-to-female ratio was 6 : 1. The prevalence of the antibody to hepatitis C was found to be 0.95% with varying rates (0.34 to 2.03) among hospitals. Females had a slightly higher rate (1.06%) than males (0.93%) ($p = 0.237$). Multivariate analyses revealed the following factors to be related to HCV infection: HIV infection, odds ratio (OR) = 3.0 ($p = 0.003$); history of liver disease, OR = 8.9 ($p = 0.001$); and age 30 years and above, OR = 2.6 ($p = 0.001$). We discuss the varying prevalences of HCV around the world.

Key words: Myanmar, hepatitis C prevalence, blood donors, associated factors

Hepatitis C virus (HCV) infection is an important health problem in many countries [1]. According to a review in 2000, 3 percent of the world's population or almost 200 million persons are infected with HCV [2]. HCV has been shown to be associated with blood transfusion; although the acute infection is usually asymptomatic, the subsequent chronic infection is usually life-long and may lead to chronic liver disease, liver cirrhosis, and hepatocellular carcinoma. In fact in Japan where the incidence of hepatitis B virus infection is low, HCV has been

proven as the single most important etiological factor for the development of hepatocellular carcinoma, [3].

HCV is regarded as an emerging health problem in Myanmar. Reports in the early 90s showed HCV infection in one-third of patients with hepatocellular carcinoma (Khin-Pyone-Kyi *et al.*, (1992) presented at the 38th Myanmar Medical Conference, Yangon), and in 2.5% of apparently healthy subjects (Khin-Pyone-Kyi *et al.*, (1995) presented at the Myanmar Health Research Congress, Yangon) in Myanmar.

In May, 2000, anti-HCV testing was introduced at blood banks in Yangon. From May, 2000 to October, 2003, a total of 102,632 donors were screened, and the overall anti-HCV positivity rate was found to be 2.8% (Paing-Soe *et al.*, (2002) presented at the

Myanmar Health Research Congress, Yangon). In 2005, hepatitis C screening of blood donors was extended to Upper Myanmar. Blood banks from major hospitals in Mandalay and Magwe were included in the protocol. Although more data on the prevalence of HCV infection among blood donors became available (Myo-Khin *et al.*, (2006) presented at the Myanmar Health Research Congress, Yangon), blood bank personnel have emphasized a lack of information on the factors associated with HCV infection in blood donors.

We carried out this study to determine the associated factors of HCV infection in blood donors for use in the deferral of high-risk blood donors.

Subjects and Methods

Blood donors from 3 hospitals in Central Myanmar (Mandalay General Hospital, Magwe General Hospital, Pyinmana Township Hospital) and 7 hospitals (Yangon Children Hospital, Central Women Hospital, East Yangon General Hospital, West Yangon General Hospital, Insein General Hospital, Thingangyun Sanpya Hospital and North Okkalapa General Hospital) in the Yangon area were included. Prior to the study, informed consent was obtained from each donor. The study was conducted in accordance with the principles of Good Clinical Practice and the Declaration of Helsinki.

The OneStep HCV RapiDip test (CORTEZ Diagnostics Inc. Calabasas, CA, USA) was used at the hospitals for HCV seropositivity screening. This one-step test is based on the principle of a double antigen sandwich immunoassay for determination of anti-HCV in serum. It is very sensitive and only takes 10–20 min for the result to be read. Test results are read visually without any instrument.

The study period was from November, 2005 to June, 2007. A pre-tested questionnaire was used to obtain information on age, ethnic group, marital status, tattooing, body piercing, history of receiving transfusions, and liver diseases in self and in spouses. Data was collected by the laboratory personal from the participating hospitals. Data on seropositivity to hepatitis C, hepatitis B and human immunodeficiency virus infections were also recorded.

Statistical methods. Data analysis was performed with computer software MINITAB Statistical

Package Version 14.1; MINITAB Inc., (PA, USA) The chi-square test or Fisher's exact test was used as appropriate to evaluate the statistical significance in the distribution of each factor between blood donors who were seropositive and seronegative hepatitis C. Multiple logistic regression analysis was used to determine the independent association between potential factors and HCV seropositivity after controlling for potential confounding factors. Crude and adjusted odd ratios (ORs) and 95 percent confidence intervals (CIs) were calculated [4].

Results

General characteristics. A total of 65,240 blood donors participated in the study. Their ages ranged from 18 years to 60 years (mean \pm SD = 29.5 \pm 9.3). Male blood donors dominated female blood donors with a ratio of 6 : 1. Females were significantly older than the males (Table 1).

HCV antibody prevalence. The prevalence of the antibody to hepatitis C among the study population was found to be 0.95%. The HCV seropositivity rate in blood donors varied from 0.34 to 2.03 among hospitals (Table 2).

Females had a slightly higher prevalence of anti-HCV seropositivity than males (1.06% vs 0.93%; Student's *t* test, *p* = NS). The anti-HCV seropositivity rate increased with increasing age group with the exception of the oldest group. It was found to be the lowest (0.53%) in the 18–20 years age group and highest (1.88%) in the 41–50 years age group (Table 3).

Associated risk factors for anti-HCV seropositivity. After univariate analysis, it was found that the significant associated factors among the population were: (a) marital status, (b) history of liver disease, (c) age of 30 years and above, (d) hepatitis B surface antigen seropositivity, and (e) presence of the antibody to HIV. There was no significant association

Table 1 Mean age of blood donors in relation to gender

Sex	Number	Mean age (yrs) \pm SD	Student's 't' test
Males	55920	29.4 \pm 9.3	T-value 7.25
Females	9320	30.2 \pm 9.2	df = 11981
Both	65240	29.5 \pm 9.3	P-value = 0.0001

Table 2 HCV seropositivity rate in blood donors by participating hospitals

Hospital	Number of donors	Number HCV positive	HCV seropositive rate
Children Hospital Yangon	6,716	58	0.86
Central Women HospitalH	6,022	44	0.73
East Yangon General Hospital	1,847	16	0.87
West Yangon General Hospital	919	12	1.31
North Okkalapa General Hospital	8,397	45	0.54
Insein General Hospital	2,274	9	0.40
Sanpya General Hospital	4,630	23	0.50
Mandalay General Hospital	31,079	351	1.13
Magwe General Hospital	3,060	62	2.03
Pyinmana Hospital	293	1	0.34

with sex, history of receiving transfusions, history of body piercing, history of tattooing, or history of hepatitis in spouse. (Table 4).

Multiple logistic regression was applied for con-

Table 3 Anti-HCV seropositivity by age group

Age group	Total	Anti-HCV seropositivity	
		Positive (%)	Negative (%)
18–20 yrs	10907	58 (0.5%)	10,849 (99.3%)
21–30 yrs	27596	186 (0.7%)	27,410 (99.3%)
31–40 yrs	14923	211 (1.4%)	14,712 (98.6%)
41–50 yrs	7489	141 (1.9%)	7,348 (98.1%)
51–60 yrs	1608	23 (1.4%)	1,585 (98.6%)

Chi-Square = 142.907, DF = 4, *P*-Value = 0.0001

trolling confounders and for evaluating the effects of associated factors on HCV infection in the blood donors. After analysis, 3 variables history of liver disease, age of 30 years and above, and presence of the antibody to HIV were found to have an effect on the presence of HCV infection. (Table 5).

Table 4 Associated factors for anti-HCV seropositivity by univariate analysis

Associated factors		Anti-HCV seropositivity		<i>p</i> value
		No. tested	No. positive (%)	
Sex	Male	56,383	526 (0.93)	<i>p</i> = 0.237
	Female	8,834	94 (1.42)	
Marital status	Never	14,942	183 (1.22)	<i>p</i> = 0.001
	Ever	48,628	426 (0.88)	
History of liver diseases	Yes	47	3 (6.38)	<i>p</i> = 0.01
	No	65,189	618 (0.95)	
History of receiving transfusion	Yes	59	0 (0.00)	<i>p</i> = 0.907
	No	65,177	621 (0.95)	
History of body piercing	Yes	638	2 (0.31)	<i>p</i> = 0.095
	No	64,598	619 (0.96)	
History of tattooing	Yes	408	4 (0.98)	<i>p</i> = 0.953
	No	64,828	617 (0.95)	
History of hepatitis in spouse	Yes	5	0 (0.00)	<i>p</i> = 0.90
	No	65,230	620 (0.95)	
Age 30 years and above	Yes	23,645	375 (1.56)	<i>p</i> = 0.001
	No	38,502	244 (0.63)	
Hepatitis B surface antigen seropositivity	Positive	2,221	11 (0.50)	<i>p</i> = 0.024
	Negative	63,015	610 (0.97)	
Antibody to HIV	Positive	288	8 (2.78)	<i>p</i> = 0.001
	Negative	64,948	613 (0.94)	

Table 5 Factors associated with anti-HCV seropositivity among blood donors by multivariate analysis

Associated factors		Adjusted OR	95% CI of OR	p value
History of liver diseases	Yes	8.86	2.65–29.65	$p = 0.001$
	No	1		
Age 30 years and above	Yes	2.59	2.18– 3.08	$p = 0.001$
	No	1		
Presence of antibody to HIV	Yes	2.96	1.46– 6.02	$p = 0.003$
	No	1		

Discussion

The problem of hepatitis C virus infection in Myanmar is well recognized, and efforts to control it in the blood donor population were initiated in 2000. Hepatitis C screening of 154,161 blood donors from May, 2000 to April, 2004 demonstrated the prevalence of HCV infection among Myanmar blood donors to be 2.6% [5]. Low HCV seropositivity in blood donors (<1%) has been reported from Europe [2] and the Far East (1%) [6]. However, high rates of HCV infection had been reported in Egypt, with 4% of blood donors being positive for anti-HCV antibodies and a higher prevalence rate of 15% in the rural areas of the country [7]. The HCV prevalence in Myanmar could be termed intermediate as the global seroprevalence of HCV among blood donors varies from 0.4 to 19.2% [8].

The present study highlights the increasing prevalence of anti-HCV seropositivity with age. The lowest seropositivity was observed in the 18–20 years age group; and seropositivity tended to increase with age reaching a maximum in the 41–50 years age group. Comparable results have been observed in studies on blood donors from India [9]. A similar pattern has also been reported from Japan [10], most likely due to the tumultuous period after World War II, marked by illicit intravenous amphetamine abuse and unsafe healthcare-related procedures like the use of unsafe needles in group vaccinations. With these results in mind, efforts should be made to increase retain young motivated voluntary donors to maintain a safe blood supply.

The seroprevalence of HCV in voluntary blood donors from different hospitals in India was shown to vary between 0.12 and 4 percent [11]. In the present study, HCV seropositivity among blood donors varied from hospital to hospital. Although the differences

could be geographical in nature, a recent study highlights the benefit of earlier introduction of hepatitis C screening, as evidenced by the lower rates of hepatitis C seropositivity among blood donors in Yangon compared to Mandalay and Magwe (Myo-Khin *et al.*, (2006) presented at the Myanmar Health Research Congress, Yangon). This finding is also supported by the results of the present study where anti-HCV seroprevalence among blood donors in hospitals from Yangon was much lower than that of blood donors from Mandalay and Magwe.

The present study highlights the associated factors of HCV infection among blood donors. A previous history of liver diseases, age of 30 years and older, and seropositivity to HIV were identified as independently associated factors. Coinfection with HIV and the hepatitis C virus (HCV) has been reported in the past. The diseases are spread in similar ways, notably through shared use of needles to inject illicit drugs and sexual activities. Many people are coinfecting with HIV and HCV, HIV and HBV, or even all 3 viruses [12]. A previous study carried out among Yangon blood donors also outlined associated factors. A higher prevalence of anti-HCV positivity was found in those with a history of surgical operation, tooth extraction, ear piercing and tattooing (Thein-Saw *et al.*, (1998) presented at the Myanmar Health Research Congress, Yangon). However, due to the nature of the present study, only minimal relevant data could be collected.

Variables that could cause confusion and were difficult to recall were not included. It was decided that an individual's history of surgical operation and tooth extraction procedures might not be easily defined and thus were excluded from the data collection form. Ear piercing was also not included as it is a very common practice among the Myanmar population, especially among females.

Although HCV prevalence surveys among blood donors could not be considered to be representative of the population at large, it could be concluded that the prevalence of HCV infection in blood donors is generally less than that in the general population as demonstrated by surveys carried out among apparently healthy populations. The prevalence of the antibody to HCV was found to be 2.8% among 569 subjects (246 males, 323 females), aged 3 months to 74 years residing in Yangon [13]. It has been reported that rates of HCV infection among populations residing in border areas of Myanmar were higher ($> 10\%$) than that of blood donors from Yangon and Mandalay [14].

The risk factors for HCV seropositivity among Israelis include gum surgery, hospitalization without surgery, and therapy in injection form [15]. We did not include these items in our data form as our previous studies among blood donors and the general population did not establish them as significant factors in Myanmar [13, 14]. A study from Australia identifies the use of injected drugs as a potential risk factor [16]. However, drug use is not easy to define, and as self-revelation of such a factor is also not dependable, it was not included in the data collection form.

The present study was based on the experiences of a field study where characteristics of hepatitis C seropositive subjects from communities in the border areas of Myanmar were used to develop a simple cost-free system for assessment of HCV infection [17]. Many health care facilities in Myanmar do not have the resources to implement routine HCV testing of blood donors given the current availability and the cost of the test. The factors identified by the present study as associated with HCV infection could help local health personnel in the deferral of blood donors who are likely to be seropositive to HCV infection.

Acknowledgments. We would like to acknowledge the participation of the pathologists and staff members of the hospitals from Yangon, Mandalay and Pinyinana in the collection of data.

References

1. Cohen J: The scientific challenge of hepatitis C. *Science* (1999) 285: 26–30.
2. Wasley A and Alter MJ: Epidemiology of Hepatitis C: Geographic Differences and Temporal Trends. *Semin Liver Dis* (2000) 20: 1–16.
3. Tanaka K, Hirohata T, Koga S, Sugimachi K, Kanematsu T, Ohryohji F, Nawata H, Ishibashi H, Maeda Y, Kiyokawa H, Tokunaga K, Irita Y, Takeshita S, Arase Y and Nishino N: Hepatitis C and Hepatitis B in the Etiology of Hepatocellular Carcinoma in the Japanese Population Cancer Research (1991) 51: 2842–2847.
4. Altman DG: *Practical Statistics for Medical Research*, 1st ed, Chapman and Hall, London (1991).
5. Myo-Khin and Tin-Nu-Swe: Contributions by the Japan International Cooperation Agency to Hepatitis C Control and Research in Myanmar DMR Bull (2003) 17: 1–17.
6. Watanebe J, Minegishi K, Mitsumori T, Ishifuji M, Oguchi T, Ueda M, Tokunaga E, Tanaka E, Kiyosawa K and Furuta S: Prevalence of anti-HCV antibody in blood donors in the Tokyo area. *Vox Sang* (1990) 59: 86–88.
7. El-Zahadi A, Selim O, Rafik M and El-Haddad S: Prevalence of hepatitis C virus among non-A, non-B-related chronic liver disease in Egypt. *J Hepatol* (1992) 14: 2–3.
8. Memon MI and Memon MA: Hepatitis C: An epidemiological review. *J Viral Hepat* (2002) 9: 84–100.
9. Thakral B, Marwaha N, Chawla YK, Saluja K, Sharma A, Sharma RR, Minz RW and Agnihotri: Prevalence and significance of hepatitis C virus (HCV) seropositivity in blood donors. *Indian J Med Res* (2006) 124: 431–438.
10. Okayama A, Stuver SO, Tabor E, Tachibana N, Kohara M, Mueller NE and Tsubouchi H: Incident hepatitis C virus infection in a community-based population in Japan. *J Viral Hepat* (2002) 9: 43–51.
11. Panigrahi AK, Panda SK, Dixit RK, Rao KV, Acharya SK, Dasarathy S and Nahu A: Magnitude of hepatitis C virus infection in India: Prevalence in healthy blood donors, acute and chronic liver disease. *J Med Virol* (1997) 51: 167–174.
12. Talal AH, Cancis PW and Jacobson I: The HCV and HIV coinfecting patient: what have we learned about pathophysiology?
13. Myo-Khin: Hepatitis C infection in different population groups. *Proceedings of the Seminar on Control of Hepatitis C Infection in Myanmar* (2000) 23–29.
14. Myo-Khin: Research Studies that Highlights the Problem of Hepatitis C Infection in Myanmar. *Proceedings of the Workshop on Developing IEC Package Regarding Hepatitis C Prevention in Myanmar*; 2001 (2001) 16–21.
15. Kerzman H, Green MS and Shinar E: Risk factors for hepatitis C virus infection among blood donors in Israel: a case-control study between native Israelis and immigrants from the former Soviet Union. *Transfusion* (2007) 47: 1189–1196.
16. Wong PY, Dodd R, Kiely P, Carroll C and Whyte G: Characteristics of hepatitis C-positive blood donors in Victoria, Australia. *Transfus Med* (1999) 9: 15–19.
17. Myo-Khin, Yi-Yi-Kyaw, Win-Pa-Pa-Naing, Than-Than-Aye, Swe-Zin-Yu, San-San-Oo and Khine-Win: Use of risk scores for screening of hepatitis C of blood donors in remote areas. *Myanmar Health Sciences Res J* (2006) 18: 109–113.