Seroepidemiology of HTLV-1 and HTLV-2 Infection in Neyshabur City, North-Eastern Iran, during 2010-2014

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ABSTRACT

Background: Retroviruses of human T-lymphotropic viruses (HTLV-1 and HTLV-2) have been demonstrated to be endemic in the north-eastern region of Iran. This study was aimed to determine the HTLV-1 and HTLV-2 prevalence among healthy individuals in Neyshabur City during 2010-2014. **Methods:** A total of 8054 blood samples were collected from healthy participants in Neyshabur, North-Eastern Iran. The blood samples were screened for the presence of specific antibodies against HTLV-1 and HTLV-2 by using ELISA according to the manufacturer's instructions. **Results:** The overall seropositivity rate for HTLV-1 and HTLV-2 was found to be 6.55% (528 out of 8054) among participants. **Conclusion:** Both HTLV-1 and HTLV-2 were demonstrated to be at a high rate in healthy individuals. However, a smaller number of asymptomatic carriers were found in this study, as compared to those identified in previous investigations in the city. **DOI:** 10.6091/.21.1.57

Keywords: Human T-lymphotropic viruse, Seroepidemiology, Enzyme-linked immunosorbent assay, Iran

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INTRODUCTION

uman T-lymphotropic viruses (HTLV-1 and HTLV-2), classified in the retroviridae family, are among the first identified species^[1-3]. HTLV-1 and HTLV-2 are widespread all over the world and are endemic in different areas, including North-Eastern Iran^[4-6]. According to a previous study, the rate of HTLV-1 infection has been reported to be less than 0.26% in Mashhad, North-Eastern Iran, while it does not exceed 0.34% in other areas of the country^[7]. The prevalence of HTLV-1 infection in other countries such as Turkmenistan, Brazil, Spain, Korea and Japan was found to be 0.007%^[8], 1.9%^[9], 0.001%^[10], 0.27%^[11], and 0.12%^[12], respectively.

HTLV-1- and HTLV-2-infected carriers remain asymptomatic for a long time, serving as a potential source for the transmission of the disease^[13]. The aim of this investigation was to determine the prevalence of HTLV-1 and HTLV-2 among healthy individuals in Neyshabur, North-Eastern Iran, during 2010-2014.

MATERIALS AND METHODS

Study population

A total of 8054 healthy individuals from Neyshabur, North-Eastern Iran, were included in this study. Serum samples (5 ml) were prepared from the individuals and stored at -20°C until the ELISA test.

Table 1. Age- and sex-based distribution of individuals and overall HTLV-positive cases

Variable	No.	Positive cases (%)	Odd Ratio (OR)	OR (95%CI)	P value
Age (year)					
0-19	429	13(3.03)	Baseline		< 0.0001
20-29	2556	49(1.92)	0.625	0.336-1.163	
30-39	2018	88(4.36)	1.459	0.807-2.637	
≥40	3051	377(12.36)	4.512	2.571-7.918	
Gender					
Male	1565	130(8.31)	1.206	1 120 1 704	0.002
Female	6489	398(6.13)	1.386	1.128-1.704	0.002

Serological assays and confirmation tests

Serum samples were screened for the presence of specific antibodies against HTLV-1 and HTLV-2 by ELISA (Dia.Pro Diagnostic Bioprobes, Italy) according to the manufacturer's instructions^[14].

Statistical analysis

The SPSS software (version 20) was employed to analyze all data using chi-square and t-test. A P<0.05 was considered to be statistically significant.

RESULTS AND DISCUSSION

Of 8054 healthy individuals participated in the study, 1565 (19.4%) and 6489 (80.6%) were males and females, respectively. As shown in Table 1, the mean age of males and females was 46 ± 3 and 51 ± 3 years, respectively. The positivity of the samples was 6.55% (528 out of 8054), including 3.6% for HTLV-1 and 1.4% for HTLV-2. Table 2 indicates the total prevalence of HTLV-1 and HTLV-2 in each year.

Previous studies have revealed that HTLV-1 is endemic in North-Eastern Iran^[15]. Another study in Neyshabur has indicated that the prevalence of HTLV-

1 is 7.2% (35 out of 483)^[16]. However, the rate of HTLV-1 seropositivity has gradually decreased from 1.97% in 1996 to 0.26% in $2014^{[17-19]}$ in other regions of North-Eastern Iran. Similarly, the results of the present study demonstrated that the prevalence of HTLV-1 has decreased in Neyshabur from 2010 to 2014. In a survey carried out in Mashhad in 2012, the rate of HTLV-1 was detected to be 0.47% [20]. The seroprevalence of HTLV-1 did not exceed 0.19% in a study conducted by Safabakhsh et al. [7]. It seems that the reduction in HTLV-1 rate is mainly due to the improvement of blood donor selection and increased awareness among blood donors. However, in a study performed by Rafatpanah et al. [21] in Mashhad, it was revealed that the prevalence of HTLV-1 is 20% (10 positive samples), although no evidence of HTLV-2 infection was found among immuneblotted samples and nested PCR.

In the current study, over 3% of healthy individuals were positive for HTLV-1 in all five years. To the best of our knowledge, there is a small number of published data regarding HTLV-2 prevalence in Iran. Also, a lower rate of positive HTLV-1 infection was identified in the present investigation, when compared to a previously study in Neyshabur^[22]. This finding

Table 2. The annual prevalence of HTLV-1 and HTLV-2 investigated in this study

Year	Number	HTLV-1 (%)	HTLV-2 (%)	Total percentage		
2014	Positive: 58 Total: 1350	3.01	ND	3.01		
2013	Positive: 94 Total: 2337	4.11	ND	4.11		
2012	Positive: 115 Total: 2188	5.12	ND	5.12		
2011	Positive: 117 Total: 2057	5.13	ND	5.12		
2010	Positive: 122 Total: 1789	5.74	ND	5.74		

ND, not determined

highlights that Neyshabur is a major endemic region for HTLV-1. In addition, a higher prevalence of HTLV-1 was found in the age groups over 40 years, suggesting that there is a relationship between HTLVs and the age of individuals.

In the present study, a high rate of HTLV-1 among serum samples was detected using the ELISA test among healthy individuals in Neyshabur city during 2010-2014. The results from this study emphasize that HTLV is still an important endemic disease in Neyshabur. More importantly, the prevalence of HTLV-1 in Neyshabur was detected to be higher than other city (Mashhad) in all duration of this study, though being in a decreasing status compared to the previous reports.

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CONFLICT OF INTEREST. None declared.

REFERENCES

- Chenari M, Norouzi M, Ghalichi L, Rezaee A, Yari A, Alavian SM, Jazayeri SM. Characterization of overt and occult hepatitis B virus infection among HTLV-1 positive healthy carriers in the Northeast of Iran; An HTLV-I endemic area. *Journal of medical virology* 2014; 86(11): 1861-1867.
- 2. Pohanka M, Pavlis O, Pikula J. Galantamine effect on tularemia pathogenesis in a BALB/c mouse model. *Iranian biomedical journal*. 2012; **16**(3): 156-161.
- 3. Doan CC, Le TL, Hoang NS, Doan NT, Le VD, Do MS. Differentiation of umbilical cord lining membrane-derived mesenchymal stem cells into endothelial-like cells. *Iranian biomedical journal*. 2014; **18**(2): 67-75.
- Cook LB, Taylor GP. HTLV-1 and HTLV-2 prevalence in the United States. *Journal of infectious diseases* 2014; 209(4): 486-487.
- Farid R, Farid F, Rezaee SA. Prevalence of HTLV-1 infection in northeast of Iran. *Retrovirology* 2015; 12(Suppl 1): O7.
- Mahzounieh M, Ghorani M, Karimi A, Pourgheysari B, Nikoozad R. Prevalence of human T-lymphotropic virus types I and II in patients with hematological disorders in Isfahan, Iran. *Jundishapur journal of microbiology* 2015; 8(6):e17201.
- Safabakhsh H, Jalalian M, Karimi G. Seroepidemiology of Human T-Cell Lymphotropic Virus Type-1 (HTLV1)

- in Mashhad. Global journal of health science. 2014; **6**(5): 99-104.
- Senyuta N, Syrtsev A, Yamashita M, Stepina V, Susova O, Scherbak L, Pavlish O, Hayami M, Gurtsevitch V. Sero-epidemiologic and phylogenetic studies of HTLV-I infection in 2 countries of the Caspian Sea region. *International journal of cancer* 1998; 77(4): 488-493.
- Carneiro-Proietti ABF, Sabino EC, Leão S, Loureiro P, Sarr M, Busch M, Proietti FA, Murphy EL. HTLV-1/2 prevalence in Brazilian blood donors: regional and demographic variation. *Retrovirology* 2011; 8(Suppl 1): A83
- Toro C, Rodés B, Aguilera A, Caballero E, Benito R, Tuset C, García J, De Lejarazu RO, Eirós JM, Calderón E, Rodriguez C, Soriano V; HTLV Spanish Study Group. Clinical impact of HTLV-1 infection in Spain: implications for public health and mandatory screening. *Journal of acquired immune deficiency syndromes* 2002; 30(3): 366-368.
- 11. Kwon SY, Lim AH, Park JY, Han SH, Cho NS. Seroprevalence of human T-lymphotropic virus type 1 and 2 in Korean blood donors. *Journal of medical virology* 2008; **80**(10): 1864-1867.
- 12. Satake M, Yamaguchi K, Tadokoro K. Current prevalence of HTLV-1 in Japan as determined by screening of blood donors. *Journal of medical virology* 2012; **84**(2): 327-335.
- 13. Tanajura D, Castro N, Oliveira P, Neto A, Muniz A, Carvalho NB, Orge G, Santos S, Glesby MJ, Carvalho EM. Neurological manifestations in human T-cell lymphotropic virus type 1 (HTLV-1)-infected individuals without HTLV-1-associated myelopathy/ tropical spastic paraparesis: A longitudinal cohort study. *Clinical infectious diseases* 2015; **61**(1): 49-56.
- 14. Mortezaie Z, Bouzari M, Roghanian R. Evaluating the frequency of HTLV-I/II infection among blood donors, major thalassemic patients and individuals infected with hepatitis B and C viruses in Isfahan, Iran. *Iranian journal of blood and cancer* 2012; **4**(2): 75-80.
- 15. Rezvan H, Abolghassemi H, Kafiabad SA. Transfusion-transmitted infections among multi-transfused patients in Iran: a review. *Transfusion Medicine* 2007; **17**(6): 425-433.
- Hedayati-Moghaddam M, Fathimoghadam F, Mashhadi IE, Soghandi L, Bidkhori H. Epidemiology of HTLV-1 in Neyshabour, Northeast of Iran. *Iranian Red Crescent* medical journal 2011; 13(6):424-427.
- Safabakhsh HR, Karimi G, Hatami H. The prevalence of HTLV-1 infection in blood donation volunteers in Mashhad. *Journal of School of Public Health and Institute of Public Health Research*. 2014; 11(4): 85-94.
- Tarhini M, Kchour G, Zanjani DS, Rafatpanah H, Otrock ZK, Bazarbachi A, Farid R. Declining tendency of human T-cell leukaemia virus type I carrier rates among blood donors in Mashhad, Iran. *Pathology* 2009; 41(5): 498-499.
- 19. Sani AT. Serologic prevalence of HTLV among blood donors in Mashhad (northeastern Iran). *Archives of Iranian medicine* 2001; **4**(1): 25.
- 20. Hatami H, Karimi G, Safabakhsh H. Seroepidemiologic

- prevalence of HTLV in voluntary blood donors in Mashhad. *Scientific journal of Iranian blood transfusion organization*. 2012; **9**(2): 149-159.
- 21. Rafatpanah H, Fathimoghadam F, Shahabi M, Eftekharzadeh I, Hedayati-Moghaddam M, Valizadeh N, Tadayon M, Shamsian SA, Bidkhori H, Miri R, Bazarbachi A. No Evidence of HTLV-II Infection Among Immonoblot Indeterminate Samples Using
- Nested PCR in Mashhad, Northeast of Iran. *Iranian journal of basic medical sciences* 2013; **16**(3): 229-234.
- 22. Bidkhori H, Hedayati-Moghaddam M, Fathi-Moghaddam F, Soghandi L, Bakhtiari H, Rezaie A. High Prevalence of HTLV-1 Infection among Hemodialysis Patients in Neyshabour, Northeast of Iran. *Iran journal of allergy and asthma immunology* 2013; **12**(Suppl 9): 38.