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Mutual Predators: A Descriptive Cross-Sectional Study to Identify Prevalence of Hepatitis C Virus and Human Immunodeficiency Virus Type-1 Mono and Co-Infection

Prof. Dr. Fouzia Ashraf¹, Dalaq Aiysha², Muhammad Tajamal³, Shahzeb Javed⁴, Saamia Tahir⁵, Omar Ali⁶, Prof. Dr. Mahmood Shaukat⁷

^{1, 2, 3, 7}Allama Iqbal Medical College / Jinnah Hospital, Allama Shabbir Ahmad Usmani Road, Lahore - 54550. Pakistan

⁴King Edward Medical University and Mayo Hospital, Hospital Road, Nelagumbad, Anarkali, Lahore 54000, Pakistan

^{5, 6} CMH Lahore Medical College & Institute of Dentistry, Abdur Rehman Road, Cantt., Lahore, Pakistan-54000

Abstract: Coinfection, bacterial or viral origin, in HIV infected individuals' remains to be the only leading cause of deaths. This study was designed to analyze received plasma samples and plasma samples of referred patients for HIV testing to detect HIV and HCV mono and co-infection by real time PCR and finding co-relation of viral load of both viruses. Highlight and magnify the hidden coinfection, prior to seroconversion, of HIV type-1 and Hepatitis C Virus in received samples. <u>Methods:</u> Analyses were based on randomly selected 78 patients' stored plasmas. Plasma samples were tested for both, HIV-type 1 and HCV viral RNA by real time PCR. Statistical formulas were used to identify mean and the inter quartile range of patients age. The data were analyzed by IBM SPSS Statistics 21 (SPSS Inc., Chicago, IL). Study variables include gender, age and viral loads of HIV type-1 and HCV. Pearson correlation was used to evaluate any correlation in study variables. <u>Result:</u> Prevalence of HCV was 10.3%, HIV-type 1 was 19.2% and their co-infection was 37.2 percent. Thirty three percent individuals had no infection of both viruses. Gender based distribution showed that 74.4% (58/78) sample population was male. The mono-infection and co-infection was higher in males (39.7%) and highest viral load too. There was a positive correlation (CI= 95%) between the two variables; HIV and HCV viral loads, as r = 0.736, n=29, p = 0.001. <u>Conclusion:</u> Prevalence of HIV type-1 and HCV wono-infection and co-infection was higher among males as compared to females. Increased viral load was also evident among male co-infected individuals. This study proved the emergence of HCV coinfection in HIV infected individuals, and a need for on time diagnosis and treatment.

Keywords: HIV, HCV, Coinfection, Corelation, Viral load.

1. Introduction

Target cells of both viruses-Human Immunodeficiency Virus and Hepatitis C Virus-are different, but in co-infection their outcomes are deleterious due to increased disease progression in combined proliferation [1]. Common mode of HIV and HCV transmission is by direct contact with body fluids of an infected person [2]. Globally, prevalence of viral hepatitis among HIV infected people is seven million worldwide [3]. The prevalence rate of co-infection varies considerably between and within countries, depending on rate of risk factors in the population. Viral hepatitis is an emerging non-AIDS related, cause of morbidity and mortality among HIV patients [4].

In HCV mono-infection about fifteen to forty five percent patients with acute viral infection clear the virus, while twenty to fifty percent develop persistent viremia [5]. The persistence of HIV and HCV is attributed to high rates of replication, its $\sim 10^{12}$ and $\sim 10^9$ virion/day for HCV and HIV, respectively [1]. Transmission of communicable disease has been an influential element in acquiring infection and disease progression. The highest rate of HIV type-1 and HCV coinfection has been reported to be 90 percent among in-vitro drug users [6]. Hemophiliacs are the second dominant group of co-infection, who received contaminated blood or its products [7]. The reported prevalence of HIV and HCV among blood donors of Pakistan is 3.78 and 0.06 percent respectively [8, 9, 10]. Co-transmission of HIV and HCV can occur through percutaneous route among injection drug users, but they were usually infected first with HCV [11]. Hepatitis C Virus and HIV type-1 both belongs to two different families of viruses. This particular association has a major effect on their life cycle. HCV replicates through RNA-dependent RNA polymerase, [12] while HIV type-1 first integrates its genetic material in host DNA through the viral reverse transcriptase enzyme and then replicates [13]. There is a direct relation to inoculums size and acquiring HCV infection, after transmission through blood transfusion or by contaminated needles, viremia occurs within days or it takes six to eight weeks, respectively [14]. In America, a large cross-sectional analysis (n=1687) of two HIV trials, reported that 75% subjects had HCV RNA level above 800,000 IU/ml [5].

2. Methodology

2.1 Sampling Procedure and Sources

A cross-sectional study was conducted during the months of May, June and July 2014, among referred patients to Jinnah Hospital Lahore. JHL is the second largest teaching hospital, which provides health care facilities to the inhabitants of Lahore. A random blood samples were collected from

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patients along with their age and gender. Laboratory testing was performed in the molecular diagnostic lab, specialized for PCR testing of infectious diseases. Plasma from each sample was separated by centrifugation and stored at -20° C until analyzed.

2.2 Nucleic Acid Extraction and Amplification

Viral RNA was extracted from 140 μ l of plasma using QIAamp Viral RNA mini-kit of Qiagen (CAT. No. 52904) Germany. The PCR quantification of samples was carried out by Artus RT-PCR kit for HIV and HCV by Qiagen, for all randomly selected samples, regardless of their serological result. Standard procedures, as proposed by Kwok and Higushi [1989], were followed to avoid contamination.

2.3 Study design and statistical analysis

All study subjects were distributed in three groups, according to their age as, minors (< 18 years), adult (>18 and < 50 years) and old (>50 years). They were also distributed according to gender; in three groups as follows: male, female and transgender. The mean age and inter quartile range were also calculated by statistical formulas. All data were analyzed by IBM SPSS Statistics 21 (SPSS Inc., Chicago, IL). Study variables include gender, age and viral loads of HIV type-1 and HCV.

3. Results & Discussion

The overall prevalence of co-infection, HIV and HCV monoinfection among walk-in patients of Jinnah Hospital, was 29 (37.2%), 15 (19.2%) and 8 (10.3%) respectively. Among 78 total participants, 26 (33.3%) had no infection of HIV and HCV. Of the study population (n=78), 25.6% (20/78); 74.4% (58/78) were female and males respectively. The mean age of the sample population, irrespective of gender, was 30 years with an inter-quartile range of 24 to 40. The youngest positive co-infected person was male of seven years and the oldest one male of 59 years.



Figure 1: Prevalence of HIV and HCV mono or co-infection in sample population (n=78)

As shown in table 1 and figure 2; among 58 male participants, 23 (39.7%) had HIV and HCV co-infection, 13 (22.4%) carried the HIV mono - infection, 5 (8.6%) carried

the HCV mono - infection and rest 17 (29.3%) had no infection of both viruses. Among twenty female, the distribution of HIV or HCV mono-infection and co-infection of both (HIV and HCV) was 2 (10%), 3 (15%) and 6 (30%), respectively. Nine females, representing 45% of the female population (n=20), had no infection of both HIV and HCV. There was a positive correlation between the two variables; HIV and HCV viral loads, as r = 0.736, n=29, p= 0.001. Figure 1 shows the prevalence of HIV and HCV co-infection, HIV type 1 and HCV mono-infection in the study population, while table 2 shows the number of cases of HIV and HCV co-infection, mono-infection and no infection of both viruses in different age groups.

 Table 1: Prevalence of HIV and HCV in different gender

 groups

Broups											
	Male (n=58)		Female (n=20)		Trans-gender						
Coinfection	23	39.7 %	6	30 %	0	0 %					
HIV only	13	22.4 %	2	10 %	0	0%					
HCV only	5	8.6 %	3	15 %	0	0 %					
No infection	17	29.3 %	9	45 %	0	0%					

The results we present here show the prevalence of HIV type-1 and HCV co-infection among walk-in patients of Lahore region. The analysis revealed increased prevalence of co-infection of both persistent viruses. An estimated five to ten million people in the world are living with HIV and HCV co-infection [15]. The detection of HCV RNA, during initial HIV infection, is possible after two to fourteen days of exposure [16]. The seroconversion of Hepatitis C Virus is delayed in HIV positive co-infected individuals, possibly due to the immunosupression trade mark of HIV infection. Only three studies revealed zero to thirteen percent incidence of seronegative chronic Hepatitis C virus infection [17, 18, 19]. Such silent and persistent infection not only alter the course of HIV disease progression, but also results an ineffective treatment and its outcomes [20,21]. Therefore, on time and accurate diagnosis is vital for survival.



infection in different gender groups

This study established the presence of HIV and HCV coinfection among general, the random infected population of Punjab. Among the HIV positive individuals (n=43) of our study, 67.5 % were co-infected with Hepatitis C Virus. The

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HCV only

No infection

0

1

0%

1.3 %

highest (90%) reported HIV and HCV co-infection was among jail inmates of Sindh, Pakistan [22]; followed by 73.74% in Lahore inmates [23]. The highest prevalence of co-infection; 32.1 % has been seen among the adult population of our study. The prevalence of HCV co-infection was less in our study as compared to the reported prevalence; 53.54% of Nafees *et al* (2011) [23].

Table 2: Prevalence of HIV and HCV in different age

groups											
	Minor ≤ 18 years		Adult >18 & < 50		$Old \ge 50$ years						
				years							
Coinfection	3	3.8%	25	32.1 %	1	1.3 %					
HIV only	0	0 %	11	14.1 %	3	3.8%					

In the sample population, no significant statistical relation was found between age and gender groups with an incidence of mono or co-infections. However, in our study, the male population (74%) as compared to the female population (26%) had a higher prevalence of HIV and HCV mono/coinfections. Our study has a higher percentage of male population as compared to other recently reported study [24], but less as compared to the study of jail inmates in Lahore [23]. Three minor individuals in our study; aged 7, 15 and 17 years, had co-infection of HIV and HCV; among them one was female of 15 years.

6

23

7.7 %

29.5 %

2

3

2.6 %

3.8%



Figure 3: Prevalence of HIV and HCV in different age groups (n=78)

A prevalence of coinfection in a minor population (3.8 %) was observed in our data which was not reported before in any other studies [23,24]. The mean age of our study group was 30 years old. This mean age was equal to the sample population of a previously reported study [25] but, more as compared to other reported studies from this region irrespective of their risk population and gender [23,24].

This study was aimed to see any kind of correlation between viral load of HIV type-1 and HCV co-infected individuals. Pearson's correlation model showed evidence of significant (P<0.001) supporting results of our hypothesis (CI=95%). Positive and less than one value of r (r = 0.736), indicated strong relationship between viral load and correlation in pathogenesis of HIV type-1 and HCV. This correlation indicated that increase in HIV type-1 viral load associated with an increase in HCV viral load. Many other studies have shown similar results of correlation between HIV type-1 viral load an HCV viral load [26 - 32]. Immediate prevention and surveillance measures are needed to control the spread of mono-transmission as well as co-transmission of HIV type-1 and HCV infections.

4. Conclusion

The prevalence rate of coinfection of HIV type-1 and Hepatitis C virus is much higher in our study conducted in a metropolitan city, Lahore, in Pakistan. Co-infection of Hepatitis C Virus is associated with increase disease progression of both viruses, HIV type-1 and HCV, irrespective of gender and age, as evident from increased viral loads.

References

- [1] A.Y. Kim and R.T. Chung, "Coinfection with HIV-1 and HCV—a one-two punch," Gastroenterology, 137(3), pp. 795–814, 2009.
- [2] B. Polsky, A.Y. Kim, R.T. Chung, "Human Immunodeficiency Virus and Hepatitis B and C coinfection: pathogenic interactions, natural history and therapy," AIDS Clinical Review, pp. 263–306, 2000– 2001.
- [3] V. Soriano, E. Vispo, P. Labarga, J. Medrano, P. Barreiro, "Viral Hepatitis and HIV co-infection" Antiviral Research, 85, pp. 303–15, 2010.
- [4] I.L. Bica, B. Mcgovern, R. Dhar, D. Stone, K. Mcgowan, R. Scheib, D.R. Snydman, "Increasing mortality due to end-stage liver disease in patients with Human Immunodeficiency Virus infection" Clinical Infectious Diseases, 32, pp. 492–497, 2001.
- [5] D.L. Thomas and L. B. Seeff, "Natural history of Hepatitis C," Clinics in Liver Disease, 9(3), pp. 383– 98, 2005.
- [6] K. Sherman, S. Rouster, R. Chung, N. Rajicic, "Hepatitis C Virus prevalence among patients infected with Human Immunodeficiency Virus: A crosssectional analysis of the US adult AIDS clinical trials groups," Clinics In Liver Disease, 34, pp. 831–837, 2002.

- [7] T.yee, A. Griffioen, C. Sabin, G. Dusheiko, C. Lee, "The natural history of HCV in a cohort of hemophiliac patients infected between 1961 and 1985," Gut, 47, pp. 845–851, 2000.
- [8] Y. Waheed, T. Shafi, S.Z. Safi, I. Qadri, "Hepatitis C Virus in Pakistan: A systematic review of prevalence, genotypes and risk factors," World Journal of Gastroenterology, 15, pp.5647-5653, 2009.
- [9] Y. Waheed, T.B. Rahat, S.Z. Safi, I. Qadri, "Epidemiological patterns and risk factors associated with Hepatitis B virus in Pakistani population," Clinics in Liver Disease, 4, pp. 547-554, 2010.
- [10] F. Sultan, T. Mehmood, M.T. Mahmood, "Infectious pathogens in volunteer and replacement blood donors in pakistan: a ten-year experience," International Journal of Infectious Diseases, 11, pp. 407-412, 2007.
- [11] V. Di Martino, P. Rufat, N. Boyer, P. Renard, F. Degos, M. Martinot-Peignoux, S. Matheron, V. Le Moing, F. Vachon, C. Degott, D. Valla, P. Marcellin, "The influence of Human Immunodeficiency Virus coinfection on chronic Hepatitis C in injection drug users: a long-term retrospective cohort study," Hepatology, 34, pp. 1193–1199, 2001.
- [12] R. Bartenschlager, V. Lohmann, "Replication of Hepatitis C Virus," The Journal of General Virology, 81, pp. 1631–48, 2000.
- [13] Y.H. Zheng, N. Lovsin, B.M. Peterlin, "Newly identified host factors modulate HIV replication. Immunology Letters, 97 (2), pp. 225–34, 2005.
- [14] A. Maheshwari, S. Ray, P.J. Thuluvath, "Acute Hepatitis C. Lancet, 372, pp. 321–332, 2008.
- [15] E.A. Operskalski, A. Kovacs, "HIV/HCV co-infection: pathogenesis, clinical complications, treatment, and new therapeutic technologies," Current HIV/AIDS Reports, 8, pp.12-22, 2011.
- [16] B. Hajarizadeh, J. Grebely, G.J. Dore, "Epidemiology and natural history of HCV infection," Nature Reviews Gastroenterology & Hepatology, 10, pp. 553-562, 2013.
- [17] G. Chamie, M. Bonacini, D.R. Bangsberg, J.T. Stapleton, C. Hall, E.T. Overton, R. Scherzer, P.C. Tien, "Factors associated with seronegative chronic Hepatitis C Virus infection in HIV infection," Clinical Infectious Diseases, 44, pp. 577-583, 2007.
- [18] C.L. Thio, K.R. Nolt, J. Astemborski, D. Vlahov, K.E. Nelson, D.L. Thomas, "Screening for Hepatitis C Virus in Human Immunodeficiency Virus-infected individuals," Journal of Clinical Microbiology, 38, pp. 575-577, 2000.
- [19] E.C. Homson, E. Nastouli, J. Main, P. Karayiannis, J. Eliahoo, D. Muir, M.O. Mcclure, "Delayed anti-HCV antibody response in HIV-positive men acutely infected with HCV," AIDS, 23, pp. 89-93, 2009.
- [20] L. Hua, J.W. Andersen, E.S. Daar, M.J Glesby, K. Hollabaugh, C. Tierney, "Hepatitis C Virus/HIV coinfection and responses to initial antiretroviral treatment. *AIDS*, 27, pp. 2725-2734.
- [21] G. Antonucci, E. Girardi, A. Cozzi-Lepri, M.R. Capobianchi, A. De Luca, M. Puoti, E. Petrelli, G. Carnevale, G. Rizzardini, P.A. Grossi, P. Vigano, M.C. Moioli, F. Carletti, M. Solmone, G. Ippolito, A.D. Monforte, "Role of Hepatitis C Virus (HCV) viremia and HCV genotype in the immune recovery from

highly active antiretroviral therapy in a cohort of antiretroviral-naive HIV-infected individuals," Clinical Infectious Diseases, 40, pp. e101-e109, 2005.

- [22] S. Safdar, A. Mehmood, S.A. Abbas, "Prevalence of HIV/AIDS among jail inmates in Sindh," Journal of Pakistan Medical Association, 59(2), pp. 111-112, 2009.
- [23] M. Nafees, A. Qasim, G. Jafferi, M.S. Anwar, M. Muazzam, "HIV infection, HIV/HCV and HIV/HBV co-infections among jail inmates of Lahore," Pakistan Journal of Medical Sciences, 27(4), pp. 837-841, 2011.
- [24] Y. Khan, S. Ali, M.B. Awan, "Frequency of newly diagnosed Hepatitis B and C Viruses in patients with Human Immunodeficiency Virus and their common leading factors," Khyber Journal Of Medical Sciences, Vol. 6, No. 2, 2013.
- [25] R.N. Samo, A. Altaf, A. Agha, O. Pasha, S. Rozi, A. Memon, S. Azam, M. Blevins, S.H. Vermund, S.A. Shah, "High HIV incidence among persons who inject drugs in pakistan: greater risk with needle sharing and injecting frequently among the homeless," PLoS ONE, 8(12), e81715, 2013.
- [26] K.E. Sherman, J. O'Brien, A.G. Guteierrez, S. Harrison, M. Urdea, P. Neuwald, J. Wilber, "Quantitative evaluation of Hepatitis C Virus RNA in patients with concurrent Human Immunodeficiency Virus infection, Journal of Clinical Microbiology, 31, pp. 2679-82, 1993.
- [27] B. Cribier, D. Rey, C. Schmitt, J. M. Lang, A. Kirn, F. Stoll-Keller, "High Hepatitis C viremia and impaired antibody response in patients coinfected by HIV, *AIDS*, 9, pp. 1131-6, 1995.
- [28] D. L. Thomas, J. W. Shih, H. J. Alter, D. Vlahov, S. Cohn, D.R. Hoover, L. Cheung, K. E. Nelson, "Effect of Human Immunodeficiency Virus on Hepatitis C Virus infection among injecting drug users," Journal Of Infectious Diseases, 174, pp. 690-5, 1996.
- [29] C. Mazza, A. Ravaggi, M. Pouti, A. Albertini, E. Cariani, "Increased HCV titer and absence of selection of E2-hypervariable region (HVRI) in HCV/HIV coinfection," Journal of Hepatology, pp. 21:s34, 1994.
- [30] M. E. Eyster, M.W. Fried, A. M. Di Bisceglie, J. J. Goedert, "Increased Hepatitis C Virus RNA levels in hemophiliacs: relationship of Human Immunodeficiency Virus infection and liver disease. *Blood*, 84, pp.1020-3, 1994.
- [31] M. Beld, M. Penning, V. Lukashov, M. Mcmorrow, M. Roos, N. Pakker, J. Goudsmit, "Evidence that both HIV and HIV-induced immunodeficiency enhance HCV replication among HCV seroconverters", Virology, 244, pp. 504-12, 1998.
- [32] B. Cribier, C. Schmitt, D. Rey, G. Uhl, J. M. Lang, D. Vetter, F. Stoll-keller, "HIV increased Hepatitis C viremia irrespective of the Hepatitis C Virus genotype," Research in Virology, 148, pp. 267-71, 1997.

Author Profile



Prof. Dr. Fouzia Ashraf, MBBS, M. Phil, Ph. D; is a Professor of Pathology-teaching faculty at Allama Iqbal Medical College, Consultant Pathologist, and Molecular Biologist (Virologist) at Jinnah Hospital in

Volume 4 Issue 2, February 2015 www.ijsr.net Lahore, Pakistan.



Muhammad Tajamal received his B.Sc. in Medical Lab Technology, M.Sc. in Molecular Biology and MS in Forensic Chemistry. He is working as Medical Lab Technologist and Molecular Technologist in Molecular Diagnostics Lab of Allama Iqbal Medical College & Jinnah Hospital and Hameed Latif Hospital Lahore,

Pakistan, respectively.



Shahzeb Javed is a third year medical student at King Edward Medical University, Lahore, Pakistan. He completed his B.Sc from the University of Michigan, Ann Arbor, and has conducted research studying airway hyper-reactivity and asthma, and child psychiatric disorders at the University of Liverpool,

UK. His current research interests are in the field of Infectious Disease.



Saamia Tahir is a third year medical student at CMH Lahore Medical College, Lahore, Pakistan. She has conducted extensive bench research at National Jewish Hospital in Denver, Colorado related to airway disorders.



Omar Ali is a second year medical student at CMH Lahore Medical College, Lahore, Pakistan. He has actively been involved in assisting research projects related to the atherosclerotic process at the Punjab Institute of Cardiology.



Prof. Dr. Mahmood Shaukat received his M.B.B.S, B.Sc in English from Lahore and F.R.C.S in Paediatric Surgery from Dublin. He is currently working as principal of Allama Iqbal Medical College / Jinnah Hospital, Lahore and Professor of Paediatric Surgery. He is founding as well as board member of various

academic and social welfare organizations.