

**ORIGINAL RESEARCH**

# The spectrum of hepatic abnormalities including sero-prevalence of hepatitis B and hepatitis c co-infection in HIV infected individuals and their correlation with CD4 count

Sunil Pawar<sup>1</sup>, Gourav Bansal<sup>2</sup>, Vijay Kumar<sup>3</sup>, Priyanka<sup>4</sup><sup>1</sup>Assistant Professor, Department of Medicine, Government Medical College, Patiala, Punjab, India<sup>2</sup>DNB Resident, Department of Gastroenterology and Hepatology, Fortis Hospital, Ludhiana, Punjab, India<sup>3</sup>Professor, Department of Medicine, MM Medical College & Hospital, Kumarhatti, Solan, Himachal Pradesh, India<sup>4</sup>Assistant Professor, Department of Radiodiagnosis, Dr BR Ambedkar State Institute of Medical Sciences, Mohali, Punjab, India**Corresponding Author**

Priyanka

Assistant Professor, Department of Radiodiagnosis, Dr BR Ambedkar State Institute of Medical Sciences, Mohali, Punjab, India

**Email:** [docpriyanka.87@gmail.com](mailto:docpriyanka.87@gmail.com)

Received Date: 18 July, 2024

Acceptance Date: 16 August, 2024

**ABSTRACT**

**Background:** To study the spectrum of hepatic abnormalities including sero-prevalence of hepatitis B and hepatitis C coinfection in HIV infected individuals and their correlation with CD4 count. There had been little work done on liver function tests in HIV patients without pre-existing liver disease. **Method:** An observational study was conducted on 200 HIV patients above 18 years of age and excluded those who were already diagnosed with Hep B or Hep C, co-existing malignancy, obstructive biliary disease and those with heavy alcohol consumption. **Result:** In our study sero-prevalence of Hepatitis B co-infection among HIV individuals was 4% (n=8). In our study sero-prevalence of Hepatitis C co-infection among HIV individuals was 15% (n=30). In our study sero-prevalence of Hepatitis B and Hepatitis C co-infection among HIV individuals was 1.5% (n=3). Overall, 34% (n=68) patients had abnormal Liver Function Tests. Out of 200 patients studied, 13 patients (6.5%) had serum bilirubin of more than 1 mg/dl and 187 patients (93.5%) had serum bilirubin in normal range. 137 patients (68.5%) had AST level up to 40 IU/L and 63 patients (31.5%) had AST level of more than 40 IU/L. Out of these 63 patients, 61 patients (96.8%) were on TLE regimen and 2 patients (3.2%) were on ZLN regimen. Probable etiologies for deranged AST level in 63 patients were HIV/HBV co-infection in 7.9% patients (n=5), HIV/HCV co-infection in 28.6% patients (n=18), NAFLD in 9.6% patients (n=6) and HIV infection itself or cART or unknown causes in the remaining 53.9% patients (n=34). **Conclusion:** Elevated liver transaminases are now a common finding in HIV-infected patients. The main etiologies for hepatic morbidities in HIV-infected patients include viral hepatitis coinfection, NAFLD, ART-related liver injury and infection-related factors. Thus, regular liver function tests along with non-invasive methods of investigation of hepatobiliary system needs to be done in every individual infected with HIV.

This is an open access journal, and articles are distributed under the terms of the Creative Commons Attribution- Non Commercial-Share Alike 4.0 License, which allows others to remix, tweak, and build upon the work non-commercially, as long as appropriate credit is given and the new creations are licensed under the identical terms.

**INTRODUCTION**

Hepatobiliary diseases occur commonly in patients with human immunodeficiency virus (HIV) infection and are now the commonest causes of death in HIV positive patients on antiretroviral therapy (ART) in western countries. <sup>[1,2]</sup> Liver enzyme abnormalities have been reported in 20 – 93% of HIV-infected populations. <sup>[3,4]</sup>

Human immunodeficiency virus (HIV) infection of liver cells contributes to liver disease progression by both direct and indirect mechanisms. Gp120 binding to CXCR4 may induce hepatocyte apoptosis and activation of HSCs, both contributing to fibrosis. HIV infection of the gastrointestinal tract produces pro-inflammatory cytokines and chemokines which attract activated lymphocytes and monocytes to the liver

which may further drive fibrosis.<sup>[5]</sup>

Most liver disease among HIV-infected individuals is secondary to coinfection with HCV and/or HBV <sup>[7]</sup> due to both common modes of transmission and geographic patterns of disease. HIV infection alters the natural history of HCV in several ways. Coinfected individuals also have higher HCV RNA levels, accelerated progression to hepatic fibrosis, an increased risk of developing cirrhosis, and a higher risk of decompensated liver disease once cirrhotic. <sup>[8-10]</sup> Individuals with HIV-HBV co-infection are 3-6 times more likely to develop chronic HBV after an acute exposure than individuals without HIV infection and lower rate of spontaneous clearance of HBeAg, increased HBV replication, and a higher rate of loss of anti-HBs and reactivation of HBV.<sup>[11]</sup>

Liver toxicity is one of the most common serious adverse events associated with ART. <sup>[12]</sup> due to direct drug toxicity and/or drug metabolism, hypersensitivity reactions, mitochondrial toxicity, and immune reconstitution inflammatory syndrome [IRIS]. <sup>[12-14]</sup> Common ART Agents Associated with Liver Injury in HIV infected patients <sup>[6]</sup>. NNRTI- Nevirapine, Efavirenz, NRTI- Abacavir, Protease Inhibitor- Ritonavir, Tipranavir, CCR5 Inhibitor- Maraviroc. Other prevalent factors affecting liver function are alcohol abuse and Non-alcoholic fatty

liver disease (NAFLD)

## MATERIALS AND METHODS

A hospital based observational study was conducted in the department of Medicine at Rajindra Hospital, Patiala on 200 patients (20 in-hospital and 180 OPD/ART) above 18 years of age, after obtaining informed consent and ethical clearance. HIV patients on treatment as per NACO guidelines were included in this study. Patients who were already diagnosed with Hep B or Hep C, co-existing malignancy, obstructive biliary disease and those with heavy alcohol consumption were excluded from the study population.

Each participant was subject to detailed clinical history and examination and blood investigations including liver function tests were done. Imaging studies including Ultrasound and Fibroscan were performed. Liver function was also assessed using various scores such as APRI and FIB-4.

All indoor patients were managed as per symptoms and associated co-morbidities.

All statistical analysis was done on SPSS version 18, Pearson Chi square test was used for studying correlation of different variables. p-value less than 0.05 was considered significant and less than 0.001 was considered highly significant.

## RESULTS

**Table 1: Baseline characteristics of study population**

	Mean	Std. Deviation
Age	37.85	12.105
CD4 count	493.43	297.502
Duration of ART	4.14	2.391
S Bilirubin	.558600	.3147561
AST	39.18	30.472
ALT	39.06	33.154
Total serum protein	6.599500	.3853469
Albumin	3.628500	.4752548
Globulin	2.986000	.3816178
INR	1.031050	.0861773
Fibroscan	6.425500	6.2142000
APRI	.566800	.8941899
FIB-4	1.379250	1.7227960

The above table shows the baseline characteristics of study population. Mean age 37.85±12.105 yrs (range 18-70 yrs). Mean CD4 cell count 493.43±297.502 cells/mm<sup>3</sup> (range 22-1857 cells/mm<sup>3</sup>). Mean duration of treatment with ART 4.14±2.25 years (range 1-11 years). Mean Hemoglobin level 11.32±2.25 gm% (range 4-17.6 gm%). Mean Platelet count 225835±90305 /ul (range 29000-564000 /ul). Mean Bilirubin level was 0.55±0.31 mg/dl (range 0.2-2.1 mg/dl). Mean AST level 39.18±30.42 IU/L (range 13-

243 IU/l). Mean ALT level 39.06±33.15 IU/L (range 11-295 IU/L). Mean total serum protein level 6.59±.38 gm/dl (range 5.1-7.8 gm/dl). Mean albumin level 3.62±0.47 gm/dl (range 2-4.4 gm/dl). Mean globulin level 2.98±0.38 gm/dl (range 2.3-4.8 gm/dl). Mean INR was 1.03±.08 (range 0.9-1.5). Mean Fibroscan value 6.42±6.21 KPa (range 3.1-41.0 KPa). Mean APRI score 0.56±0.89 (range 0.07-10.43). Mean FIB 4 score 1.37±1.72 (range 0.21-16.53).

**Table 2: Liver parameters in Study Population**

Parameters		Frequency	Percent
Serum bilirubin	≤1	187	93.5
	>1	13	6.5
AST	Up to 40	137	68.5
	>40-120	56	28
	>120	7	3.5
ALT	Up to 40	132	66.0
	>40-120	62	31.0
	>120	6	3.0
USG Whole Abdomen	WNL	165	82.5
	Cholelithiasis	5	2.5
	Hepatomegaly	3	1.5
	Hepatosplenomegaly	4	2.0
	Fatty liver	13	6.5
	Cirrhosis liver	10	5.0
Fibroscan	<7.1KPa	157	78.5
	7.1-12.5KPa	31	15.5
	>12.5KPa	12	6.0
APRI Score	<0.5	140	70.0
	0.5-1.5	49	24.5
	>1.5	11	5.5
FIB-4 Score	<1.45	146	73.0
	1.45-≤3.25	44	22.0
	>3.25	10	5.0

Out of 200 patients, 6.5% (n=13) population had serum bilirubin of more than 1 mg/dl and 93.5% (n=187) had serum bilirubin in normal range.

Out of 200 patients, 68.5% (n=137) patients had AST level up to 40 IU/L and 31.5% (n=63) had AST level of more than 40 IU/L. 7 patients had AST level of >3 times upper normal limit of AST.

Out of 200 patients, 66% (n=132) patients had ALT level up to 40 IU/L and 34% (n=68) had ALT level of more than 40 IU/L. 6 patients had ALT level of >3 times upper normal limit of ALT.

Out of 200 patients, 5.0% (n=10) had features of cirrhosis of liver, 6.5% (n=13) had features of fatty liver, 2% (n=4) had hepatosplenomegaly, 2.5% (n=5) had Cholelithiasis, 1.5% (n=3) had hepatomegaly

(liver span >15 cm) and rest 82.5% (n=165) had ultrasound findings within normal limit.

Liver Stiffness values >7.1 kPa (equivalent to Metavir F ≥ 2) were considered abnormal. In this study population 21.5% had LSM >7.1KPa. On the basis of liver stiffness values, F2-F3 and F4 Metavir stages were evidenced in 15.5% (n=31) and 6.0% (n=12) of patients.

In this study, 24.5% (n=49) patients had an APRI score of 0.5-1.5 and 5.5% (n=11) patients had an APRI score of >1.5.

In this study, 22% (n=44) patients had FIB-4 score of 1.45-≤3.25 and 5% (n=10) patients had FIB-4 score of >3.25.

**Table 3: Association of CD4 Count with Duration of cART**

CD4 Count		Duration of ART			Total	
		<2	2-5	>5		
≤200	Count	6	10	7	23	
	%	26.1%	43.5%	30.4%	100.0%	
201-500	Count	17	48	25	90	
	%	18.9%	53.3%	27.8%	100.0%	
>500	Count	10	51	26	87	
	%	11.5%	58.6%	29.9%	100.0%	
Total		Count	33	109	58	200

As shown in table 3, 23 patients who had CD4 count ≤200 cells/mm<sup>3</sup>, 6 patients had duration of cART <2 years, 10 patients had duration of cART 2-5 years and 7 had duration of cART more than 5 years. Of 90 patients who had CD4 count in the range of 201-500

cells/mm<sup>3</sup>, 17 patients had duration of cART <2 years, 48 patients had duration of cART 2-5 years and 25 had duration of cART more than 5 years. Of 87 patients who had CD4 count >500 cells/mm<sup>3</sup>, 10 patients had duration of cART <2 years, 51 patients

had duration of cART 2-5 years and 26 had duration of cART more than 5 years. On Statistical analysis, the difference in CD4 count with duration of ART was not significant (p value 0.43).

**Table 4: Association of AST and ALT with type of cART regimen**

ART drugs		AST level		Total	ALT		Total
		≤40	>40		≤40	>40	
TLE	Count	120	61	181	116	65	181
	% within cART regimen	66.3%	33.7%	100.0%	64.1%	35.9%	100.0%
ZLN	Count	17	2	19	16	3	19
	% within cART regimen	89.5%	10.5%	100.0%	84.2%	15.8%	100.0%
Total	Count	137	63	200	132	68	200
	% within total pts on cART drugs	68.5%	31.5%	100.0%	66.0%	34.0%	100.0%

As shown in table 4, 23 patients who had CD4 count ≤200 cells/mm<sup>3</sup>, 6 patients had duration of cART<2 years, 10 patients had duration of cART 2-5 years and 7 had duration of cART more than 5 years. Of 90 patients who had CD4 count in the range of 201-500 cells/mm<sup>3</sup>, 17 patients had duration of cART<2 years, 48 patients had duration of cART 2-5 years and 25

had duration of cART more than 5 years. Of 87 patients who had CD4 count >500 cells/mm<sup>3</sup>, 10 patients had duration of cART<2 years, 51 patients had duration of cART 2-5 years and 26 had duration of cART more than 5 years. On Statistical analysis, the difference in CD4 count with duration of ART was not significant (p value 0.43).

**Table 5: Association of AST and ALT with patients of HIV/HBV coinfection, HIV/HCV coinfection and fatty liver**

			HBsAg			Anti-HCV antibody			Fatty Liver		
			Non-reactive	Reactive	Total	Non-reactive	Reactive	Total	No fatty liver	Fatty liver	Total
AST Level	≤40	N	134	3	135	125	12	137	130	7	137
		%	69.8	37.5	68.5	73.5	40	68.5	69.5	53.8	68.5
	>40	N	58	5	63	45	18	63	57	6	63
		%	30.2	62.5	31.5	26.5	60	31.5	30.5	46.2	31.5
Total			192	8	200	170	30	200	187	13	200
P value			0.054			<0.001			0.434		
ALT Level	≤40	N	129	3	132	118	14	132	126	6	132
		%	67.2	37.5	66	69.4	46.7	66	67.4	46.1	66
	>40	N	63	5	68	52	16	68	61	7	68
		%	32.8	62.5	34	30.6	53.3	34	32.6	53.9	34
Total			192	8	200	170	30	200	187	13	200
P value			0.082			0.015			0.228		

As shown in table 5, 31.5% (n=63) patients had AST level above upper limit. Out of 181 patients in the TLE group, 61 (33.7%) patients had deranged AST levels. Out of 19 patients in the ZLN group, 2 (10.5%) patients had deranged AST levels. This association was significant on statistical analysis (p value 0.03).

As shown in table 5, 34% (n=68) patients had ALT level above upper limit. Out of 181 patients in the TLE group, 65 (35.9%) patients had deranged ALT levels. Out of 19 patients in the ZLN group, 3 (15.8%) patients had deranged ALT levels. This association was significant on statistical analysis (p value 0.03).

**Table 6: Association of AST and ALT with patients of HIV/HBV/HCV coinfection**

No of patients with HIV/HBV/HCV coinfection	No of patients with AST Level>40 IU/L	No of patients with ALT Level>40 IU/L
3	3	3

As shown in table 6, 8 patients had HIV/HBV coinfection. Out of 8 patients with coinfection 5 (62.5%) patients had deranged AST level and 3 (37.5%) patients had AST level in normal range. This

association was insignificant on statistical analysis (p value 0.054). 30 patients had HIV/HCV coinfection. 18 (60.0%) patients with coinfection had deranged AST level and

12 (40%) patients had AST level in normal range. This association was highly significant on statistical analysis (p value <0.001).

13 patients had fatty liver. 6 patients (46.2%) had

deranged AST level and 7 patients (53.8%) had AST level in normal range. This association was insignificant on statistical analysis (p value 0.434).

**Table 7: Association of CD4 Count with AST and ALT levels**

CD4 Count		AST group		Total	ALT group		Total
		≤40	>40		≤40	>40	
≤200	Count	13	10	23	14	9	23
	% within CD4 grp	56.5%	43.5%	100.0%	60.9%	39.1%	100.0%
201-500	Count	69	21	90	65	25	90
	% within CD4 grp	76.7%	23.3%	100.0%	72.2%	27.8%	100.0%
>500	Count	55	32	87	53	34	87
	% within CD4 grp	63.2%	36.8%	100.0%	60.9%	39.1%	100.0%
Total	Count	137	63	200	132	68	200
	% within CD4 grp	68.5%	31.5%	100.0%	66.0%	34.0%	100.0%

As shown in table, above 10 patients had cirrhosis. 8 patients (80.0%) had deranged AST level and 2 patients (20%) had AST level in normal range. This association was significant on statistical analysis (p value 0.001).

As shown in table above, 7 patients (70.0%) had deranged ALT level and 3 patients (30%) had ALT level in normal range. This association was significant on statistical analysis (p value 0.014)

## DISCUSSION

The present study was conducted to study the spectrum of hepatic abnormalities and Sero-prevalence of Hepatitis-B and Hepatitis-C coinfection in HIV infected individuals and their correlation with CD4 count. In the present study mean age was found to be 37.85±12.1 which was comparable to studies conducted by Ocama P et al, Pathania MS et al and Bajpai S et al. In the present study, mean of CD4 cells was found to be 493.43±297.5 cells/mm<sup>3</sup> which was comparable to study conducted by Pathania MS et al. Mean CD4 count was 493.43±297.5 cells/mm<sup>3</sup> (range 22-1857 cells/mm<sup>3</sup>). In our study Sero-prevalence of Hepatitis B co-infection among HIV Individuals was 4% (n=8). In our study Sero-prevalence of Hepatitis C co-infection among HIV Individuals was 15% (n=30). In our study Sero-prevalence of Hepatitis B and Hepatitis C co-infection among HIV Individuals was 1.5% (n=3).

Overall, 34% (n=68) patients had abnormal Liver Function Tests. Out of 200 patients studied, 13 patients (6.5%) had serum bilirubin of more than 1 mg/dl and 187 patients (93.5%) had serum bilirubin in normal range. 137 patients (68.5%) had AST level up to 40 IU/L and 63 patients (31.5%) had AST level of more than 40 IU/L. Out of these 63 patients, 61 patients (96.8%) were on TLE regimen and 2 patients (3.2%) were on ZLN regimen. Probable etiologies for deranged AST level in 63 patients were HIV/HBV co-infection in 7.9% patients (n=5), HIV/HCV co-infection in 28.6% patients (n=18), NAFLD in 9.6% patients (n=6) and HIV infection itself or cART or unknown causes in the remaining 53.9% patients

(n=34). 132 patients (66%) had ALT level up to 40 IU/L and 68 patients (34%) had ALT level of more than 40 IU/L. Out of these 68 patients, 65 patients (95.6%) were on TLE regimen and 3 patients (4.4%) were on ZLN regimen. Probable etiologies for deranged ALT level in 68 patients were HIV/HBV co-infection in 7.4% patients (n=5), HIV/HCV co-infection in 23.5% patients (n=16), NAFLD in 10.3% patients (n=7) and HIV infection itself or cART or unknown causes in the remaining 58.8% (n=40) patients. Mean Total serum protein was 6.59±.38 gm/dl (range 5.1-7.8 gm/dl). Mean INR value 1.03±.08 (range 0.9-1.5).

In our study, 5.0% (n=10) patients had features of cirrhosis of liver, 6.5% (n=13) had features of fatty liver, 2% (n=4) had hepatosplenomegaly, 2.5% (n=5) had Cholelithiasis, 1.5% (n=3) had hepatomegaly and rest 82.5% (n=165) had ultrasound findings with in normal limit. Out of 10 patients who had cirrhosis, 9 patients had HCV/HIV coinfection and 1 patient had HBV/HIV co-infection. In this study population 21.5% had LSM>7.1KPa on fibroscan. 5.5% (n=11) patients had advanced fibrosis as per APRI score >1.5. 5% (n=10) patients had advanced fibrosis as per FIB-4 score >3.25. No statistically significant association was found in CD4 count with AST levels, CD4 count with ALT levels

In the present study, 4% of patients had both HIV/HBV coinfection. The discordance with other studies as conducted by Rathi PM et al(1997)<sup>[16]</sup>, Ocama P et al(2008)<sup>[17]</sup>, Luma HN et al(2016)<sup>[18]</sup> can be explained as in India, chronic hepatitis B affects around 2- 8 % of PLHIV, though there are wide variations in different parts of the country.<sup>[159]</sup>

In the present study, 15% of patients had both HIV/HCV coinfection. Hepatitis C virus (HCV) affects 5- 15% of PLHIV, rising to 90% among people who inject drugs. Because of shared routes of transmission, certain groups, particularly persons who inject drugs (PWID) have high rates of co-infection with HIV and HCV. Globally around 67% of PWID are infected with HCV. Other high-risk groups are those who have a high frequency of injections and a

low level of infection control, transmission from HCV infected mothers and people with HCV infected sexual partners. Persons with HIV infection, in particular MSM, are at increased risk of HCV infection through unprotected sex. Tattoo recipients have higher prevalence of HCV compared with persons without tattoos.<sup>[59]</sup>

In present study, 10 (43.5%) out of 23 patients with CD4 counts  $\leq 200$  cells/mm<sup>3</sup> and 53 (29.94%) out of 177 patients with CD4 counts  $>200$  cells/mm<sup>3</sup> had deranged AST level. 9 (39.1%) out of 23 patients with CD4 counts  $\leq 200$  cells/mm<sup>3</sup> and 59 (33.33%) out of 177 patients with CD4 counts  $>200$  cells/mm<sup>3</sup> had deranged ALT level. While in study conducted by Pathania MS et al<sup>[15]</sup> in 2016, 49 out of 97 patients (50.5%) with CD4 counts  $<200$  cells/mm<sup>3</sup> and 79 out of 150 patients (52.7%) with CD4  $>200$  cells/mm<sup>3</sup> had abnormal LFTs. This difference in the results of our study and previous study could be attributed to fact that our study had excluded patients with heavy alcohol intake, chronic hepatotoxic drug, any obstructive biliary pathology, co-existent malignant disease.

In the present study, 5% patients had cirrhosis, 6.5% had fatty liver, 1.5% had hepatomegaly and 2% had hepatosplenomegaly on Ultrasound abdomen. In study conducted by Savita M et al<sup>[19]</sup> 24% had fatty liver whereas in present study fatty liver was found in 6.5% of patients as patients with significant amount of alcohol consumption (more than 1 drinks per day in women and more than 2 drinks per day in men, 1 drink was defined as 10gm of ethanol) were excluded from our study. 2.5% had cholelithiasis which was an incidental finding.

The difference in fibroscan findings in present study as compared with previous study could be attributable to the fact that the number of HCV/HIV co-infected patients were 30 out of 200 and HBV/HIV co-infected patients were 8 out of 200. While, the number of HCV/HIV co-infected patients in previous study by Maggi P et al<sup>[20]</sup> were 148 out of 228 and remaining were HIV mono-infected.

In the present study, 10 out of 12 patients with LSM  $> 12.5$  KPa were noted to have cirrhosis on USG whole abdomen, of whom 9 patients were HIV/HCV co-infected and 1 was HIV/HBV co-infected.

The assessment of asymptomatic liver damage using LSM in patients mono-infected with HIV receiving long-term antiretroviral therapy has been rarely addressed. Even if liver biopsy is still considered the gold-standard method for assessment of liver fibrosis, Fibroscan is presently the main non-invasive method for diagnosis, staging, and follow-up of both viral and non-viral hepatopathies due to optimum accuracy standards, operator independency, absence of complications, and ease of use. Nevertheless, experience in the literature is still lacking regarding the use of this method for patients mono-infected with HIV.

In the present study, APRI scores of  $< 0.5$ , 0.5-1.5 and

$>1.5$  were noted in 70%, 24.5% and 5.5% of patients respectively. Of 5.5% of these, 5% were noted to have cirrhosis on ultrasound of the whole abdomen.

In the present study, FIB-4 scores of  $<1.45$ , 1.45-3.25 and  $>3.25$  were noted in 73%, 22% and 5% of patients respectively. All the patients with FIB-4 scoring of  $>3.25$  were noted to have cirrhosis on ultrasound of the whole abdomen.

There is increased seroprevalence of HBV and HCV coinfection in HIV/AIDS patients due to common mode of transmission. Overall, 34% patients had abnormal liver function tests which can be attributed to HIV infection itself, co-infection with HBV, HCV, cART and NAFLD. These findings confirm hepatic involvement to be a feature of HIV infection.

Alcohol abuse and Anti-tubercular drugs are also one of the common causes of liver injury but they were excluded from our study.

Ours is a small study, larger studies are needed to confirm the presence of hepatic morbidities in HIV infection.

## CONCLUSION

Elevated liver transaminases are now a common finding in HIV-infected patients. The main etiologies for hepatic morbidities in HIV-infected patients include viral hepatitis coinfection, NAFLD, ART-related liver injury and infection-related factors. Several simple fibrosis biomarkers based on liver transaminases like APRI, FIB-4 score and non-invasive investigations like ultrasonography whole abdomen and fibroscan can help in the management of the individual patient. Thus, regular liver function tests along with non-invasive methods of investigation of the hepatobiliary system needs to be done in every individual infected with HIV.

Alcohol abuse and Anti-tubercular drugs are also one of the common causes of liver injury but they were excluded from our study.

## BIBLIOGRAPHY

1. Lizardi-Cervera J, Soto Ramirez LE, Poo JL et al. Hepatobiliary diseases in patients with human immunodeficiency virus (HIV) treated with non-highly active antiretroviral therapy: frequency and clinical manifestations. *Ann Hepatol.* 2005; 4:188–91.
2. Lodenyo H, Schoub B, Ally R et al. Hepatitis B and C infections and liver function in AIDS patients at Chris Hani Baragwanath hospital, Johannesburg. *East Afr Med J.* 2000;77(1):13–15.
3. Dieterich DT, Robinson PA, Love J et al. Drug-induced liver injury associated with the use of nonnucleoside reverse-transcriptase inhibitors. *Clin Infect Dis.* 2004;(38 Suppl 2): S8.
4. Wnuk AM. Liver damage in HIV-infected patients. *Med Sci Monit.* 2001;7: 729–36.
5. Crane M, Iser D, and Lewin SR Human immunodeficiency virus infection and the liver *World J Hepatol.* 2012 Mar 27; 4(3): 91–98.
6. Price JC, Thio LL. Liver disease in HIV infected individuals. *Clinical Gastroenterology and Hepatology* 2010;8: 1002-12.

7. Weber R, Sabin CA, Friis-Moller N, et al. Liver-related deaths in persons infected with the human immunodeficiency virus: the D:A: D study. *Arch Intern Med* 2006;166: 1632–41.
8. Telfer P, Sabin C, Devereux H, et al. The progression of HCV-associated liver disease in a cohort of haemophilic patients. *Br J Haematol* 1994;87: 555–61.
9. Soto B, Sanchez-Quijano A, Rodrigo L, et al. Human immunodeficiency virus infection modifies the natural history of chronic parenterally-acquired hepatitis C with an unusually rapid progression to cirrhosis. *J Hepatol* 1997;26: 1–5.
10. Benhamou Y, Bochet M, Di Martino V, et al. Liver fibrosis progression in human immunodeficiency virus and hepatitis C virus coinfecting patients: the Multivirc Group. *Hepatology* 1999;30: 1054–58.
11. Thio CL. Hepatitis B and human immunodeficiency virus coinfection. *Hepatology* 2009;49(Suppl):S138–S45.
12. Reisler RB, Han C, Burman WJ, et al. Grade 4 events are as important as AIDS events in the era of HAART. *J Acquir Immune Defic Syndr* 2003;34: 379–86.
13. Puoti M, Nasta P, Gatti F, et al. HIV-related liver disease: ARV drugs, coinfection, and other risk factors. *J Int Association Physicians AIDS Care* 2009;8: 30–42.
14. Soriano V, Puoti M, Garcia-Gasco P, et al. Antiretroviral drugs and liver injury. *AIDS* 2008;22: 1–13.
15. Pathania MS, Kaur SLN, Kumar MS. A cross-sectional study of liver function tests in HIV-infected persons in Western India. *medical journal armed forces India* 73 (2017) 23–28.
16. Rathi PM, Amarapurkar DN, Borges NE et al. Spectrum of liver diseases in HIV infection. *Indian J Gastroenterol.* 1997 Jul;16(3):94-5
17. Ocama P, Katwere M, Piloya T et al. A. The spectrum of liver diseases in HIV infected individuals at an HIV treatment clinic in Kampala, Uganda. *Afr Health Sci.* 2008 Mar; 8(1): 8–12.
18. Luma HN, Eloumou SAFB, Ekaney DSM et al. Seroprevalence and Correlates of Hepatitis B and C Co-infection Among HIV-infected Individuals in Two Regional Hospitals in Cameroon. *Open AIDS J.* 2016; 10: 199–208.
19. Savita M, Singh RB, Vengadkrishnan K et al. Liver Function Abnormalities in Human Immunodeficiency Virus Positive Individuals and its Correlation with Disease Severity. *Int J Sci Stud* 2015;3(8):15-18.
20. Maggi P, Altizio S, Biagio AD et al. Prevalence and Risk Factors for Significant Liver Fibrosis in Patients with HIV Infection. *in vivo* 2015;29: 771-76.