



## Kidney Function Impairment in HIV Subjects On Art

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Date of Submission: 20-01-2024

Date of Acceptance: 30-01-2024

**ABSTRACT:** The number of people living with HIV/AIDS is on the rise, so are the problems like renal dysfunction which needs to be addressed. The study was aimed at providing data regarding prevalence and the association of certain modifiable and non-modifiable factors with renal function. This was a cross sectional study on 390 HIV positive subjects. Renal function was assessed by means of eGFR, calculated using Cockcroft-Gault equation, and proteinuria, estimated by spot urine albumin creatinine ratio (UACR). Renal dysfunction was identified by either eGFR < 60 ml/min or UACR > 30 mg/gm. The data was analyzed based on demographic characteristics, type of ART, height, weight, blood pressure and fasting lipid profile between renal dysfunction group and normal renal function group.

Different types of ART regimen, height, weight, systolic and diastolic blood pressure did not show any effect on renal function in this study. High triglyceride levels and high LDL levels are associated with decreased renal function.

**KEYWORDS:** HIV Seropositivity; Renal Insufficiency; Proteinuria; CD4 Cell Count ; Lipid profile;

### I. INTRODUCTION

Human Immunodeficiency Virus (HIV) was identified in 1983 as a causative agent for Acquired Immunodeficiency Syndrome (AIDS)<sup>1</sup>. This virus progressively damages cells of the body's immune system thus predisposing the infected individual to opportunistic infections, cancers, and different organ pathology.

In 1984 Rao<sup>2</sup> et al reported a renal lesion in patients infected with HIV that was characterized with severe proteinuria, biochemical features of nephrotic syndrome, and renal function impairment that rapidly progresses to end stage renal disease. Various types and severities of renal disorders have been encountered at all stages of HIV infections, ranging from mild and transient renal impairment to end stage renal disease requiring renal replacement therapy<sup>3</sup>. The CD4 cell count is used as a surrogate marker of immune status in HIV infected patients. However infection with HIV leads to destruction and depletion of CD4 cells and this predisposes the individual to various

disease conditions. CD4 cell count has been used to monitor, follow up and determine the severity of HIV infection<sup>4</sup>.

The renal function impairment (RFI) has been encountered at various stages of HIV infection, from the stage of seroconversion to advanced AIDS. The prevalence, severity, morbidity and mortality of renal function impairment in these patients have been associated with the degree of immunosuppression<sup>5,6</sup>. Various studies have reported that CD4 cell count of less than 200/ul is a poor prognostic index in HIV infected patients with renal disease<sup>7,8</sup>. Early detection and intervention delays progression and in many situations reverses impairment of renal function in HIV infected patients. The CD4 cell count of HIV infected patients is assessed on presentation in most health facilities. However renal function assessment is sparse in most of the centres in attending to HIV patients. It is necessary to objectively assess the relationship between renal function and degree of immunosuppression in these patients. Renal function is usually assessed by means of serum creatinine measurement and then estimated glomerular filtration rate calculation by Cockcroft-Gault formula or Modification of diet in renal disease equation (MDRD)<sup>9</sup>. Proteinuria and persistent microalbuminuria is also used for assessing renal function. Urine albumin creatinine ratio which is a surrogate marker for microalbuminuria can detect early renal dysfunction patients. Other determinants which contribute to renal dysfunction may include comorbidities such as hypertension, diabetes, hyperlipidemia, age, sex and ethnicity<sup>10-12</sup>. It might also be affected by many opportunistic infections, duration of HAART, type of drug regimen as shown in many previous studies<sup>13,14</sup>. Most of the studies regarding renal impairment in HIV patients have been done outside India and data regarding prevalence and pattern of renal involvement and its correlation with various variables in this part of India is not available. So the aim of this study is to provide valuable data regarding renal dysfunction in HIV infected individuals.

### OBJECTIVES



1. To determine the prevalence of renal impairment in HIV seropositive subjects attending ART clinic.
2. To determine the association between renal impairment and variables such as height, weight, systolic and diastolic blood pressure, type of ART regimen, and lipid profile.

## II. REVIEW OF LITERATURE HISTORICAL ASPECTS

The first indication of a new syndrome began in the summer of 1981 reported from New-York and California of a sudden increase in the incidence of two very rare diseases. Kaposi's sarcoma and pneumocystis carinii pneumonia's in young adults who were homosexuals or addicted to heroin or other IV narcotic<sup>15-17</sup>.

In May 1983, doctors from Dr. Luc Montagnier's team at the Pasteur Institute in France reported that they had isolated a new retrovirus from lymphoid ganglions that they believed was the cause of AIDS. The virus was later named lymphadenopathy-associated virus 1,<sup>18</sup>.

In May 1984 a team led by Robert Gallo of the United States confirmed the discovery of the virus, but they renamed it human T lymphotropic virus type III (HTLV-III).<sup>19,20</sup>

In January 1985, a number of more-detailed reports were published concerning LAV and HTLV-III, and by March it was clear that the viruses were the same, were from the same source, and were the etiological agent of AIDS<sup>21</sup>.

In India the first case of AIDS was reported in 1985 in Madras<sup>22</sup>.

In May 1986, the International Committee on Taxonomy of Viruses ruled that both names should be dropped and a new name, HIV (Human Immunodeficiency Virus), be used<sup>23</sup>.

The association between HIV and renal disease was first reported in 1984 by investigators in New York City and Miami, who reported a series of HIV-1- seropositive patients who developed a renal syndrome characterized by progressive renal failure and proteinuria<sup>2</sup>. The most common kidney biopsy finding was focal segmental glomerulosclerosis (FSGS)<sup>25,26</sup>. During the next several years, the existence of a specific HIV-associated nephropathy (HIVAN) was debated, in part, because of its similarity to heroin nephropathy and the frequent occurrence of intravenous drug use in this population<sup>27</sup>. Finding HIVAN in patients in whom a history of IVDU could be ruled out definitively helped to establish HIVAN as a distinct clinical entity<sup>28,29</sup>.

## III. METHODOLOGY

### Source of data

The material for the study will be collected from subjects, who are HIV seropositive and who fulfill the inclusion and exclusion criteria, attending ART clinic,

### Method of collection of data

- a) **Study design:** Single centered cross sectional study in a tertiary care hospital
- b) **Sample size:** 390 (As reviewed with statistician)
- c) **Sampling Method:** Simple Random Sampling
- d) **Duration of study:** January 2021 to August 2023
- e) **Method of collection of specimens and processing:** Blood and urine sample will be collected at a single sitting with universal precautions into separate sterile containers. Blood sample will be used for testing CD4 count, haemoglobin, total count using automated counters, and used to test serum creatinine by Jaffe method. Urine sample will be tested for urine albumin creatine ratio by immunoturbidometric method.

### f) Inclusion criteria

1. HIV seropositive subjects attending ART clinic.
2. Age more than or equal to 18 years.
3. Subjects either on Antiretroviral therapy or not on any drug therapy.

### g) Exclusion criteria

1. Subjects with preexisting urinary tract infection.
2. Pregnancy
3. Subjects not willing to take part in the study.
4. Patients on steroid intake or antituberculous treatment.

### h) Investigations

- 1) Haemoglobin
- 2) Urine albumin- creatinine ratio (Urine ACR).
- 3) Blood urea and serum creatinine
- 4) CD4 cell count
- 5) Lipid profile
- 6) RBS (Random Blood Sugar)
- 7) LFT (Liver Function Test)
- 8) White blood cell count

Data will be collected using a pretested proforma meeting the objectives of the study. Detailed history, physical examination and necessary investigation will be undertaken. The purpose of the study will be explained to the patient and informed consent obtained.

After collecting data subjects will be assessed for renal function impairment based on two criteria given below and classified into two groups, one with renal dysfunction and one with normal renal function. If a study subject satisfies any one of these two criteria, he/she will be



considered to have renal function impairment or renal dysfunction.

Criteria are 1.) Estimated GFR <60 ml/min as calculated using Cockcroft Gault formula and 2.) Microalbuminuria or Macroalbuminuria based on urine albumin creatine ratio (UACR) >30 mg/gm. The prevalence of renal dysfunction will be found out using this definition and the rest of the data will be used for further analysis and interpretation.

**Data Analysis and Interpretation**

Data was entered into Microsoft excel and analyses were done using the Statistical Package for Social Sciences (SPSS) for Windows software (version 18.0; SPSS Inc, Chicago). Descriptive statistics such as mean and standard deviation (SD)

for continuous variables, and frequency and percentage for categorical variables were determined. The level of significance was set at 0.05. Correlation analysis will be done wherever appropriate.

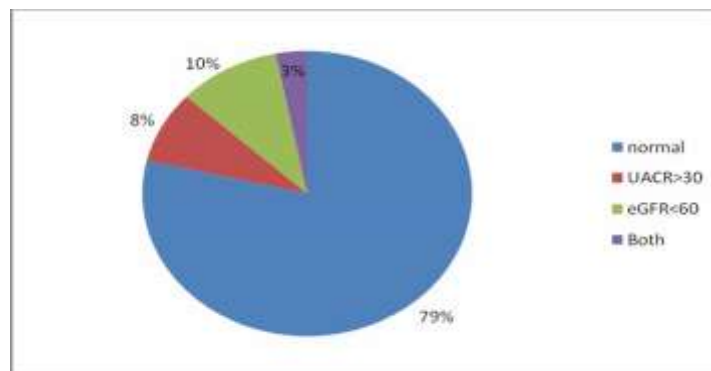
**IV. RESULTS**

In this study conducted in ART clinic, 390 subjects fulfilling inclusion and exclusion criteria were included. Out of them 83 subjects (21.28%) were found to have renal dysfunction according to the criteria used. Out of the 21.28% subjects with renal function impairment, 10 % had only low eGFR, 8.2% had only microalbuminuria and 3.08% had both decreased eGFR and microalbuminuria.

**Table 1: Renal function distribution**

Normalrenalfunction		307
Renaldysfunction	eGFR<60	39
	UACR>30	32
	Both	12

**Figure 1: Renal function distribution**



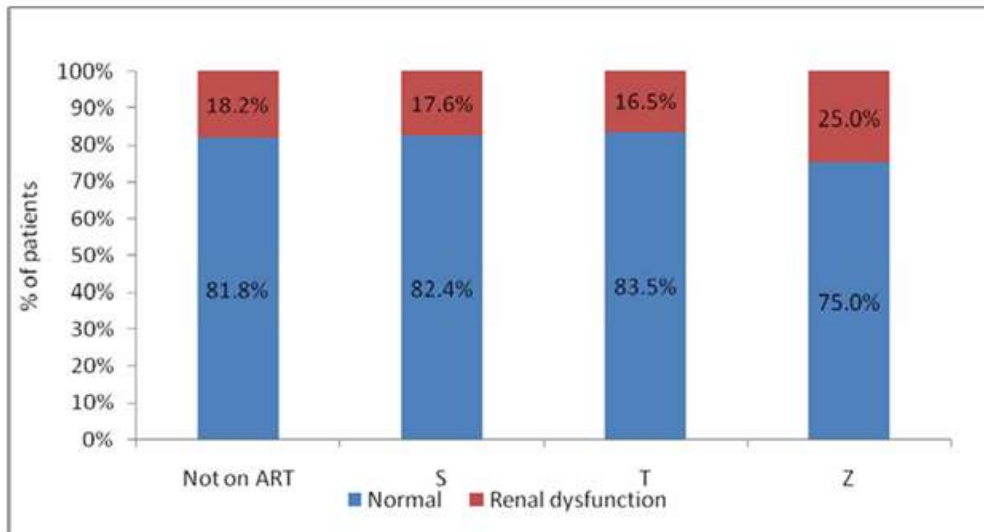
**Table 2: ART Regimen distribution**

ART Regimen	Not on ART	Stavudine(S)	Tenofovir(T)	Zidovudine(Z)
	%ofpatients (No.)	%ofpatients (No.)	%ofpatients (No.)	%ofpatients (No.)
Normal function	81.82%(72)	82.35%(14)	83.53%(71)	75.00%(150)
Renal dysfunction	18.18%(16)	17.65%(3)	16.47%(14)	25.00%(50)

P-value between Not on ART and S	P-value between Not on ART and T	P-value between Not on ART and Z	P-value between S and T	P-value between S and Z	P-value between T and Z
0.96012	0.7253	0.2048	0.90448	0.4983	0.1144



**Figure 2: ART Regimen distribution**



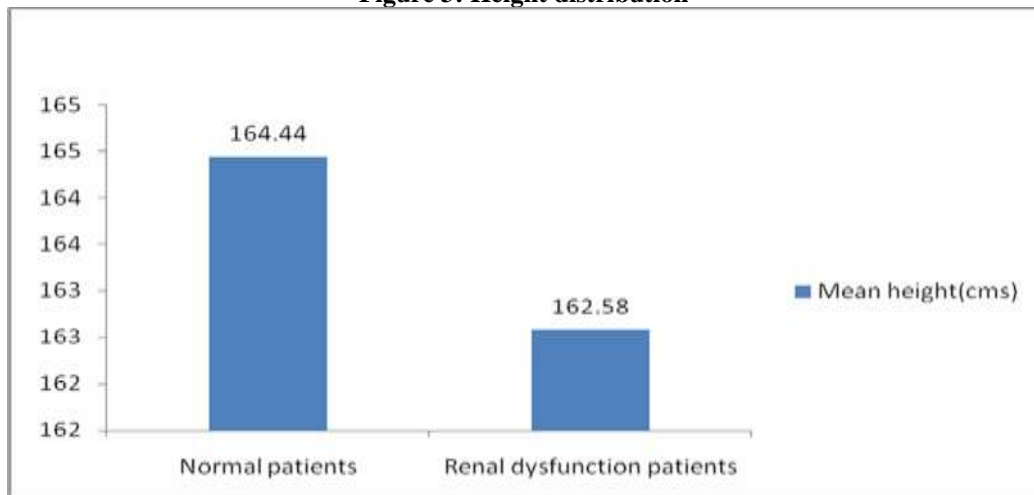
The proportions of patients with renal dysfunction across the entire regimen whether subjects were not on ART or on ART were comparison across all the regimens, the proportions do not differ and are statistically same.

statistically same with p-value greater than 0.05 at 5% significance level. Since the p-value is greater than 0.05 for any two proportion

**Table 3: Height distribution**

Height distribution	Number of patients	Mean height (cms)	Minimum height (cms)	Maximum height (cms)	Standard deviation	P-value
Normal patients	307	164.44	150	182	6.75	0.032
Renal dysfunction patients	83	162.58	152	180	7.02	

**Figure 3: Height distribution**



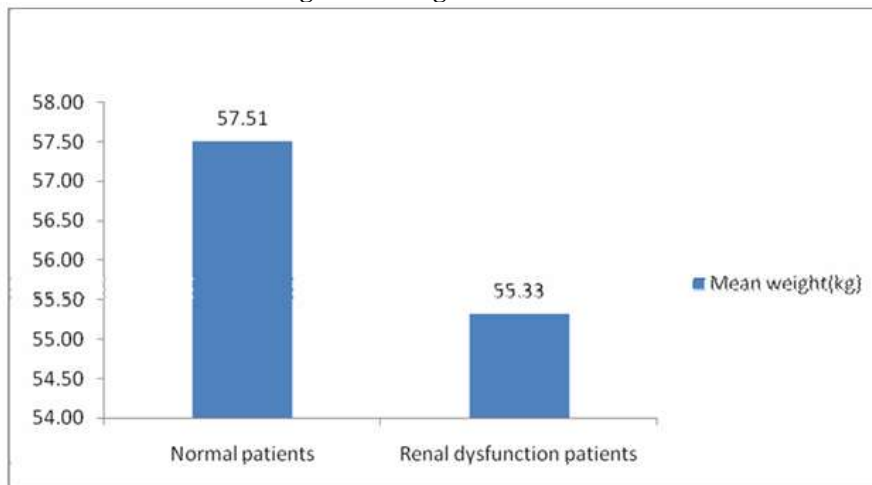


Mean height among normal patients and renal dysfunction patients is statistically different with p-value 0.032 which is significant at 5% significance level.

**Table 4: Weight distribution**

Weightdistribution	Mean weight(kg)	Minimum weight(kg)	Maximum weight(kg)	Standard deviation	P-value
Normalpatients	57.51	40	71	6.02	0.004
Renaldysfunctionpatients	55.33	43	68	6.04	

**Figure 4: Weight distribution**

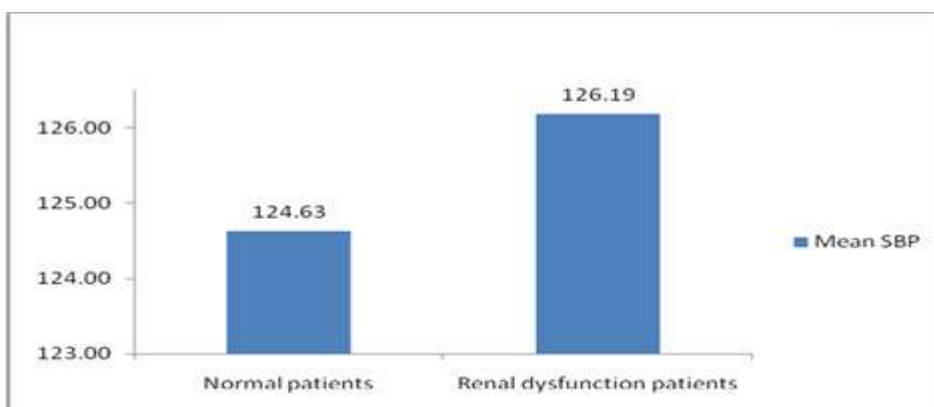


The mean weight is statically different among normal patients and renal dysfunction patients.

**Table 5: Systolic BP distribution**

SBPdistribution	Mean SBP	Minimum SBP	Maximum SBP	Standard deviation	P-value
Normalpatients	124.63	108	154	8.62	0.232
Renaldysfunctionpatients	126.19	12	150	15.71	

**Figure 5: Systolic BP distribution**





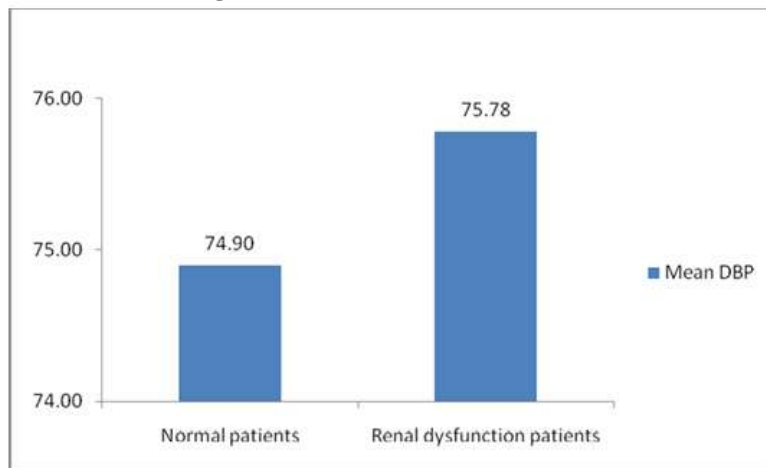
Mean SBP is statistically same among normal patients and also patients with renal dysfunction with p-value 0.232. The p-value is

statistically insignificant at 5% significance level. It indicates that the mean SBP is statistically same across both the groups.

**Table 6: Diastolic BP distribution**

DBPdistribution	Mean DBP	Minimum DBP	Maximum DBP	Standard deviation	P-value
Normalpatients	74.90	62	90	5.57	0.204
Renaldysfunctionpatients	75.78	68	90	5.75	

**Figure 6: Diastolic BP distribution**



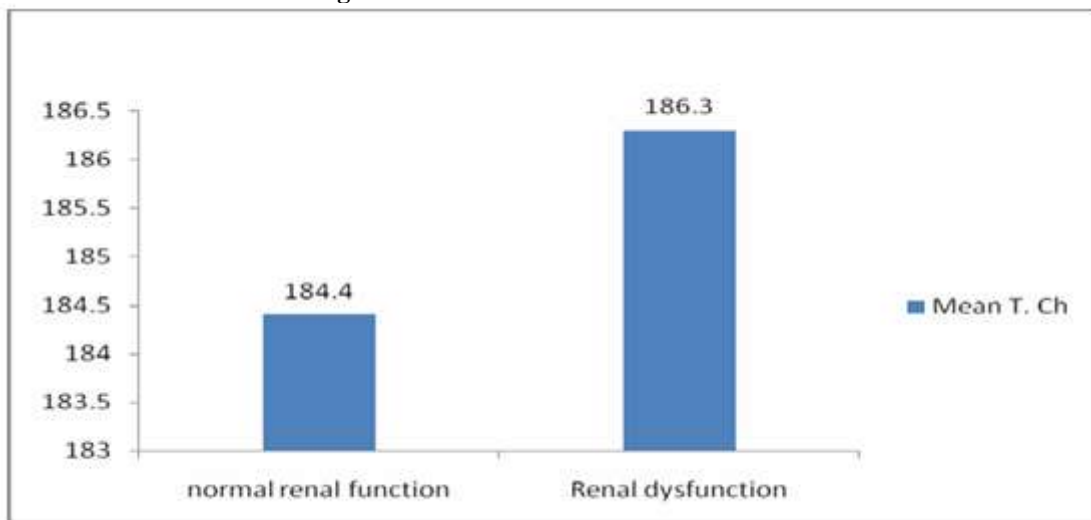
Mean DBP is statistically same among normal patients and also patients with renal dysfunction with p-value 0.204. The p-value is

statistically insignificant at 5% significance level. It indicates that the mean DBP is statistically same across both the groups.

**Table 7: Serum Cholesterol distribution**

SerumCholesterol	Mean T.Ch	Minimum T.Ch	Maximum T.Ch	Standard deviation	P-value
Normalpatients	184.39	106	355	33.05	0.667
Renaldysfunctionpatients	186.27	96	341	42.62	

**Figure 7: Serum Cholesterol distribution**



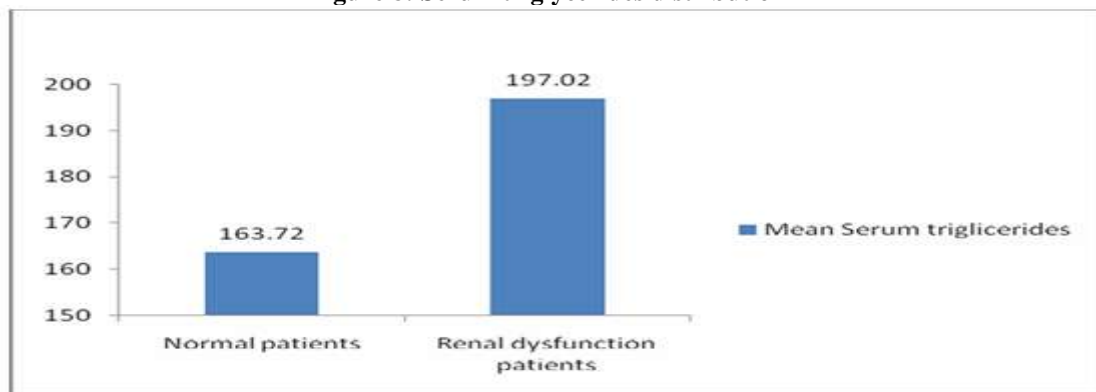


P-value 0.667 suggests that mean Serum total cholesterol is statistically same at 5% significance level.

**Table 8: Serum triglycerides distribution**

Serum triglycerides	Mean Serum triglycerides	Minimum Serum triglyceride	Maximum Serum triglyceride	Standard deviation	P-value
Normal patients	163.72	13	488	64.67	0.0
Renal dysfunction patients	197.02	84	580	88.84	

**Figure 8: Serum triglycerides distribution**



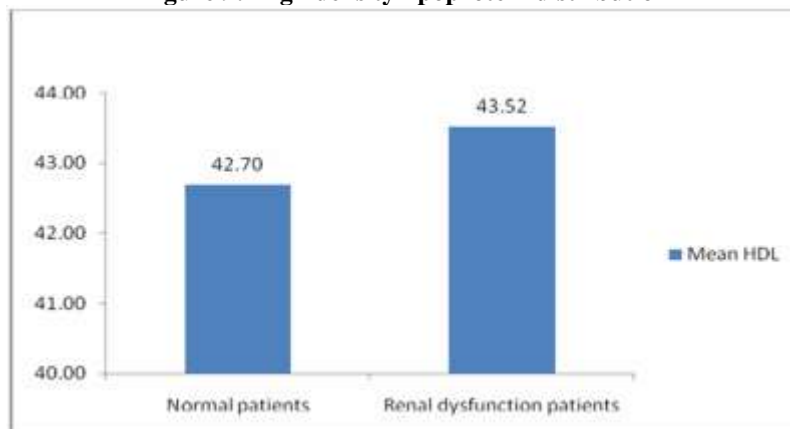
Mean Serum triglycerides is statistically different at 5% significance level with p-value 0.0. It indicates the mean of Serum triglycerides is

different and it is significantly higher among the patients with renal dysfunction.

**Table 9: High density lipoprotein distribution**

HDL distribution	Number of patients	Mean HDL	Minimum HDL	Maximum HDL	Standard deviation	P-value
Normal patients	307	42.70	3	70	7.44	0.396
Renal dysfunction patients	83	43.52	27	88	9.07	

**Figure 9: High density lipoprotein distribution**







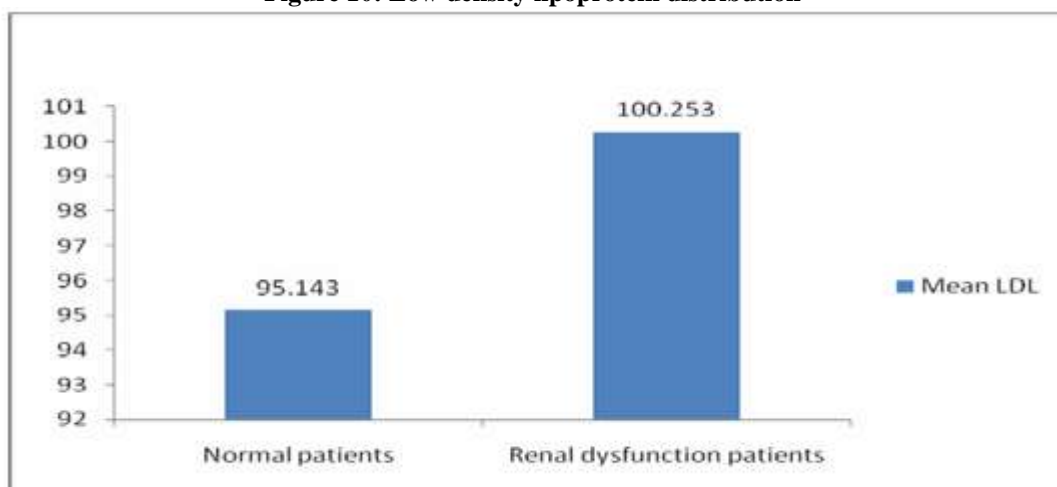
Mean HDL is statistically same among normal patients and also patients with renal dysfunction with p-value 0.396. The p-value is

statistically insignificant at 5% significance level. It indicates that the mean HDL is statistically same across both the groups.

**Table 10: Low density lipoprotein distribution**

LDLdistribution	Mean LDL	Min. LDL	Max. LDL	Standard deviation	P-value
Normalpatients	95.143	25	153	14.77	0.005
Renaldysfunctionpatients	100.253	66	151	14.20	

**Figure 10: Low density lipoprotein distribution**



Mean LDL is statistically different among both the groups with p-value 0.05 at 5% significance level. It indicates that mean LDL is statistically higher among the patients with renal dysfunction.

## V. DISCUSSION

In this study conducted in ART clinic, 390 subjects fulfilling inclusion and exclusion criteria were included. Out of them 83 subjects (21.28%) were found to have renal dysfunction according to the criteria used. Renal function impairment was identified based on two criteria, first one was estimated GFR<60 ml/min calculated using Cockcroft Gault formula and second one was microalbuminuria based on urine albumin-creatinine ratio>30 mg/gm. Patient was considered to have renal dysfunction if any one of the criteria was fulfilled.

### ART REGIMEN

In this study subjects were divided into 4 groups based on their ART regimen namely Stavudine regimen, Tenofovir regimen, Zidovudine regimen and subjects who were not initiated on

ART. The proportions of patients with renal dysfunction across the different regimens were statistically same with p-value greater than 0.05 at 5% significance level. So no statistically significant difference was seen between various regimens and renal dysfunction.

This was consistent with studies conducted by Keith Rawlings<sup>81</sup> et al (2009) in USA and H.M.F. Kamga<sup>82</sup> et al (2011) in Cameroon which showed no difference with different ART regimens. But study conducted by Menezes<sup>10</sup> et al (2009) showed tenofovir use as a risk factor for renal impairment.

### HEIGHT

In this study mean height was 164.4 (S.D =6.75) cms in normal renal function subjects and 162.6 (S.D =7.02) cms in renal dysfunction subjects. P-value was 0.032 which is significant at 5% significance level. So less height was statistically associated with renal dysfunction i.e. decrease in eGFR.





## WEIGHT

In this study mean weight was 57.5 (S.D =6.02) kgs in normal renal function subjects and 55.3 (S.D =6.04) kgs in renal dysfunction subjects. P-value was 0.004 which is significant at 5% significance level. Decline in renal function was significantly associated with less weight. This result is consistent with the study by Menezes<sup>10</sup> et al in Brazil where higher weight was significantly associated with normal kidney function (RR= 0.89, p = 0.005). In the study by Struik<sup>83</sup> et al also lower weight was associated with lower eGFR.

## BLOOD PRESSURE

In the present study mean systolic blood pressure (SBP) among normal renal function subjects and renal dysfunction subjects were 124.6(S.D=8.6) and 126.2(S.D=15.7) mms of Hg respectively. It was found to be statistically insignificant, p-value=0.232. Mean diastolic blood pressure (DBP) was 74.9(S.D=5.6) and 75.8(S.D=5.8) among normal function group and renal dysfunction group respectively. It was also found to be insignificant, p-value being 0.204. These findings were consistent with studies by Struik<sup>83</sup> et al in Malawi, Okafor<sup>5</sup> et al in Nigeria and Han<sup>36</sup> et al in South Africa which showed no statistically significant difference in blood pressure between the two groups.

## LIPID PROFILE

In this study mean total cholesterol was 184.9 (SD=33.05) mg/dl and 186.2 (SD=42.62) mg/dl among normal and renal dysfunction group respectively. The difference was statistically insignificant, p-value = 0.667. Mean triglyceride values were 163.7 (S.D =64.7) mg/dl and 197.0 (S.D =88.8) mg/dl respectively and p value was 0.00. So higher triglyceride value was associated with renal dysfunction. The difference in mean HDL value of 42.7 (S.D =7.4) mg/dl among normal renal function group and 43.5 (S.D =9.0) among renal dysfunction group was found to be statistically insignificant, p-value = 0.396. LDL means were 95.1(S.D =14.8) mg/dl and 100.3 (S.D =14.2) mg/dl among normal and renal dysfunction group and was found to be statistically significant, p-value = 0.005.

In the study by Overton<sup>11</sup> et al conducted among 845 subjects in USA it was found that higher total cholesterol and higher triglyceride level was associated with decline in renal function whereas HDL and LDL level did not show any significant association.

## VI. CONCLUSIONS

1. A total of 390 HIV infected subjects were included in the study.
2. The prevalence of renal dysfunction in the study was 21.28% (N= 83). So around one fifth of HIV infected patients will have renal function impairment.
3. Around half of the renal dysfunction subjects (50.16%) were in the age group 38-47 years.
4. Though the study population was predominantly males, 60.24% of the renal dysfunction group was females.
5. HIV positive patients whether pre-ART or on different ART regimens did not have any implication on renal function.
6. There is increased chance of renal function impairment with increased duration of anti-retroviral therapy.
7. Lesser height and weight is associated with more chance of renal dysfunction.
8. Body mass index did not show any significant effect on renal function.
9. Systolic and diastolic blood pressure did not show any association with renal dysfunction in this study.
10. Lower CD4 count which indicates more immunosuppression is associated with decreased renal function and more so with decline in estimated GFR rather than proteinuria.
11. Abnormal lipid profile in the form of high triglycerides and high LDL contributes to renal function impairment.

## SUMMARY

HIV infected population and AIDS is a global health concern and more so in a populated country like India. With increasing awareness and advent of ART, the number of people living with HIV/AIDS is on the rise, so are the problems like renal dysfunction which needs to be addressed. The study was aimed at providing data regarding prevalence and the association of certain modifiable and non-modifiable factors with renal function.

The study was conducted in ART clinic. A total of 390 subjects were studied and prevalence of renal dysfunction was found to be 21.28%. Around 3/5th of renal dysfunction population were females. Lower height, lower weight, increased duration of ART therapy were associated with decreased renal function. Different types of ART regimen, body mass index, systolic and diastolic blood pressure did not show any effect on renal function in this study. Decrease in CD4 count was shown to have positive correlation with decrease in estimated GFR. High triglyceride levels and high



LDL levels are associated with decreased renal function.

#### LIMITATIONS OF THE STUDY

1. No attempt was made to differentiate between causes of renal dysfunction in terms of acute/chronic or reversible /irreversible in the group studied.

2. This study did a onetime assessment of renal function which would have overestimated or underestimated the prevalence of renal dysfunction. Multiple assessments would have been better.

#### REFERENCE

- [1]. Barré-Sinoussi F, Chermann JC, Rey F, Nugeyre MT, Chamaret S, Gruest J et al. Isolation of a T-lymphotropic retrovirus from a patient at risk for acquired immune deficiency syndrome (AIDS). *Science* 1983;220(4599):868–71.
- [2]. Rao TKS, Fillipone EJ, Nicastrì AD. Associated focal and segmental glomerulosclerosis in Acquired Immunodeficiency Syndrome. *N Engl J Med* 1984;1631–45.
- [3]. Roling J, Schmid H, Fischereeder M, Draenert R, Goebel FD. HIV-associated renal diseases and highly active antiretroviral therapy induced nephropathy. *Clin Infect Dis* 2007;42(10):1488–95.
- [4]. Abdul K Abbar. Diseases of immunity: AIDS. In: Robbins and Cotron Pathologic Basis of diseases. 7th ed. 2004. p.245–65.
- [5]. Okafor UH, Unuigbo EI, Ojogwu LI, Oviasu E, Wokoma FS, Renal disease in HIV infected patients at University of Benin Teaching Hospital in Nigeria. *African Health Sciences* 2011;11(S1):S28 - S33.
- [6]. Jonathan A Winston, Mary E Klotman, Paul E Klotman. HIV associated nephropathy is a late, not early, manifestation of HIV-1 infection. *Kidney International* 1999;55:1036–40.
- [7]. Andrew Reid, Wolfgang Stohr, A Sarah Walker, Ian G Williams, Cissy Kityo, Peter Hughes et al. Severe renal dysfunction and risk factors associated with renal impairment in HIV infected adults in Africa initiating antiretroviral therapy. *Clinical Infectious Diseases* 2008;46:1271–81.
- [8]. Winston JA, Klotman PE. The human immunodeficiency virus (HIV) and HIV associated nephropathy. *Semin Nephrol* 1998;18:373–7.
- [9]. Joanne MB, Karl Skorecki. Chronic kidney disease: Harrison's principles of internal medicine. 18th ed. 2012;2308–21.
- [10]. Menezes AM, Torelly J Jr., Real L, Bay M, Poeta J, Sprinz E. Prevalence and Risk Factors Associated to Chronic Kidney Disease in HIV-Infected Patients on HAART and Undetectable Viral Load in Brazil. *PloS one* 2011;6:e26042.
- [11]. Overton ET, Nurutdinova D, Freeman J, Seyfried W, Mondy KE. Factors associated with renal dysfunction within an urban HIV infected cohort in the era of HAART. *HIV Medicine* 2009;10:343–50.
- [12]. Cantor ES, Kimmel PL, Bosch JP. Effect of race on expression of acquired immunodeficiency syndrome-associated nephropathy. *Arch Intern Med* 1991; 151:125–8.
- [13]. Philippe Flandre, Pascal P, Lise C, Corinne IB, Ivan T, Andre C, et al. Risk Factors of Chronic Kidney Disease in HIV-infected Patients. *Clin J Am Soc Nephrol* 2011 Jul 6;1700–7.
- [14]. Schwartz EJ, Szczech LA, Ross MJ, Klotman ME, Winston JA, Klotman PE. Highly Active Antiretroviral Therapy and the Epidemic of HIV+ End-Stage Renal Disease. *J Am Soc Nephrol* 2005;16(8):2412–20.
- [15]. Pneumocystis pneumonia. Los Angeles. *MMWR Morb Mortal Wkly Rep* 1981;30(21):250–2.
- [16]. Update on Kaposi's sarcoma and opportunistic infections in previously healthy persons—United States. *MMWR Morb Mortal Wkly Rep* 1982 Jun; 31(22):294,300–1.
- [17]. A cluster of Kaposi's sarcoma and Pneumocystis carini pneumonia among homosexual male residents of Los Angeles and Orange Counties, California. *MMWR Morb Mortal Wkly Rep* 1982 Jun;31(23):305–7.
- [18]. Kingman, Sharon; Connor, Steve. The search for the virus. Harmondsworth, England: Penguin; 1989.
- [19]. Popovic M, Sarngadharan MG, Read E, Gallo RC. Detection, isolation, and continuous production of cytopathic retroviruses (HTLV-III) from patients with AIDS and pre-AIDS. *Science* 1984;224(4648):497–500.
- [20]. Marx JL. A virus by any other name. *Science* 1985 Mar;227(4693):1449–51.



- [21]. Chang SY, Bowman BH, Weiss JB, Garcia RE, White TJ. The origin of HIV-1 isolate HTLV-IIIB. *Nature* 1993 Jun;363(6428):466–9.
- [22]. Sajithkumar R. Basic considerations and Epidemiology of HIV infection. *API textbook of medicine*. 7th ed. 2003. p.148-9.
- [23]. Coffin J, Haase A, Levy JA, Montagnier L, Oroszlan S, Teich N, et al. What to call the AIDS virus? *Nature* 1986;321(6065):10.
- [24]. Peter Piot. HIV infection and AIDS. A global overview. Chapter 407. *Goldman's Cecil Medicine* 2007.
- [25]. Agati V, Suh JI, Carbone L, Cheng JT, Appel G. Pathology of HIV-associated nephropathy: A detailed morphologic and comparative study. *Kidney Int* 1989; 35:1358–70.
- [26]. Agati V, Appel GB. Renal pathology of human immunodeficiency virus infection. *SeminNephrol* 1998;18:406–21.
- [27]. Mazbar SA, Schoenfeld PY, Humphreys MH. Renal involvement in patients infected with HIV: Experience at San Francisco General Hospital. *Kidney Int* 1990;37:1325–32.
- [28]. Strauss J, Abitbol C, Zilleruelo G, Scott G, Paredes A, Malaga S, et al. Renal disease in children with the acquired immunodeficiency syndrome. *N Engl J Med* 1989;321:625–30.
- [29]. Pardo V, Meneses R, Ossa L, Jaffe DJ, Strauss J, Roth D, et al. AIDS-related glomerulopathy: Occurrence in specific risk groups. *Kidney Int* 1987;31:1167–37.
- [30]. Agati V, Appel GB. HIV infection and the kidney. *J Am SocNephrol* 1997; 8:138–52.
- [31]. Monahan M, Tanji N, Klotman PE. HIV-associated nephropathy: An urban epidemic. *SeminNephrol* 2001;21:394–402.
- [32]. US Renal Data System (USRDS). *USRDS 2001 Annual Data Report*. Bethesda MD, The National Institutes of Health. National Institute of Diabetes and Digestive and Kidney Diseases 2001.
- [33]. Winston JA, Klotman PE. Are we missing an epidemic of HIV associated nephropathy? *J Am SocNephrol* 1996;7:1–7.
- [34]. Centers for Disease Control and Prevention (CDC). *HIV/AIDS Surveillance Report*. Atlanta; 2001. p.5–35.
- [35]. SreepadaRao TK, Eli A Friedman, Anthony D Nicastrì. The Types of Renal Disease in the Acquired Immunodeficiency Syndrome. *New England Journal Med* 1987;316:1062-8.
- [36]. Han TM, Naicker S, Ramial PK, Assounga AG. A cross sectional study of HIV seropositive patients with varying degrees of proteinuria in South Africa. *Kidney International* 2006;69:2243-50.
- [37]. Anthony S Fauci, Clifford H Lane. Human Immunodeficiency Virus disease: AIDS and related disorders. In: *Harrison's Principles of internal medicine*. 18th ed. 2012. p.1506-87.
- [38]. Joint United Nations Programme on HIV/AIDS. *UNAIDS report on the global AIDS epidemic 2012*;2012.
- [39]. AIDS. *Park's Textbook of Preventive and Social medicine* 18th ed. p.271.
- [40]. Jawetz, Melnick, Adelberg. *Medical Microbiology, AIDS and Lentiviruses*. Chapter 44. 2007;24e.
- [41]. Riesenber MD. HIV virus and Pathogenesis of AIDS. *JAMA* 1989 May 26; 261(20).
- [42]. Sanjay Pujari. Pathophysiology of HIV infection. *API Textbook of medicine* 7th ed. 2003. p.149-50.
- [43]. Sivaraman V, Gibert Fernandez, SambasivaRao R. HIV infection and pulmonary tuberculosis report of 6 cases. *Ind J Tub* 1992,35,39.
- [44]. Shantanu Doe. Prevalence of HIV infection in patients with tuberculosis. *Ind J Tub* 1995;42:183.
- [45]. Narain JP. HIV associated tuberculosis in developing countries, epidemiology and strategies for prevention. *Tuberc and Lung Dis* 1992;73:311-21.
- [46]. Nisar M, Narula M. HIV related tuberculosis in England and Wales. *Tuberc and Lung Dis* 1992;73:200-2.
- [47]. Atul K Patel. Laboratory diagnosis of HIV infection/AIDS. *API Textbook of Medicine*. 7th ed. 2003. p.155-6.
- [48]. Jehangir S Sorabjee. Clinical Approach to a patient with HIV infection. *API Textbook of medicine*. 7th ed. 2003. p.152-4.
- [49]. Yeolekar ME. Opportunistic infections in AIDS. *API Textbook of medicine*. 7th ed. 2003. p.156-8.
- [50]. Human Immunodeficiency Virus. In: *Ananthanarayanan's Textbook of Microbiology*. 6th ed. 2000. p.539-54.
- [51]. Sushrut SW, Joseph VB. Acute Kidney injury: Chapter 279. *Harrison's principles of Internal Medicine*. 2012. p.2293-308.
- [52]. SreepadaRao TK. Human immunodeficiency virus infection and renal failure. *Infect Dis Clin North Am* 2001;15(3):833-50.



- [53]. Kimmel P. The nephropathies of HIV infection: pathogenesis and treatment. *Curr Opin Nephrol Hypertens* 2000;9(2):117-22.
- [54]. Weiner NJ, Goodman JW, Kimmel PL. The HIV-associated renal diseases: current insight into pathogenesis and treatment. *Kidney Int* 2003;63(5):1618-31.
- [55]. Kimmel P, Maslo C, Akposso K, Mougenot B, Rondeau E. Acute renal failure in the course of HIV infection: a single-institution retrospective study of ninety-two patients and sixty renal biopsies. *Nephrol Dial Transplant* 1999;14:1578-85.
- [56]. Franceschini N, Napravnik S, Finn WF, Szczech LA, Eron JJ Jr. Immunosuppression, hepatitis C infection, and acute renal failure in HIV-infected patients. *J Acquir Immune Defic Syndr* 2006;42(3):368-72.
- [57]. Bourgoignie JJ. Renal complications of human immunodeficiency virus type 1. *Kidney Int* 1990;37(6):1571-84.
- [58]. Pardo V, Aldana M, Colton RM. Glomerular lesions in the acquired immunodeficiency syndrome. *Ann Intern Med* 1984;101(4):429-34.
- [59]. Pardo V, Meneses R, Ossa L. AIDS-related glomerulopathy: occurrence in specific risk groups. *Kidney Int* 1987;31(5):1167-73.
- [60]. Hailemariam S, Walder M, Burger HR. Renal pathology and premortem clinical presentation of Caucasian patients with AIDS: An autopsy study from the era prior to antiretroviral therapy. *Swiss Med Weekly* 2001;131:412-7.
- [61]. Shahinian V, Rajaraman S, Borucki M, Grady J, Hollander WM, Ahuja TS. Prevalence of HIV-associated nephropathy in autopsies of HIV-infected patients. *Am J Kidney Dis* 2000;35(5):884-8.
- [62]. diBelgiojoso GB, Ferrario F, Landriani N. Virus-related glomerular diseases: histological and clinical aspects. *J Nephrol* 2002;15(5):469-79.
- [63]. Gerntholtz TE, Goetsch SJ, Katz I. HIV-related nephropathy: a South African perspective. *Kidney Int* 2006;69(10):1885-91.
- [64]. Wools-Kaloustian K, Gupta SK, Muloma E. Renal disease in an antiretroviral naive HIV-infected outpatient population in Western Kenya. *Nephrol Dial Transplant* 2007;22(8):2208-12.
- [65]. Andia I, Pepper LM, Matthieson P. Prevalence of renal disease in outpatients with HIV/AIDS in Mbarara Hospital. 3rd IAS (International Aids Society) Conference on HIV Pathogenesis and Treatment, Rio de Janeiro 2005 Jul 24-27.
- [66]. Szczech LA, Grunfeld C, Scherzer R. Microalbuminuria in HIV infection. *AIDS* 2007;21(8):1003-9.
- [67]. Wali RK, Drachenberg CI, Papadimitriou JC, Keay S, Ramos E. HIV-1-associated nephropathy and response to highly-active antiretroviral therapy. *Lancet* 1998;352(9130):783-4.
- [68]. Betjes MG, Verhagen DW. Stable improvement of renal function after initiation of highly active anti-retroviral therapy in patients with HIV-1-associated nephropathy. *Nephrol Dial Transplant* 2002;17(10):1836-9.
- [69]. Levin ML, Palella F, Shah S, Lerma E, Butter J, Kanwar YS. HIV-associated nephropathy occurring before HIV antibody seroconversion. *Am J Kidney Dis* 2001;37(5):E39.
- [70]. Gardner LI, Holmberg SD, Williamson JM. Development of proteinuria or elevated serum creatinine and mortality in HIV-infected women. *J Acquir Immune Defic Syndr* 2003;32(2):203-9.
- [71]. WHO. Case Definitions of HIV for Surveillance and Revised Clinical Staging and Immunological Classification of HIV-related Disease in Adults and Children. Geneva: World Health Organization; 2007.
- [72]. Kopp JB, Winkler C. HIV-associated nephropathy in African Americans. *Kidney Int* 2003;83:S43-9.
- [73]. Freedman BI, Soucie JM, Stone SM, Pegram S. Familial clustering of end-stage renal disease in blacks with HIV-associated nephropathy. *Am J Kidney Dis* 1999;34(2):254-8.
- [74]. Cohen AH. HIV-associated nephropathy: Racial difference in severity of renal damage. *J Am Soc Nephrol* 1990;1:305.
- [75]. Ross MJ, Klotman PE. Recent progress in HIV-associated nephropathy. *J Am Soc Nephrol* 2002;13(12):2997-3004.
- [76]. Smith MC, Austen JL, Carey JT. Prednisone improves renal function and proteinuria in human immunodeficiency virus-associated nephropathy. *Am J Med* 1996;101(1):41-8.
- [77]. Eustace JA, Nuermberger E, Choi M, Scheel PJ Jr, Moore R, Briggs WA. Cohort study of the treatment of severe HIV-associated nephropathy with corticosteroids. *Kidney Int* 2000;58(3):1253-60.



- [78]. Ifudu O, SreepadaRao TK, Tan CC, Fleischman H, Chirgwin K, Freidman EA. Zidovudine is beneficial in human immunodeficiency virus associated nephropathy. *Am J Nephrol* 1995;15:217-21.
- [79]. Michel C, Dosquet P, Ronco P, Mougnot B, Viron B, Mignon F. Nephropathy associated with infection by human immunodeficiency virus: a report on 11 cases including 6 treated with zidovudine. *Nephron* 1992;62(4):434-40.
- [80]. Ahuja TS, Borucki M, Funtanilla M, Shahinian V, Hollander M, Rajaraman S. Is the prevalence of HIV-associated nephropathy decreasing? *Am J Nephrol* 1999; 19(6):655-9.
- [81]. Rawlings MK, Jennifer K, Edna PT, Jeanée Queen E, Lauren R, Linda HY, et al. Impact of comorbidities and drug therapy on development of renal impairment in a predominantly African American and Hispanic HIV clinic population. *HIV/AIDS – Research and Palliative Care* 2011;3 1–8.
- [82]. Kanga HLF, Assob JCN, Njunda AL, Fon PN, Nsagha DS, Atanga MBS, et al. The kidney function trends in human immunodeficiency virus/AIDS patients at the Nylon District Hospital, Douala. *Cameroon Journal of AIDS and HIV Research* 2011;3(2):30-7.
- [83]. Struik GM, den Exter RA, Munthali C, Chipeta D, van Oosterhout JJG, Nouwen JL, Allain TJ. The prevalence of renal impairment among adults with early HIV disease in Blantyre, Malawi. *International Journal of STD and AIDS* 2011; 22:457-62.