

1 **SEROPREVALENCE OF HCV AND HIV ANTIBODIES IN TUBERCULOSIS**
2 **CONFIRMED PATIENT IN EKITI STATE, NIGERIA**

3
4 **ABSTRACT**

5 This study was conducted to determine the seroprevalence of HCV and HIV antibodies
6 in TB confirmed patient attending the Federal Medical Centre (FMC), Ido Ekiti, Ekiti
7 State, Nigeria. A total of 500 tuberculosis confirmed patients were selected by random
8 sampling. Their blood samples were collected and assayed for HCV and HIV antibodies
9 using Clinotech diagnostic Anti-HCV detection test and Abbot determine HIV ½ in
10 conjunction with Chembio HIV ½ STAT-PAK assay kit respectively. Out of 500 TB
11 patients tested, 10(2.0%), 21(4.2%) and 3(0.6%) tested positive to HCV, HIV, and
12 HCV/HIV antibodies respectively. Age group 36-45 was the most prevalence of HCV,
13 HIV, and HCV/HIV antibodies with P-value 0.000, 0.000 and 0.002 respectively. The
14 associated risk factors were alcoholism 14 (45.2%), being the highest identified risk
15 factor, followed by previous unprotected sex, multiple sex partner, previous blood donor,
16 previous transfusion, tattoos, and history of the Sexually transmitted disease being the
17 least risk factor 3 (9.68%). The degree of disparity in regards to HCV, HIV and co-exists
18 of HCV/HIV antibodies between 302 male and 198 female that participated were not
19 statistically significant. (P-value 0.531, 0.549, and 0.824 for HCV, HIV and HCV/HIV
20 antibodies respectively). These findings confirmed that both HCV and HIV can co-exist in
21 TB patients, and may increase the risk of antituberculosis drug-induced hepatotoxicity, if
22 overlooked, there will be a greater risk for TB patients, and these infections will continue
23 to spread through the associated risk factors. However, in managing the TB patients,
24 there is a need to screen for Anti- HCV, as it has been for HIV antibody.

25 **Keywords:** Nigeria, HCV and HIV antibodies, tuberculosis, Seroprevalence

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27 **INTRODUCTION**

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29 **Hepatitis C virus (HCV)** belongs to the family of Flaviviridae, genus- hepacivirus and it
30 is 50nm in size with enveloped, single-stranded, positive-sense RNA [1]. It infects an
31 estimated 170 million person's worldwide. The institution of blood screening measures in

32 industrialized countries has reduced the risk of transfusion-associated hepatitis to a
33 minimal level, but the transfusion-related transmission still occurs in developing
34 countries that have not fully implemented blood screening procedures. Globally, new
35 cases of HCV also continue to occur as a result of injecting drug use (IDU) and through
36 other means of percutaneous or mucous membrane exposure [2]

37 HIV infection in humans is now pandemic as of January 2006, the Joint United
38 Nations Programme on HIV/AIDS (UNAIDS) and the world health organization (WHO)
39 estimates that AIDS has killed more than 25 million people since it was first recognized
40 in 1981, making it one of the most destructive pandemics in recorded history (. It is
41 estimated that about 0.6% of the world's population is infected with HIV [3]. HIV
42 prevalence varies widely by geographic region and vulnerable population, Nigeria has an
43 overall national prevalence of 3.1% but statewide; HIV prevalence among pregnant
44 women has ranged from as low as 1.6% in Ekiti in the west to 10 % in Benue in south-
45 east [4].

46 However, the current estimate state that HIV is set to infect 90 million people in
47 Africa, resulting in a minimum estimate of 18 million orphans. Antiretroviral drug reduce
48 mortality and morbidity of HIV infection, but routine access to antiretroviral medications
49 are not available in all countries [5]. HCV co-infection with HIV is common and rates
50 among HIV positive population are higher [6]. About 10,000-20,000 death yearly in the
51 US is from HCV; expectations are that this mortality rate will increase, as those who
52 were infected by transfusion before HCV testing become apparent. It is responsible for
53 90-95% of all transfusion-related hepatitis [7].

54 Tuberculosis (TB) has been major public health problems for centuries. The
55 implementation of effective public health interventions for the prevention and control of

56 TB has significantly contributed to a substantial reduction of the global disease but the
57 emergence of the HIV epidemic has posed major challenges to TB control effort globally.
58 In a country with almost 40 % population already infected with TB, increasing
59 prevalence of HIV will be jeopardized TB central effort with such consequences [8]. HIV
60 has been thought to account for much of the recent increase in the global TB burden,
61 especially in Africa. [9].

62 HIV is the most important risk factor for the development of TB among person
63 infected with *M. tuberculosis* and both Centres for Disease Control and Prevention
64 (CDC) and WHO guidelines recommend offering HIV testing to those person diagnosed
65 with TB disease [2].

66 The prevalence of HCV infection among persons with TB has been poorly
67 defined, and few data are available from most areas around the world. One recent study
68 from the US has suggested that veterans with HCV infection are at risk for other selected
69 infectious disease including TB. Part of the lack of data on HCV seroprevalence stems
70 from the fact that there is no recommendation for universal screening of person with TB
71 for HCV infection as there are for HIV testing [2].

72 Richard *et al*, [2] reported that HIV and HCV are both global public health
73 problems infections with HIV and or HCV may have an impact among those with TB.
74 The high presence of HCV co-infection among patients with TB in Georgia has the
75 potentials to have a major impact on TB management, treatment and control.

76 Hepatitis C virus is one of the deadly blood-borne viruses that has almost the
77 same route of transmission as of HIV, it is noted to have its major activity in the liver
78 where it causes inflammation of the liver, on the other hand, HIV and TB are closely
79 connected that they are often referred to as co-epidemics among confirmed TB patient,

80 however, drug or regimes given to TB or TB/HIV co-infection patient has hepatotoxicity
81 effect and can eventually lead to hepatocellular carcinoma

82 In this study, the seroprevalence of HCV and HIV in confirmed TB patient were
83 determined to provide an updated reference data for effective empiric management of
84 Tuberculosis patients with co-infection of HCV and HIV. Also, the possible
85 predisposition factor(s) to HCV and HIV coinfection in TB patients were identified.

86 **2.0. METHODOLOGY.**

87 **2.1 Study Area.**

88 The study area for this work was Federal Medical Centre (FMC), Ido Ekiti located
89 in Ekiti North senatorial district of Ekiti State, Nigeria.

90 **2.2. Study population**

91 The study population is Tuberculosis confirmed patients attending FMC, Ido
92 Ekiti. A total number of 500 samples were collected from TB confirmed patient after due
93 consultation with the patients.

94 **2.3 Sample collection**

95 About 5mls of blood was collected by venous puncture from the antecubital foci
96 of the arm after disinfecting the area with 70% alcohol. The blood was allowed to clot
97 and was spun at 1000rpm for 5 minutes and the serum was aseptically collected into
98 sterile cryovials bottles, appropriately labelled and stored at -20°C until the test was
99 performed.

100 **2.4 Sample processing**

101 **2.4.1 HCV Detection**

102 Clinotech diagnosis anti-HCV cassette detection test was used which is a rapid
103 direct binding procedure, which visually determines antibodies to hepatitis C infection.

104 2.4.2. Detection of HIV

105 The Abbot Determine HIV-1/2 was used in conjunction with STAT-PAK which
106 are in-vitro, visually ready, qualitative immunoassays for the detection antibodies to
107 HIV-1 and HIV-2 in human serum, plasma or whole blood.

109 2.5. Statistical analysis

110 The data generated from this study were analysed using SPSS version 16 (SPSS Inc.
111 Chigago IL).

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113 3.0 RESULT

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115 3.1. Response rate

116 A total of 500 questionnaires and consent forms were distributed to the patients
117 screened and out of 500 questionnaires distributed, 500 were returned indicating a 100%
118 response rate.

119 The overall seroprevalence of HCV, HIV and HCV/HIV antibodies in tuberculosis
120 confirmed patients are shown in table 1. It shows that out of 500 samples tested for HCV,
121 HIV antibodies, 10(2.0%) are positive for HCV, 21(4.2%) are positive for HIV and
122 3(0.6%) are positive for both HCV and HIV antibodies.

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124 **TABLE 1: OVERALL SEROPREVALENCE OF HCV, HIV AND HCV/HIV**
125 **ANTIBODIES IN TB CONFIRMED PATIENT.**

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INFECTION	NO OF SAMPLES	NO OF POSITIVE (%)
HIV	500	21(4.2)
HCV	500	10(2.0)
HIV/HCV	500	3 (0.6)

The demographic relationships in respect to sex are shown in table 2, table 3 and table 4 for HCV, HIV and HCV/HIV co-infection respectively. They revealed that out of 302 male subjects that participated, 7(2.32%) positive for HCV (Table 2), 14(4.64%) positive for HIV (Table 3) and 2 (0.66%) were positive for HCV/HIV antibodies (table 4), while out of 198 female, 3(1.52%) positive for HCV (Table 2), 7(3.54%) positive for HIV (Table 3), 1(0.51%) positive for HCV/HIV antibodies (table 4).

TABLE 2: SEROPREVALENCE OF HCV IN RELATION TO SEX.

SEX	NO POSITIVE (%)	P VALUE
Male	7.0 (2.32)	0.531
Female	3.0 (1.52)	

TABLE 3: SEROPREVALENCE OF HIV CONCERNING SEX

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SEX	NO POSITIVE (%)	P VALUE
Male	14.0 (4.64)	0.549
Female	7.0 (3.54)	

TABLE 4: SEROPREVALENCE OF HCV/HIV CO-INFECTION IN RELATION TO SEX

SEX	NO POSITIVE (%)	P VALUE
Male	2.0 (0.66)	0.824
Female	1.00 (0.51)	

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Age group distributions for HCV, HIV and HCV/HIV antibodies are shown in table 5, table 6 and table 7 respectively. In the age group 18-25, out of 4 (0.8%) subjects that participated, no subjects were positive for HCV and HIV antibodies as shown in table 5 and table 6 respectively. In age group 26-30, 20 (4.0%) subjects participated, 4 (0.8%) positive for HCV(Table 5), 2(0.4%) positive for HIV (Table 6), but no subject had HCV and HIV together (table 7). 70(14%) subjects are within 36-45 age group, 5(1%)

207 had HCV (Table 5), 10(2%) had HIV(table 6), 2(0.4%) had HCV/HIV antibodies (table
 208 7). In the age group of 46-55, 136(27.2%) participated, 1(0.2%), 5(1%), and 1(0.2%)
 209 were seropositive for HCV, HIV, and HCV/HIV antibodies and these were shown in
 210 table 5, table 6 and table 7 respectively. Out of 150(30%) subjects within 56-65 age
 211 group, 3(0.6%) were seropositive for HIV (Table 6), no subject was positive for HCV
 212 (Table 5) and HCV/HIV antibodies (table 7). In age 66-75, 70 (14%) participated,
 213 1(0.2%) was positive for HIV (Table 6), no seropositivity in HCV and HIV/HCV as
 214 shown in table 5 and table 7 respectively. 50(10%) subjects participated in age group 75-
 215 above, no seropositivity was recorded in both HCV and HIV as shown in table 5and table
 216 6 respectively.

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220 **TABLE 5: SEROPREVALENCE OF HCV IN RELATION TO AGE.**

223 AGE GROUP 224 VALUE 225 IN YEARS	222 NOT EXAMINED (%)	222 NO POSITIVE (%)	222 P
229 18-25	4 (0.8)	0 (0.0)	
230 26-35	20 (4.0)	4 (0.8)	
231 36-45	70 (14.0)	5 (1.0)	
232 46-55	136 (27.2)	1 (0.2)	0.000
233 56-65	150 (30.0)	0 (0.0)	
234 66-75	70 (14.0)	0 (0.0)	
235 75-above	50 (10.0)	0 (0.0)	
237 TOTAL	500 (100.0)	10 (2.0)	

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TABLE 6: SEROPREVALENCE OF HIV IN RELATION TO AGE.

AGE GROUP IN YEARS	NO EXAMINED (%)	NO POSITIVE (%)	P VALUE
18-25	4 (0.8)	0 (0.0)	0.000
26-35	20 (4.0)	2 (0.4)	
36-45	70 (14.0)	10 (2.0)	
46-55	136 (27.2)	5 (1.0)	
56-65	150 (30.0)	3 (0.6)	
66-75	70 (14.0)	1 (0.2)	
75-Above	50 (10.0)	0 (0.0)	
TOTAL	500 (100.0)	21 (4.2)	

TABLE 7: SEROPREVALENCE OF HCV/HIV CO-INFECTION IN RELATION TO AGE

AGE GROUP IN YEARS	NO EXAMINED (%)	NO POSITIVE (%)	P VALUE
18-25	4 (0.8)	0 (0.0)	0.002
26-35	20 (4.0)	0 (0.0)	
36-45	70 (14.0)	2 (0.4)	
46-55	136 (27.2)	1 (0.2)	
56-65	150 (30.0)	0 (0.0)	
66-75	70 (14.0)	0 (0.0)	
75-above	50 (10.0)	0 (0.0)	
TOTAL	500 (100.0)	3 (0.6)	

287 The risk factors associated with HCV, HIV and co-infection of HIV/HCV in TB
288 patient was based on patient self-report. Alcoholism, previous unprotected sex, multiple
289 sex partner, previous blood donation, Previous transfusion, Tattoos and History of Sexual
290 Transmitted disease are the risk factors. Out of 31 infected subjects, 14(45.2%) identified
291 with alcoholism, previous unprotected sex 11(35.5%), Multiple sex partner 10(32.3%),
292 Previous blood donation 8(25.8%), previous transfusion 7(22.6%), others are Tattoos
293 7(22.6%) and history of STD 3(9.68%). The risk factors were represented in the pie chart
294 shows in Figure I.

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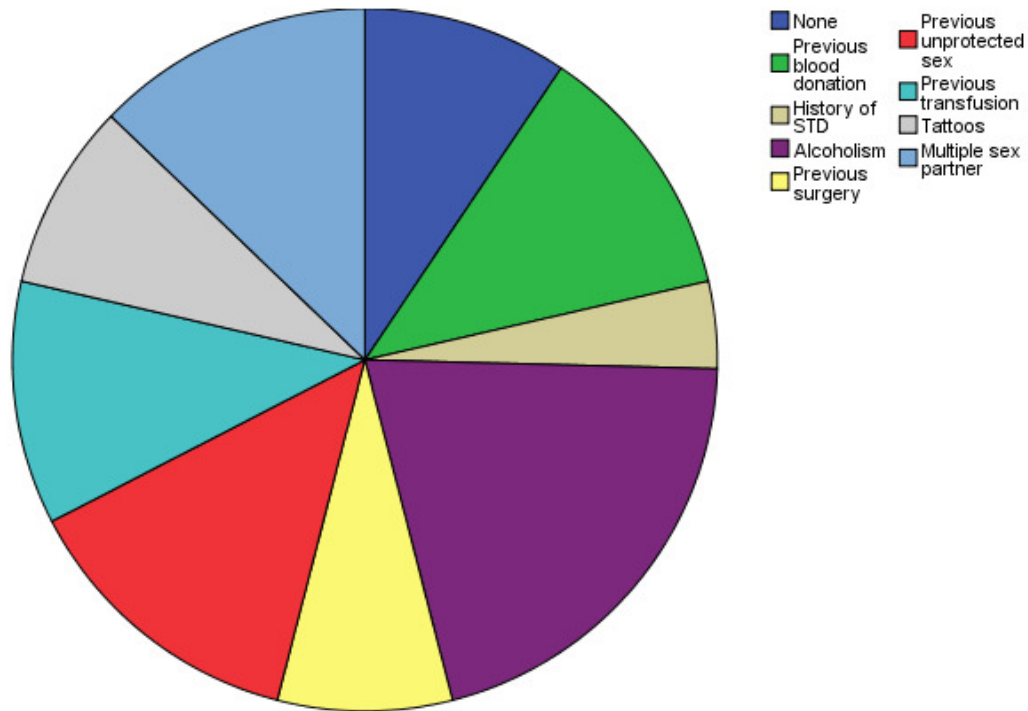
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UNDER PEER REVIEW

PATIENTS AS PREDISPOSING TO HIV AND HCV INFECTIONS



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304 **FIGURE I: PIE CHART OF RISK FACTORS WITH HCV AND HIV**

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307 **4.0 DISCUSSION**

308 HIV and HCV are both global public health problems. Infections with HIV and or HCV
309 may have a major impact on those with TB. HIV is the most important risk factor for the
310 development of TB among person infected with *M. tuberculosis* and both CDC and WHO
311 guidelines recommended offering HIV testing to those person diagnosed with TB disease
312 [2, 10]

313 However, the prevalence of HCV infection among persons with TB has been
314 poorly defined and few data are available from the most area around the world. One study
315 in the US suggested that veterans with HCV infection are at risk for other selected
316 infectious disease including TB [2]. Of 500 (100%) samples collected and tested against
317 HCV and HIV antibodies, HIV antibodies were positive in 21(4.2%) which might be

318 because HIV prevalence in Ekiti State is low and this agreed with previous work of
319 USAIDS [11]. This also agreed with the work of Idigbe *et al*, [12] that prevalence of HIV
320 in TB in Nigeria, Lagos to be specific is 5.3%. However, HCV antibodies were positive
321 in 10(2.0%) patients which also agreed with previous work of Mwangi [13] that the
322 prevalence in Ekiti State and Nigeria is low. This might be due to the proper screening of
323 donor which is one of the major predisposing factors to increase in the incidence of HCV
324 [13]. However, the prevalence of HCV among TB has been poorly defined and few data
325 are available around the world. Part of the lack of data on HCV seroprevalence stem from
326 the fact that there are no recommendations for universal screening of TB patient for HCV
327 as there for HIV testing [2]. Although, Halim and Ajayi [14] reported 12.3%
328 seroprevalence of HCV in Nigeria among the donor and the findings from Richard *et al*,
329 [2] reveals that patient with TB may have among the highest prevalence of HCV
330 infection.

331 The prevalence of HCV and HIV antibodies together in TB patient is 3(0.6%) which
332 appears to be low but can pose a major threat to the management of TB patients and this
333 agreed with previous work of Richard *et al*, [2] who reported a 0.4% prevalence rate of
334 HCV and HIV antibodies in TB patient in Georgia.

335 Gender wise distribution of seroprevalence of HCV and HIV in TB patients revealed that
336 although the number of males that participated in more than female but there was no
337 significant difference (0.531, 0.549, 0.824) between male and female for HCV, HIV, and
338 HCV/HIV co-infection respectively, which shows that HCV, HIV, and HCV/HIV can
339 infect any sex and this agreed with previous work of Richard *et al*, [2].

340 Age distribution revealed that age group 36-45 had the highest prevalence of HCV, HIV
341 and both HCV/HIV antibodies and this is statically significant (P-value 0.000 for HCV,
342 0.000 for HIV and HCV/HIV is 0.002). This might be because, at this age, subjects are
343 sexually active and are involved in some of the risk factor(s) that predispose them to the
344 infections. This agreed with the work of Watanabe *et al*, [15] which revealed 25-45years
345 as the most predispose age group to HIV. The predisposing risk factors to the
346 seroprevalence of HCV, HIV and HCV/HIV antibodies in the study population, showed
347 in figure I. Alcoholism is the highest of the factors, followed by previous unprotected
348 sex, multiple sex partner, previous transfusion, tattoos and the least is a history of

349 sexually transmitted disease. Although HCV and HIV are blood-borne diseases, the route
350 of transmission is similar, nevertheless, a number of investigations have indicated that
351 acquisition of HCV through sexual contact is uncommon and have suggested that HCV
352 is inefficiently transferred through this mechanism [16, 17] Despite these findings, a
353 number of studies had found that high-risk sexual behaviour or history of STD are
354 associated with an increased risk of HCV infection [16] and so, high-risk sexual
355 behaviours and /or a history of STI may be a maker for other risks that have been
356 implicated as mechanism of transmission of HCV. However, Richard *et al*, [2] revealed
357 that most common route of HCV transmission worldwide is through hematogenous
358 transmission, tattoos and nevertheless, hematogenous transmission may not be too
359 implicated in this study because in Ekiti state, the WHO guidelines of screening of blood
360 donor are followed strictly and this has contributed to the low prevalence of HCV, HIV in
361 this part of Nigeria [11].

362 **CONCLUSION**

363 Since, HCV and HIV co-infection in TB patients increased the risk of antituberculosis
364 drug-induced hepatotoxicity and that there is an even greater risk for drug-induced
365 hepatotoxicity among those undergoing treatment for TB who had both HCV and HIV
366 co-infection, to this end, more active screening for HCV should be done in this
367 population (TB) as was done for HIV. There is also a need to know the underlying health
368 status of TB patient as regards the HCV and HIV before administering drugs. Above all,
369 there is a need for a sample of TB patient to send to the laboratory for a liver function test
370 because of the effects of the regimes on the liver.

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372 **RECOMMENDATION**

373 It is recommended that there should be a universal screening of a person with TB for
374 HCV infection as there are for HIV testing.

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377 **Ethical consideration**

378 The ethical clearance for this research was granted by the Federal Medical Centre
379 (FMC) (Now Federal Teaching Hospital) Ido-Ekiti ethical committee after due processes
380 had been followed. Before the collection of the sample, information regarding the study
381 was explained to the subjects.

382 **Questionnaire and informed consent**

383 Questionnaire to obtain the demographic characteristics, possible risk factors and
384 other relevant information to the study as well as an informed consent were administered
385 to the participant.

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