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# Triple Infections of HBV, HCV and Malaria *Plasmodium Falciparum* Among HIV-Infected Individuals in Yenagoa, Bayelsa State, Nigeria

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# ABSTRACT

Infectious diseases continue to remain life-threatening and a significant public health problem globally. Patients with HIV frequently have concomitant HBV, HCV, and malaria infections; thus, this study was undertaken to describe the prevalence of HBV, HCV, and Malaria triple infection with HIV among patients presenting at the Federal Medical Centre, Yenagoa, Bayelsa State, Nigeria. In this study, 104 HIV-positive patients were recruited and evaluated for the presence of HBsAg, HCV, and Plasmodium falciparum with HBsAg rapid strips, anti-HCV antibodies ELISA kit (Dia. Pro), and SD Bioline RDT, following the respective manufacturer's instructions. The triple infection rate was 1.0% for HIV/HBV/HCV/Plasmodium falciparum. Other co-infections were 1.9% for HIV/HCV/Plasmodium falciparum, 2.9% for HIV/HBV/Plasmodium falciparum, and 1.9% for HIV/HBV/HCV, respectively. A higher HIV/HBV/HCV/Plasmodium falciparum triple infection occurred in the age group 21years-40 years (2.0%), females (1.3%), being single (2.3%), tertiary education holders (2.4%), students (4.3%), CD4 counts>500 cells/µl (4.0%) and Viral Load (VL)<20 copies/ml (2.0%). Higher HIV/HBV/HCV triple infections occurred in the age group>41 years (2.2%), males (3.5%), being single (2.3%), tertiary education holders (4.8%) and students (4.3%), having CD4 count 350 Cells/ µl- 499 Cells/µl (7.1%), viral load 20 copies/ml-999 copies/ml (2.1%) and being on TLD ART (1.9%). Higher HIV/HBV/MPF triple infections occurred in the age group 21 years-40 years (3.9%), males (3.5%), being married (3.6%), tertiary education holders (4.8%) and students (8.7%), having CD4 count>500 cells/µl (7.7%), viral load<20 copies/ml (3.8%) and being on TLD ART (2.9%). Higher HIV/HCV/MPF triple infections occurred in the age group 21 years-40 years (3.9%), females (2.7%), being single (4.7%), tertiary education holders (4.8%) and students (8.7%), having CD4 count>500 cells/µl (4.0%), viral load <20 copies/ml (2.0%) and being on TLD ART (1.9%). None of the sociodemographic and clinical variables was significantly associated (p>0.05) with triple infections. The present study has further confirmed the low occurrence (1.0%) of HIV/HBV/ Plasmodium falciparum among HIV-infected individuals in Yenagoa, Nigeria. Ages 21 years-40 years, females, being single, tertiary education holders, and students were more prone to triple infections. The concurrency of HIV/HCV/HBV and Malaria exists in Yenagoa, Nigeria. Therefore, it is recommended to perform routine screening of HIV-infected patients for simultaneous infection with HBV, HCV, and Malaria.

**Keywords:** Co-infections, HIV/HBV, HIV/HCV, HIV/Plasmodium falciparum, HIV/HBV/HCV/ Plasmodium falciparum, Triple infections

## INTRODUCTION

Malaria remains a significant health threat worldwide. Infectious diseases remain life-threatening and a significant public health problem globally [1]. Endemic regions for malaria are endemic to other infectious diseases that might affect the malaria infection [2,3]. Examples of such a common endemic infection sharing the same territory with malaria are Hepatitis B Virus (HBV)

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#### and Hepatitis C (HCV) [3-6].

Worldwide, HBV and HCV represent significant public health problems; around 300 million people are infected with HBV and HCV [7-9]. HBV and HCV infections are responsible for the majority of cases of chronic liver diseases, namely liver cirrhosis and hepatocellular carcinoma [9,10]. HBV and HCV co-infection is common among people living with Human Immunodeficiency Virus (HIV) infection (PLWH) because of common routes of transmission such as the exchange of blood or other body fluids during Intravenous Drug Use (IVDU), sexual contact, or mother-to-child-transmission-during the perinatal period [11-13]. If undiagnosed and untreated, this co-infection accelerates liver deterioration and increases morbidity and mortality in PLWH [13,14].

HIV remains a significant public health threat, with an additional risk of HBV and HCV co-infection [13]. Based on the report of the Joint United Nations Programme on HIV/ AIDS 2019 and the World Health Organization (WHO, 2017), approximately 38 million people have been diagnosed with HIV, 257 million with HBV, and 71 million with HCV globally at the end of 2020 [13,15,16]. Therapeutic advancements have greatly aided the eradication of the HBV, HCV, and HIV epidemics during the past few decades. Strategies have been put in place by the WHO and the Joint United Nations Programme on HIV/AIDS to encourage the global eradication of these viruses by 2030 [13,17,18]. Among other factors, the success of these strategies relies upon the testing and diagnosis of at least 90% of all persons living with HBV, HCV, and HIV infections as a necessary first step towards engagement in care and treatment [13,19,20].

To the best of our knowledge, there is a dearth of literature on the prevalence of HBV/HCV/*Plasmodium falciparum* triple infection among HIV-infected individuals in Bayelsa State, Nigeria. Studies on the triple infections of HBV, HCV, and *Plasmodium falciparum* among patients with HIV are limited in Bayelsa State, Nigeria. Therefore, this study was created to ascertain the prevalence of triple infections with HBV, HCV, and *Plasmodium falciparum* in HIV-positive patients.

#### MATERIALS AND METHODS

#### Study area

The study was conducted at the Federal Medical Centre, Yenagoa, Bayelsa State, Nigeria. This hospital is one of the main treatment facilities for HIV-infected patients in Bayelsa State, Southern Nigeria.

#### Study design

A hospital-based cross-sectional study design was adopted for the present study, which seeks to determine HBV, HCV, and *Plasmodium falciparum* triple infections among HIV-infected individuals attending Federal Medical Centre, Yenagoa, Bayelsa State, Nigeria.

#### **Ethics statement**

Administrative approval for this study was obtained from the management of the Federal Medical Centre, Yenagoa, Nigeria. The University of Port Harcourt Research Ethics Committee reviewed the work for ethical issues and approved the standards for research involving human beings. Before samples were taken and processed, everyone who participated gave informed consent.

#### Study population

We recruited 104 HIV-infected individuals attending the HIV outpatient clinic of Federal Medical Centre, Yenagoa, Nigeria, who willingly gave informed consent and volunteered to examine their blood samples. The study entailed screening for co-infections, clinical evaluation, and recording of demographic information such as the age of the participants, sex, marital status, educational background, occupation, and use of art.

#### Sample collection

The method of sample collection employed was the vein puncture technique. About 3 ml of venipuncture blood was collected in EDTA BA Vacutainer TM anti-coagulant tubes (BD, Franklin Lakes, USA), labeled with each patient's details. Plasma specimens were separated by centrifugation at 3000 rpm (revolution per minute) for 5 minutes. The plasma was stored at -20°C and used for laboratory analyses.

#### Serological analysis

Blood samples were taken and examined at the Virus & Genomics Research Unit, Department of Microbiology, University of Port Harcourt, for HBsAg, HCV, and Plasmodium falciparum antigen using appropriate test kits. Laboratory testing was carried out according to the manufacturer's instructions, and all tests were run using quality controls according to standard operating procedures.

#### Data analysis

SPSS version 20.0 was used to analyze the data (SPSS Inc. Chicago, IL, USA). Using Pearson's chi-square (X2) test or Fisher's exact test, as applicable, the prevalence of co-infections among HIV-infected individuals was compared to CD4+ T cell count, viral loads, sociodemographic factors, and the use of ART. A 5% significance level was used to establish the statistical significance for each analysis.

# RESULTS

#### Study population characteristics

Characteristics of the study group are highlighted in Table 1.

#### **Overall prevalence of triple infections**

Figure 1 represents the results obtained from the study. A total HIV seropositivity of 100% was obtained for all samples, reconfirming the HIV status of the study participants. Further analysis for HBsAg, anti-HCV antibodies, and malaria *Plasmodium falciparum* revealed 2.9% for HIV/HBV/Plasmodium falciparum, 1.9% for HIV/HBV/HCV, 1.9% for HIV/HCV/*Plasmodium falciparum*, and 1.0% for HIV/HBV/HCV, 1.9% for HIV/HCV/*Plasmodium falciparum*, and 1.0% for HIV/HBV/HCV, 1.9% for HIV/HCV/*Plasmodium falciparum*, and 1.0% for HIV/HBV/HCV.



## Age-specific HBV/HCV/Plasmodium falciparum triple infections

Higher HBV/HCV/*Plasmodium falciparum* triple infections occurred among age groups 21years–40 years (2.0%) than other age groups with 0.0% (Figure 2). These differences were not statistically associated (p=0.59).



No. Tested HBV/HCV/Plasmodium falciparum (%)

Figure 2. HBV/HCV/Plasmodium falciparum triple infections in relation to age

# Sex-specific HBV/HCV/Plasmodium falciparum triple infections

Higher HBV/HCV/Plasmodium falciparum triple infections were observed among females (1.3%) than in males (0.0%) (Figure 3). No significant association existed between HBV/HCV/*Plasmodium falciparum* triple infections and sex (p=0.53).



Figure 3. HBV/HCV/Plasmodium falciparum Triple Infections in relation to Sex

#### Marital Status-specific HBV/HCV/Plasmodium falciparum triple infections

Higher HBV/HCV/*Plasmodium falciparum* triple infections occurred among singles (2.3%) than the married (0.0%) and divorced (0.0%) (Figure 4). No significant association existed between HBV/HCV/*Plasmodium falciparum* triple infections and marital status (p=0.49).



Figure 4. HBV/HCV/Plasmodium falciparum Triple Infections in relation to marital status

#### Educational Background-specific HBV/HCV/Plasmodium falciparum triple infections

Higher HBV/HCV/*Plasmodium falciparum* triple infections were observed among those with tertiary educational background (2.4%) than those with secondary education (0.0%) and primary education with 0.0% (Figure 5). No significant association was found between HBV/HCV/*Plasmodium falciparum* triple infections and educational background (p=0.68).





#### Occupation-specific HBV/HCV/Plasmodium falciparum triple infections

Higher HBV/HCV/*Plasmodium falciparum* triple infections were observed among students (4.3%) than in other occupations with 0.0% (Figure 6). No significant association existed between HBV/HCV/*Plasmodium falciparum* triple infections and occupations (p=0.61).





#### CD4 counts-related specific HBV/HCV/Plasmodium falciparum triple infections

In terms of CD4 counts, higher HBV/HCV/*Plasmodium falciparum* triple infections (4.0%) were observed for participants with CD4 counts>500 cells/ $\mu$ l than <200, 200-349 and 350-499 cells/ $\mu$ l had the least, 0.0% (Figure 7). No significant association existed between HBV/HCV/*Plasmodium falciparum* triple infections and CD4 counts (p=0.39).



**Figure 7.** HBV/HCV/*Plasmodium falciparum* Triple Infections in relation to CD4 Counts

# Viral loads-related specific HBV/HCV/Plasmodium falciparum triple infections

In terms of viral loads, higher HBV/HCV/*Plasmodium falciparum* triple infections (0.0%) were recorded for participants that had<20 copies/ml than those with 20-999 copies/ml and>1000 copies/ml with 0.0% (Figure 8). No significant association existed between HBV/HCV/*Plasmodium falciparum* triple infections and viral loads (p=0.62).





#### Sociodemographic Characteristics and other triple infections

Higher HIV/HBV/HCV triple infections occurred in the age group >41 years (2.2%), males (3.5%), being single (2.3%), tertiary education holders (4.8%) and students (4.3%), having CD4 count 350 Cells/ $\mu$ l-499 Cells/ $\mu$ l (7.1%), viral load 20-999 copies/ml (2.1%) and being on Tenofovir, Lamivudine, and raltegravir (TLD) ART (1.9%) (Table 1).

Higher HIV/HBV/ Malaria *Plasmodium falciparum* triple infections occurred in the age group 21 years-40 years (3.9%), males (3.5%), being married (3.6%), tertiary education holders (4.8%) and students (8.7%), having CD4 count >500 cells/µl (7.7%), viral load <20 copies/ml (3.8%) and being on Tenofovir, Lamivudine, and Dolutegravir (TLD) ART (2.9%), as in Table 1.

Higher HIV/HCV/Malaria *Plasmodium falciparum* triple infections occurred in the age group 21-40 years (3.9%), females (2.7%), being single (4.7%), tertiary education holders (4.8%) and students (8.7%), having CD4 count >500 cells/ $\mu$ l (4.0%), viral load <20 copies/ml (2.0%) and being on Tenofovir, Lamivudine, and dolutegravir (TLD) ART (1.9%) as in Table 1.

Variables	Categories	No. Tested	HIV/HBV/HCV (%)	HIV/HBV/MPF (%)	HIV/HCV/ MPF (%)	HIV/HBV/HC/ MPF (%)	Chi-Square analysis
Age groups (Years)	20-Aug	8	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	P = 0.59
	21-40	51	1(2.0)	2 (3.9)	2 (3.9)	1(2.0)	
	41 & above	45	1 (2.2)	1 (2.2)	0 (0.0)	0 (0.0)	
Sex	Females	75	1(1.3)	2 (2.7)	2 (2.7)	1(1.3)	P = 0.53
	Males	29	1 (3.5)	1 (3.5)	0 (0.0)	0 (0.0)	
Marital Status	Singles	43	1(2.3)	1(2.3)	2 (4.7)	1(2.3)	P = 0.49
	Married	56	1 (1.8)	2 (3.6)	0 (0.0)	0 (0.0)	
	Divorced	5	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	
Educational Background	Primary	17	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	P = 0.68
	Secondary	43	0 (0.0)	1(2.3)	0 (0.0)	0 (0.0)	
	Tertiary	42	2 (4.8)	2 (4.8)	2 (4.8)	1(2.4)	
	None	2	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	
Occupations	Self-Employed	27	1 (3.7)	0 (0.0)	0 (0.0)	0 (0.0)	P = 0.61
	Unemployed	10	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	
	Business/ Trader	29	1 (3.5)	1 (3.5)	0 (0.0)	0 (0.0)	
	Students	23	1(4.3)	2 (8.7)	2 (8.7)	1(4.3)	
	Artisans	7	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	
	Civil Servants	8	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	
CD4 Counts (Cells/ µl)	<200	54	1(1.9)	1(1.9)	0 (0.0)	0 (0.0)	P = 0.39
	200-349	10	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	
	350-499	14	1(7.1)	0 (0.0)	0 (0.0)	0 (0.0)	
	500 & above	26	0 (0.0)	2(7.7)	1 (4.0)	1(4.0)	
Viral Loads (Copies/ml)	<20	53	1(2.0)	2 (3.8)	1(2.0)	1(2.0)	P = 0.62
	20-999	41	1 (2.4)	1 (2.4)	0 (0.0)	0 (0.0)	
	1000 & above	10	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	
ART Drugs	TLD	103	2 (1.9)	3 (2.9)	2 (1.9)	1(0.9)	
	ABC/BTC/ EFC	1	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	
Total		104	2(1.9)	3(2.9)	2(1.9)	1 (1.0)	
HBV = Hepatitis B Virus, HCV = Hepatitis C Virus, HIV = Human Immunodeficiency Virus, MPF = Malaria Plasmodium falciparum							

Table 1. Patients Characteristics

# DISCUSSION

HIV and malaria have similar risk factors, and HIV/HBV/HCV co-infection speeds up the development of AIDS, which results in millions of deaths annually throughout the world [21]. The study shows interactions between HBV, HCV, and malaria *Plasmodium falciparum* infections among HIV-infected people in Yenagoa, Bayelsa State, Nigeria, with some developing co-infections. As far as we know, no studies have been conducted in Yenagoa or the South-South region that considered these four illnesses simultaneously. Unfortunately, epidemiological information on HIV/HBV, HIV/HCV, and HIV/HBV/HCV co-infections among HIV patients is scarce and inconsistent domestically and internationally [13].

The study presents the fundamental demographic and clinical traits of co-infections with HBV, HCV, and malaria (*Plasmodium falciparum*) in HIV-infected patients. This study's total prevalence of co-infection with HIV and HBV/HCV/malaria (*Plasmodium falciparum*) was 1.0%. HIV mono-infection was 92.3% among them, while the co-infection rates for HIV/HBV/malaria, HIV/HCV/malaria, and HIV/HBV/HCV were 2.9%, 1.9%, and 1.9%, respectively. The results of our study agreed with those of other research from Nigeria and elsewhere.

The 1.0% HBV/HCV/malaria *Plasmodium falciparum* prevalence reported in this study is comparable to the 1.9% HIV/HBV/HCV and malaria co-infection reported by Mohammed in Gedarif, Eastern Sudan [16]. In our analysis, triple infections with HIV, HBV, HCV, and *Plasmodium falciparum* only occurred in those aged 21 to 40 (2.0%), women (1.3%), singles (2.3%), those with university education (2.4%), and students (4.3%). As revealed in Gedarif, Eastern Sudan, none of these sociodemographic factors was substantially linked (p>0.05) with triple infections.

The 2.9% reported for triple infections with HIV, HBV, and *Plasmodium falciparum* in this study is higher than in previous studies. Kogi State had a 0.5% HBV/*Plasmodium falciparum* co-infection, according to Omatola & Okolo [22]. 1.0% co-infection between malaria and HBV was observed by Adeleke, and 2.2% was reported by Afolabi and Bakare [23,24]. However, the 5.0% Okonko recorded in a previous study in Port Harcourt is lower than it [25].

This study's 2.9% reported rate of triple infections with HIV, HBV, and Plasmodium falciparum is greater than the 0.0% reported

by Okonko in Port Harcourt, Nigeria [26]. It also did not match the conclusions of other earlier investigations. According to Helegbe, Ghana has a 0.7% prevalence of triple infections with HIV, HBV, and *Plasmodium falciparum* malaria [1]. The prevalence rates of 3.3%, 4.3%, and 6.0% were previously reported in various places in Nigeria and the Gambia, respectively, they were higher than the figure recorded in the current study [27-29]. That varies from Anabire results, who reported that 1.9% of patients had *P. falciparum*/HBV co-infection and similarly recorded opposing values in their study [30,31].

In terms of sociodemographic factors linked to HIV/HBV/Malaria *Plasmodium falciparum* triple infections, males (3.5%), married individuals (3.6%), those with university education (4.8%), and students (8.7%) all experienced greater rates. As revealed by Mohammed in Gedarif, Eastern Sudan, none of these sociodemographic factors was substantially linked (p > 0.05) with triple infections [9].

Of these triple infections, HIV/HBV/HCV was 1.9%. This value is less than the 16.9% recorded for triple infection with HBV, HCV, and HIV in Abeokuta, Ogun State, Nigeria [32]. Several seroprevalences of HIV/HBV/HCV infections were reported in Nigeria in earlier research. According to Forbi, 7.2% of people worldwide have HIV, HBV, or HCV [33]. In Central Nigeria, Balogun reported a seroprevalence of 3.9% for HIV/HBV/HCV, Okeke recorded a seroprevalence of 6.5%, and Oti et al. (2021) reported a seroprevalence of 2.8% [34,35].

In the same study area, Aaron and Cookey reported values of 0.0% and 0.0%, respectively (Port Harcourt, Nigeria) [36,37]. Moreover, the 1.9% reported here exceeds the 1.0% frequency in Central Nigeria that Pennap reported [38]. It is also greater than the 1.3% and 0.5% found in Edo-Ekiti and Benin, respectively [39,40]. That is greater than the 0.34% reported for triple infection with HIV, HBV, and HCV by Bhattarai [41]. In contrast to the 1.83% reported by Agarwal, the seroprevalence of 1.9% found here is favorable [42].

In other places, rates of HIV, HBV, and HCV triple infection of 19.1% and 10.4%, respectively, were observed in China and Myanmar [43]. According to Gupta and Singh, North India has a triple infection prevalence of 5.32% and 2.43%, respectively [44]. According to Musyoki , the prevalence of HIV/HBV/HCV infections in South Africa was 29.4% [45]. According to Lonita, 4.4% and 19.0% of Nepal's population are triply infected with HIV, HBV, and HCV [46]. At Kumasi, Ghana, Boateng found a prevalence of co-infections with HIV, HBV, and HCV of 18.0% [47]. According to Pappoe, Ghana had a 6.1%, 0.5%, and 0.0% prevalence of HIV, HBV, and HCV co-infections, respectively [48]. Similarly, the combined HIV/HBV/HCV prevalence reported by Shrestha in a systematic review study of Nepal from 1990 to 2020 was 1.3% [49]. In Eastern Nepal, 2.53% of cases of HIV/HBV/HCV triple infections were reported by Shrestha in 2022.

It is essential to note the increased prevalence of HBV and HCV dual infection in this study compared to prior studies [35]. Furthermore, these studies have shown that triple HIV/HBV/HCV infections or dual HIV/HBV/HCV infections are the most prevalent. However, the incidence of these infections depends on risk categories, the type of exposure involved, and geographic locations [43, 50-52].

In terms of sociodemographic factors, greater rates of HIV/HBV/HCV triple infections occurred in people over the age of 41 (2.2%), in men (3.5%), in singles (2.3%), in those with university education (4.8%), and in students (4.3%). None of these sociodemographic factors showed a meaningful correlation (p>0.05). This result contrasts with that of studies by Shrestha, which found that people under 35 were more at risk of co-infection, and Choy, which found that people in their 30s, 40s, and 50s had considerably higher rates of co-infection [13,53]. The current finding conflicts with that of Oti, who claimed that only educational status was statistically associated with HIV/HCV co-infection in Central Nigeria [35].

*Plasmodium spp.* and HCV are known to infect liver cells; hence, given that the hepatitis C and malaria epidemics overlap in some parts of the world, it is conceivable that they might infect and reproduce in the same cell. Moreover, it is conceivable that these two illnesses could co-infect, in which case one pathogen could influence the other's severity to rise or fall and vice versa. Furthermore, despite substantial research on HCV attachment and entrance due to the virus's rising prevalence worldwide, more research needs to be done on plasmodium entry into host hepatocytes. Investigating the co-infection of the three diseases was imperative since it would aid in creating novel treatments and testing equipment. In Yenagoa, Bayelsa State, Nigeria, the study found 1.9% triple infections with HIV, HCV, and *Plasmodium falciparum*.

In contrast to the 10.9% reported by Asaga in Abuja, Central Nigeria, this study's 1.9% reported rate of triple infections with HIV, HCV, and *Plasmodium falciparum* is lower. Sociodemographic indices revealed numerous variations in prevalence across the various groupings. HIV/HCV/malaria *Plasmodium falciparum* triple infections were more common in people aged 21years to 40 years (3.9%), in women (2.7%), in singles (4.7%), in those with tertiary education (4.8%), and in students (8.7%). As revealed by Mohammed in Gedarif, Eastern Sudan, none of these sociodemographic factors was substantially linked (p>0.05) with triple infections [9]. This observation disputes the assertion made by Ouwe-Missi-Oukem-Boyer that age is a significant confounding factor in their setting. Multivariate analysis, on the other hand, suggests that *P. falciparum* and HCV interact at the hepatic level, with *P. falciparum* emerging more slowly in HCV chronic carriers [54]. According to Asaga, in HIV-infected people, co-infection rates for malaria and HCV were 8.7% and 11.2%, respectively.

Contrary to a study by Asaga, which found higher rates in the younger age range of 15 years to 25 years in Abuja, Nigeria, the higher rates of HIV/HCV/*Plasmodium falciparum* triple infections were recorded in older age groups >41 years (2.2%). HCV's tendency could explain the discrepancy in the results to self-limit as a child gets older and eventually acquires the status of a chronic carrier. Also, this observation represents a complete departure from a related study in Egypt, where the prevalence ranged from 0.2% to 5.0% [55].

Asaga study, which found higher rates in married women than singles and agrees with our study in that 0.0% prevalence was also reported for participants who had recently divorced, contradicts the higher rates of HIV/HCV/*Plasmodium falciparum* triple infections reported in singles than other marital groups. The results of Todd, who found that participants with HCV co-infection were less likely

to be married, are consistent with our data [56]. In the current study, lifestyle, social activities, and awareness may be the primary deductive reasoning factors contributing to this [57-59].

In this study, triple infections with HIV, HCV, and Plasmodium falciparum only occurred in those with tertiary education. Asaga showed higher rates in women who attended Islamic Quranic schools (20%) compared to postsecondary education level (18.6%). This observation is in contrast to their findings. This finding might be explained by a person's lifestyle, social interactions, and awareness. The results of this study showed complete agreement with the claim made by Oni that greater levels of education increase the likelihood of sexual adventures, which frequently involve several partners [60]. The results of this investigation showed a complete divergence from those of the Asaga study. However, it contrasts with the findings of Todd. However, the results of Todd showed that people with HCV co-infection had a lower likelihood of having a tertiary education [56].

In addition, our findings indicate that females are more likely than males to have co-infections with HIV/HCV/Plasmodium falciparum and HIV/HBV/HCV/Plasmodium falciparum. The study by Shrestha, which indicated a higher likelihood among males than females, did not support this finding [13]. In contrast, the males were more likely to be coinfected in the study by Lonita [46]. This observation could be a result of female sexual promiscuity. Age and gender of the male were risk variables for co-infection [3]. It is necessary to determine the precise cause of these differences from other prospective cohort studies.

To the best of our knowledge, there is no information on the relationship between CD4+ T cell count, viral load, and coinfections with HIV, HBV, HCV, and *Plasmodium falciparum* in Bayelsa State, Nigeria. Only HIV-infected patients with CD4 counts > 500 cells/l (4.0%) and viral loads (VL) 20 copies/ml (2.0%) had a triple infection. Increased HIV/HBV/HCV triple infections were seen in people with CD4 counts between 350 and 499 cells/l (7.1%), viral loads between 20 copies/ml and 999 copies/ml (2.1%), and TLD ART use (1.9%). Having a CD4 count>500 cells/l (7.7%), a viral load of<20 copies/ml (3.8%), and being on Tenofovir, Lamivudine, and Dolutegravir (TLD) ART (2.9%) were also associated with an increased risk of HIV/HBV/MPF triple infections.

Additionally, having a CD4 count >500 cells/l (4.0%), a viral load of <20 copies/ml (2.0%), and being on Tenofovir, Lamivudine, and Dolutegravir (TLD) ART (1.9%) were associated with greater HIV/HCV/Malaria *Plasmodium falciparum* triple infections. In support of Kakisingi who noted that none of the CD4 count and viral load tested in Lubumbashi, DR Congo, exhibited any statistically significant difference, none of these clinical variables was significantly linked (p > 0.05) with triple infections [61]. Our findings go counter to those of Ebong, who claimed that correlations between the CD4 count and the kind of illness (*Plasmodium*) or even co-infection (HIV/HCV) had existed [21]. This finding conflicts with research by Bhattarai and Shrestha, which found co-infection was less common in HIV patients with CD4 cells above 200 cells/mm<sup>3</sup> [13]. However, a CD4 count under 200 cells/l still carries a substantial risk for opportunistic infections [62,63]. According to George and Wang, the declining CD4 cell count is a sign of HIV progression and immunological dysfunction, as well as co-infections and various opportunistic infections [13,64,65]. Nevertheless, this study's scenario was the opposite. However, research carried out in regions where HIV and malaria are highly endemic discovered a very high viral load and a meagre CD4 count in related individuals [66,67].

These pathogens' overlap is a significant health concern [35,68]. However, more research is required on the seroprevalence of HBV, HCV, and malaria (*Plasmodium falciparum*) among HIV-positive patients in Nigeria. When compared to the WHO norm, we discovered that the overlapping infections of HIV/HBV/*Plasmodium falciparum*, HIV/HCV/*Plasmodium falciparum*, HIV/HBV/HCV, and HIV/HBV/HCV/*Plasmodium falciparum* were 2.9%, 1.9%, 1.9%, and 1.0%, respectively [35]. Chronic HCV carriers' interactions with malaria parasites may prevent the latter from spreading, which may aid in developing novel malaria treatment strategies [3]. The study also urges establishing a routine HBV, HCV, and Plasmodium screening program employing malaria Rapid Diagnostic Tests (RDT) to identify the preventative actions that must be performed in the treatment of HIV-infected people [69].

# CONCLUSION

According to this study, the rates of triple infection for HIV/HBV/HCV/*Plasmodium falciparum* were 1.0%, 1.9%, 2.9%, and 1.9%, respectively. This study reveals for the first time that Yenagoa, Bayelsa State, Nigeria, has concurrent HIV/HCV/HBV and malaria cases. Hence, systematic testing for co-infection with HIV, HBV, HCV, and malaria in HIV-infected individuals should be adopted. It is necessary to do additional research on clinical patterns and risk variables to comprehend how HIV/HCV/HBV and malaria co-infection emerged in the study area.

# REFERENCES

- 1. Helegbe, G.K., et al., Seroprevalence of malaria and hepatitis B coinfection among pregnant women in tamale metropolis of Ghana: a cross-sectional study. *Canadian Journal of Infectious Diseases and Medical Microbiology*, **2018**.
- 2. Boraschi, D., et al., Immunity against HIV/AIDS, malaria, and tuberculosis during co-infections with neglected infectious diseases: recommendations for the European Union research priorities. *PLoS Neglected Tropical Diseases*, **2008**.2(6): p. e255.
- 3. Gasim, G.I., and Adam, I., Hepatitis B Hepatitis C virus and Malaria co-infection. Int J Vaccine Immunizat, 2015.1(1): p. 1-3.
- 4. World Health Organization. Anonymous Global distribution of hepatitis A. B and C, 2001. 2002: p. 45-47.
- 5. Morel, C.M., Toure, Y.T., and Dobrokhotov, B., The mosquito genome--a breakthrough for public health. *Science*, **2002**. 298(5591): p. 79.

- Snow, R.W., et al., The global distribution of clinical episodes of Plasmodium falciparum malaria. *Nature*, 2005. 434(7030): p. 214-217.
- 7. Jiwani, N., and Gul, R.B., A silent storm: hepatitis C in Pakistan. Journal of pioneering medical sciences, 2011.1(3): p. 89.
- 8. El-Serag, H.B., Epidemiology of viral hepatitis and hepatocellular carcinoma. Gastroenterology, 2012. 142(6): p. 1264-1273.
- 9. Mohammed, H.B., et al., Prevalence of Hepatitis B, Hepatitis C, HIV and Malaria Co Infection among Patients Infected with Visceral Leishmaniasis in Gedarif, Eastern Sudan. *Glob J Infect Dis Clin Res*,**2013**. 2(1): p. 021-024.
- 10. Abdallah, T.M., Mohamed, M.H., and Ali, A.A., Seroprevalence and epidemiological factors of hepatitis B virus (HBV) infection in Eastern Sudan. *International Journal of Medicine and Medical Sciences*, **2011**. 3(7): p. 239-241.
- 11. Vallet-Pichard, A., and Pol, S., Hepatitis viruses and human immunodeficiency virus co-infection: pathogenisis and treatment. *Journal of hepatology*, **2004**.41(1): p. 156-166.
- 12. Modi, A.A., and Feld, J.J., Viral hepatitis and HIV in Africa. AIDS reviews, 2007. 9(1): p. 25-39.
- Shrestha, L.B., et al., Co-infection of Hepatitis B and Hepatitis C among HIV-infected patients: A cross-sectional study from tertiary care hospital of eastern Nepal. *Plos one*, 2022.17(3): p. e0264791.
- 14. Joshi, D., et al., Increasing burden of liver disease in patients with HIV infection. The Lancet, 2011. 377(9772): p. 1198-1209.
- 15. World Health Organization. Global health sector strategy on viral hepatitis 2016-2021. Towards ending viral hepatitis, 2016.
- Case, K.K., et al., Summarizing the results and methods of the 2019 Joint United Nations Programme on HIV/AIDS HIV estimates. AIDS (London, England), 2019. 33(3):p. S197.
- 17. World Health Organization. Consolidated guidelines on the use of antiretroviral drugs for treating and preventing HIV infection: recommendations for a public health approach. *World Health Organization*, **2016**.
- Assefa, Y., and Gilks, C.F., Ending the epidemic of HIV/AIDS by 2030: Will there be an endgame to HIV, or an endemic HIV requiring an integrated health systems response in many countries?. *International Journal of Infectious Diseases*, 2020.100: p. 273-277.
- 19. Levi, J., et al., Can the UNAIDS 90-90-90 target be achieved? A systematic analysis of national HIV treatment cascades. *BMJ global health*, **2016**. 1(2): p. e000010.
- Bain, L.E., Nkoke, C., and Noubiap, J.J., UNAIDS 90–90–90 targets to end the AIDS epidemic by 2020 are not realistic: comment on "Can the UNAIDS 90–90–90 target be achieved? A systematic analysis of national HIV treatment cascades". *BMJ global health*, 2017.2(2): p. e000227.
- 21. Ebong, S.B., et al., Biological and Immunological Profile of HIV Voluntary Consultants with HCV and Plasmodium Coinfection in One Accredited Treatment Center in Cameroon.
- 22. Omatola, C.A., and Okolo, M.L., Hepatitis B and Asymptomatic Malaria Infection among Pregnant Women in a Semiurban Community of North-Central Nigeria. *Journal of environmental and public health*, **2021**.
- 23. Adeleke, M.A., et al., Sero-prevalence of malaria, hepatitis b and syphilis among pregnant women in Osogbo, Southwestern Nigeria. *J Infect Dis Immun*,2013.5(2): p. 13-17.
- 24. Afolabi, O.J., and Bakare, T.P., Malaria, hepatitis B and their co-infection among pregnant women visiting maternity centers in Akure, Nigeria. *World News of Natural Sciences*, **2022**. 45: p. 93-102.
- 25. Okonko, I.O., et al., Dual infection of HIV and malaria among HIV-infected individuals in Port Harcourt, Nigeria. *South Asian Journal of Parasitology*, **2021**.5(1): p. 26-31.
- Okonko, I.O., et al., Serological Prevalence of Hepatitis B Virus Among Patients Attending OB Lulu Briggs Health Centre in Port Harcourt, Rivers State, Nigeria. Academia Arena, 2023.15(3): p. 1-9.
- 27. Abah, A.E., Onoja, H., and Amadi, F.I., Prevalence of malaria and hepatitis B virus infections among pregnant women attending federal medical center, Owerri. *South Asian J Parasitol*, **2019.3**: p. 1-5.
- Abah, A.E., and Udoidang, I.N., Co-infection of malaria and hepatitis B virus in Port Harcourt, Rivers State, Nigeria. *International Journal of Infection*, 2019. 6(4).
- 29. Adeleke, M,A, et al., Sero-prevalence of malaria, hepatitis b and syphilis among pregnant women. *American Journal of Microbiology*, **2013**.4(1): p. 20.
- 30. Anabire, N.G., et al., Prevalence of malaria and hepatitis B among pregnant women in Northern Ghana: Comparing RDTs with PCR. *PloS one*, **2019**.14(2): p. e0210365.
- 31. Anabire, N.G., et al., Impact of malaria and hepatitis B co-infection on clinical and cytokine profiles among pregnant women. *PloS one*, 2019. 14(4): p. e0215550.

- 32. Ogwu-Richard, S.O., et al., Triple positivity of HBsAg, anti-HCV antibody, and HIV and their influence on CD4+ lymphocyte levels in the highly HIV infected population of Abeokuta, Nigeria. *African Health Sciences*, **2015**. 15(3): p. 719-727.
- 33. Forbi, J.C., et al., The role of triple infection with hepatitis B virus, hepatitis C virus, and human immunodeficiency virus (HIV) type-1 on CD4+ lymphocyte levels in the highly HIV infected population of North-Central Nigeria. *Memorias do Instituto Oswaldo Cruz*, 2007.102: p. 535-537.
- Okeke, T.C., et al., Coinfection with hepatitis B and C viruses among HIV positive pregnant women in Enugu south east, Nigeria. Nigerian Journal of Medicine, 2012.21(1): p. 57-60.
- 35. Oti, V., et al., Epidemiologic survey of HBV, HCV and HIV infections in a pregnant women population in Central Nigeria: a cross-sectional study. *J Infect Dis Epidemiol*, **2021**.7: p. 194.
- 36. Aaron, U.U., Okonko, I.O., and Frank-Peterside, N., The Prevalence of Hepatitis E, Hepatitis C and Hepatitis B Surface Antigenemia in HAART Experienced People Living with Human Immunodeficiency Virus (HIV) in Rivers State, Nigeria. *Journal* of Biomedical Sciences, 2021. 10 (S4).
- Okonko, I.O., Cookey, T.I., and Frank-Peterside, N., Zero Prevalence of HIV and HCV Coinfection in the Highly HIV-infected Population of Rivers State, Nigeria. Asian Journal of Research in Medical and Pharmaceutical Sciences, 2021. 10(3): p. 9-16.
- Pennap, G.R., Muazu, I.F., and Fatima, M., Parallel and Overlapping Hepatitis B and C Virus Infection among Pregnant Women Attending Antenatal in a Rural Clinic in Northern Nigeria. *International Journal of Current Microbiology and Applied Sciences*, 2015.4(5): p. 16-23.
- 39. Ugbebor, O., et al., The prevalence of hepatitis B and C viral infections among pregnant women. North American journal of medical sciences, 2011.3(5): p. 238.
- 40. Esan, A.J., et al., Sero-prevalence of hepatitis B and hepatitis C virue co-infection among pregnant women in Nigeria. Am J Biomed Res, 2014. 2(1): p. 11-15.
- 41. Bhattarai, M., et al., Epidemiological profile and risk factors for acquiring HBV and/or HCV in HIV-infected population groups in Nepal. *BioMed research international*, **2018**.
- 42. Agarwal, N., et al., Hep atitis B or hepatitis C: the bigger threat in multiple infected HIV positive blood donors. *J Clin and Diagn Res*, **2011**.5: p. 766-768.
- 43. Zhou, Y.H., et al., Comparison of HIV-, HBV-, HCV-and co-infection prevalence between Chinese and Burmese intravenous drug users of the China-Myanmar border region. *PloS one*, **2011**. 6(1): p. e16349.
- 44. Gupta, S., and Singh, S., Hepatitis B and C virus co-infections in human immunodeficiency virus positive North Indian patients. World Journal of Gastroenterology: WJG, 2006. 12(42): p. 6879.
- Musyoki, A.M., et al., Active co-infection with HBV and/or HCV in South African HIV positive patients due for cancer therapy. Journal of medical virology, 2015.87(2): p. 213-221.
- 46. Ionita, G., et al., Seroprevalence of hepatitis B virus and hepatitis C virus co-infection among people living with HIV/AIDS visiting antiretroviral therapy centres in Nepal: a first nationally representative study. *International Journal of Infectious Diseases*, 2017.60: p. 64-69.
- 47. Boateng, R., et al., Sero-prevalence of Hepatitis B and C viral co-infections among HIV-1 infected ART-naïve individuals in Kumasi, Ghana. *PloS one*, **2019**.14(4): p. e0215377.
- 48. Pappoe, F., et al., Sero-prevalence of hepatitis B and C viral infections in Ghanaian HIV positive cohort: a consideration for their health care. *BMC Infectious Diseases*, **2019**.19(1): p. 1-8.
- 49. Shrestha, D.B., et al., Prevalence of Hepatitis B and C among HIV Infected Patients in Nepal over 1990-2020. *Kathmandu University Medical Journal*, **2021**.19(1):p. 132-139.
- 50. Zhang, C., et al., High prevalence of HIV-1 and hepatitis C virus coinfection among injection drug users in the southeastern region of Yunnan, China. *Journal of acquired immune deficiency syndromes (1999)*, **2002**. 29(2): p. 191-196.
- 51. Wang, Y.C., et al., A study on the prevalence rates of human immunodeficiency virus, hepatitis B virus and hepatitis C virus infections in intravenous drug users. *Zhonghua liu Xing Bing xue za zhi= Zhonghua Liuxingbingxue Zazhi*, **2006**. 27(9): p.777-779.
- 52. Bao, Y.P., and Liu, Z.M., Systematic review of HIV and HCV infection among drug users in China. *International journal of STD* & *AIDS*, 2009.20(6): p. 399-405.
- Choy, C.Y., et al., Factors associated with hepatitis B and C co-infection among HIV-infected patients in Singapore, 2006–2017. *Tropical Medicine and Infectious Disease*, 2019. 4(2): p. 87.
- Ouwe-Missi-Oukem-Boyer, O., et al., Hepatitis C virus infection may lead to slower emergence of P. falciparum in blood. *PLoS One*, 2011.6(1):p. e16034. https://doi.org/10.1371/journal.pone.0016034.

- 55. Galal, S.M., et al., Chronic viral hepatitis C in pediatric age group; assessment of viral activity and hepatic fibrosis by 1H magnetic resonance spectroscopy and diffusion weighted imaging in asymptomatic patient. *The Egyptian Journal of Radiology and Nuclear Medicine*, **2016**. 47(3): p. 739-748.
- 56. Todd, C.S., et al., HIV, hepatitis C, and hepatitis B infections and associated risk behavior in injection drug users, Kabul, Afghanistan. *Emerging infectious diseases*, **2007**.13(9): p. 1327.
- Atrah, H.I., and Ahmed, M.M., Hepatitis C virus seroconversion by a third generation ELISA screening test in blood donors. *Journal of clinical pathology*, 1996.49(3): p. 254-255.
- 58. Brady, C.W., and Muir, A.J., The impact of race and ethnicity on the treatment of hepatitis C disease. *Current Hepatitis Reports*, **2006**. 5: p. 79-85.
- 59. Costa, L.M., et al., Hepatitis C as a risk factor for diabetes type 2: lack of evidence in a hospital in central-west Brazil. *Brazilian Journal of Infectious Diseases*, **2008**. 12: p. 24-26.
- 60. Oni, A.A., et al., Sero-prevalence of hepatitis C virus amoung patients attending STD clinic in Ibadan, Nigeria. *African Journal of Clinical and Experimental Microbiology*, **2005**.6(1): p. 53-59.
- 61. Kakisingi, C., et al., Immunological, virological, parasitic and biological profile of malaria/HIV co-infection in 18 years old and above patients in Lubumbashi (DR Congo). *Open Access Library Journal*, **2016**. 3(5): p. 1-7.
- 62. Browne, E.N., et al., Malariometric update for the rainforest and savanna of Ashanti region, Ghana. Annals of tropical Medicine & Parasitology, 2000. 94(1): p. 15-22.
- 63. Tay, S.C., et al, The prevalence of malaria among HIV seropositive individuals and the impact of the co-infection on their hemoglobin levels. *Annals of clinical microbiology and antimicrobials*, **2015**.14(1): p. 1-8.
- 64. George, J., et al., Significant depletion of CD4+ T cells occurs in the oral mucosa during Simian Immunodeficiency Virus infection with the infected CD4+ T cell reservoir continuing to persist in the oral mucosa during antiretroviral therapy. *Journal of immunology research*, **2015**.
- 65. Wang, S., et al., Modeling the slow CD4+ T cell decline in HIV-infected individuals. *PLoS computational biology*, **2015**.11(12): p. e1004665.
- 66. Mermin, J., et al., Effect of co-trimoxazole prophylaxis on morbidity, mortality, CD4-cell count, and viral load in HIV infection in rural Uganda. *The Lancet*, **2004**.364(9443): p. 1428-1434.
- 67. Cuadros, D.F., Branscum, A.J., and Crowley, P.H., HIV-malaria co-infection: effects of malaria on the prevalence of HIV in East sub-Saharan Africa. *International journal of epidemiology*, **2011**. 40(4): p. 931-939.
- 68. Avettand-Fenoel, V., et al., Total HIV-1 DNA, a marker of viral reservoir dynamics with clinical implications. *Clinical microbiology reviews*, **2016**. 29(4): p. 859-880.
- Cyrille, N.D., et al., Hepatitis B and C, HIV, Syphilis Seroprevalences and Asymptomatic Carriage of Hemoparasites Among Blood Donors at the Douala General Hospital in Cameroon, Central Africa. *Biomedical Journal of Scientific & Technical Research*, 2019.18(5): p. 13968-13974.