**Original Research Article** 

# Awareness and prevalence of hepatitis C virus infection among pregnant women in Nigeria: A national pilot cross-sectional study

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#### Abstract

**Background:** There are no national data on hepatitis C virus awareness and burden among pregnant women to justify its routine screening.

**Objectives:** To investigate awareness, seroprevalence and risk factors for hepatitis C virus infection among pregnant women in Nigeria.

**Methods:** A total of 159 pregnant women from antenatal clinics across six geopolitical zones in Nigeria consented to anti-hepatitis C virus testing which was confirmed using polymerase chain reaction technique. Confirmed hepatitis C virus positive women were further tested for hepatitis B and HIV. Participants were evaluated for risk factors for hepatitis C virus. Odds ratios, adjusted odds ratios, and their 95% confidence intervals (CIs) were determined, and p-values of <0.05 were considered significant.

**Results:** Of 159 participants, 77 (48.4%; 95% confidence interval = 38.2%-60.5%) were aware of hepatitis C virus infection and awareness of hepatitis C virus was associated with young age (odds ratio = 2.21; 95% confidence interval = 1.16-4.21), high educational level (odds ratio = 3.29; 95% confidence interval = 1.63-6.64), and participants' occupation (odds ratio = 0.51; 95% confidence interval = 0.26-0.99). In multivariable logistic regression, adjusted for confounders, the association between awareness of hepatitis C virus and participants' young age (adjusted odds ratio = 1.60; 95% confidence interval = 1.09-2.35; p = 0.018) and high educational level (adjusted odds ratio = 1.48; 95% confidence interval = 1.17-1.86; p = 0.001) remained significant. Hepatitis C virus seroprevalence was found to be 1.3% (95% confidence interval = 0.2%-4.5%). All (100.0%, 95% confidence interval = 12.1%-100.0%) the hepatitis C virus-positive participants and 99 (63.1%, 95% confidence interval = 51.3%-76.8%) hepatitis C virus-negative participants had identifiable hepatitis C virus risk factors. Dual seropositivity of anti-hepatitis C virus/anti-HIV and anti-hepatitis C virus/hepatitis B surface antigen each accounted for 0.6%. The most identified risk factors were multiple sexual partners (15.7%), shared needles (13.8%), and blood transfusion (11.3%). There was no significant association between the risk factors and hepatitis C virus positive status.

**Conclusion:** Awareness of hepatitis C virus infection among pregnant women in Nigeria is low and those aware are positively influenced by young age and high educational level. The prevalence of hepatitis C virus infection is high and provides preliminary evidence to justify antenatal routine screening.

#### Keywords

awareness, hepatitis C virus, pregnancy, prevalence, risk factor, screening

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## Introduction

Hepatitis C virus (HCV) may be one of the common infections among pregnant women in Nigeria, and worldwide approximately 8% of pregnant women have HCV infection, with the prevalence being as high as 4% in the United States.<sup>1</sup> According to the US Centers for Disease Control and Prevention, the estimated 23,000–46,000 children in the United States live with HCV infection and values are expectedly more in East Asia and sub-Saharan Africa.<sup>2</sup> Although there is regional variability in the prevalence of HCV infection in Nigeria, the national data on its awareness and burden among pregnant women are lacking.<sup>2</sup>

Since there is no guideline for routine screening of HCV and health promotional strategies during antenatal care in Nigeria, some pregnant women may not be aware of their HCV infection status, as HCV may not be mentioned by their healthcare providers.<sup>2</sup> In addition, HCV

infection is usually asymptomatic, so, most infected women may be unaware of their status until the incidental diagnosis of chronic HCV is made.<sup>3</sup>

Although universal antenatal screening for HCV is largely debatable, prenatal diagnosis of HCV has a twofold benefit for mother and child.<sup>3</sup> Prenatal HCV screening helps to diagnose an unacknowledged infection by the mother. Furthermore, screening during pregnancy discloses HCV exposure status of the newborn, and provides opportunity for management strategies to mother-to-child transmission (MTCT) and other adverse fetal outcomes potentially associated with maternal HCV infection. According to a recent review, the rate of MTCT of HCV is approximately 5% and perinatally infected children develop cirrhosis at an earlier age than those who acquire HCV as adolescents.<sup>3</sup> More so, HCV infection during pregnancy is associated with an increased risk of adverse fetal outcomes such as fetal growth restriction and low birth weight.<sup>3</sup> Therefore, there is need to intensify efforts at prevention of mother-to-child transmission (PMTCT) of HCV.3,4

Eradication of the virus in pregnant women and women of childbearing age is the main target in the prevention and control of HCV infection. Surprisingly, not all hospitals in Nigeria offer prenatal screening for HCV and the awareness of the HCV infection among pregnant women has not been assessed nationally. Awareness of the risks associated with HCV infection during pregnancy will create more demands for HCV screening and PMTCT interventions among them. Some authorities have argued that routine antenatal screening for HCV is not a viable option in resource-constrained settings because the information on the efficacy and safety of antivirals for HCV in pregnancy is generally lacking, and the treatment of HCV infection during pregnancy is not currently recommended.<sup>4,5</sup> Nevertheless, intrauterine and peripartum transmission of HCV is both possible and higher rates are associated with a high maternal serum viral load and concomitant HIV or hepatitis B virus (HBV) infection, prolonged or difficult delivery, and invasive fetal monitoring during delivery. Therefore, infection during pregnancy and infancy needs to be investigated more to design management strategies most effectively.<sup>6</sup> This may help impact pregnancy care decisions and may involve limiting obstetrical practices that increase fetal exposure to maternal blood such as avoiding the prolonged rupture of membranes, invasive fetal monitoring, and episiotomy (in women who screen positive for HCV).<sup>7,8</sup> In addition, knowledge of HCV infection during pregnancy may inform counseling about associated pregnancy risks such as cholestasis of pregnancy and preterm birth.7

HCV prevalence in the maternal population mirrors that of the general population and obstetrics data could also help healthcare planners and caregivers to design and implement evidence-based interventions for HCV elimination.<sup>9</sup> The World Health Organization has called for HCV elimination as one of the major public health threats by 2030.<sup>4</sup> Accordingly, the US Preventive Services Task Force and the American Association for the Study of Liver Disease-Infectious Diseases Society of America 2019 guidelines recommend universal HCV screening for all individuals aged 18 years and older including pregnant women.<sup>4</sup> Routine universal screening for HCV in pregnancy at the first prenatal visit is therefore recommended so as to institute other nonpharmacological interventions.<sup>10</sup> More so, HCV screening will promote Sustainable Development Goal 3, and general wellness for both mother and child.

There are currently no national data on the awareness and prevalence of HCV infection among pregnant women in Nigeria. This study was therefore undertaken to assess the awareness, seroprevalence, and potential risk factors of HCV infections within the Nigerian national obstetric population.

## **Materials and methods**

#### Study design

This was a multicenter national hospital-based cross-sectional pilot baseline study designed to explore the HCV awareness, the prevalence of HCV and factors affecting both awareness, and prevalence rates of HCV among pregnant women.

## Study area

The study was conducted in Nigeria, a country made up of 36 states and the Federal capital territory grouped into six geopolitical zones. Participants were recruited from one randomly selected tertiary level health facility in each of the five geopolitical zones in Nigeria apart from the South East zone where the lead institution for the Tertiary Education Trust Fund (TETFund) National Research Fund 2019 was located. The sites randomly selected are: Aminu Kano Teaching Hospital, Kano (North West zone); Obafemi Awolowo University Teaching Hospital Complex, Ile-Ife (South West zone); University of Port Harcourt Teaching Hospital, Port Harcourt (South South zone); University of Abuja Teaching Hospital, Gwagwalada (North Central zone); and University of Maiduguri Teaching Hospital, Maiduguri (North East zone) in addition to the lead institution, Nnamdi Azikiwe University Teaching Hospital, Nnewi (South East zone). A summary of the protocol is described here, and the complete protocol has previously been published.<sup>11</sup>

#### Study population

The study was carried out among consenting pregnant women, who were attendees of the antenatal clinic in randomly selected healthcare facilities in each of the six geopolitical zones of Nigeria.

#### Inclusion criteria

Pregnant women registered for their antenatal care in the study sites were eligible to participate in the study.

#### Exclusion criteria

Non-pregnant women and women whose pregnancy could not be confirmed by ultrasound or blood test were excluded.

#### Recruitment of participants

We employed random sampling by selecting six tertiary hospitals across the six geopolitical zones in Nigeria. The research assistants were adequately trained in recruiting and screening pregnant women in the survey. Pregnant women were recruited from the antenatal clinics of each hospital over a 1-month (between 2 June 2020 and 1 July 2020) period. After detailed explanation of the objectives, procedures, and possible benefits of the study, only those who accepted to participate in the study and gave a written informed consent were enrolled into the study. The participants were then interviewed using study specific tool. The study tool contained questions asking women whether they had heard of HCV infection, including whether they had specific risk factors for HCV such as past history of HBV, intravenous drug use, and multiple sex partners. Past history of HBV was defined as participants who previously tested positive to hepatitis B surface antigen (HBsAg). Intravenous drug use was defined as injection of chemicals into the body through a hypodermic needle into a vein. A person was defined to have multiple sex partners, when the person had sexual intercourse with more than one person in the past 12 months. Pregnant women who completed the survey were asked for their contact details and given a sole identifier; a research team member crosschecked these details to ensure each woman finished the survey questions. Thereafter, blood sample was collected for HCV determination.

#### Laboratory procedure and analysis

Five milliliters of the blood sample was aseptically collected by venipuncture from each pregnant woman into a plain specimen bottle after consent was sought and obtained. The participant's code, age, time, and date of collection were labeled on the bottle for proper identification. Each of the samples was centrifuged at 3000 r/min for 5 min and the serum portion was used on the test strip for antigen or antibody detection. Sera samples were stored at  $-25^{\circ}$ C in line with the kit manufacturer's instruction until screened for HBsAg, anti-HIV, and anti-HCV antibodies. Screening for anti-HCV antibodies was conducted using the enzyme-linked immunosorbent assay (ELISA) kit manufactured by LabACON (Hangzhou Biotest Biotech Company, Ltd., China) which has a specificity of 99.0% and a sensitivity of 99.9% according to the manufacturer's declared figures. The kit has in-built controls. The manufacturer's instruction was strictly followed and executed by trained research assistants in each facility. The results were reported as positive or negative. The anti-HCV antibodies tests were confirmed using RNA polymerase chain reaction (PCR) analyzed centrally at Molecular Virology Laboratory, NAUTH, Nnewi, Nigeria. HCV-positive was defined specifically as anti-HCV positive and HCV RNA detected. Serial rapid HIV testing was done according to the Nigerian National HIV testing guidelines, namely, Alere Determine HIV-1/2 (Alere Medical Co., Ltd., Matsudo, Japan) test kit as a screening test, followed by the Uni-Gold Recombigen® HIV-1/2 (Trinity Biotech, Ireland) assay if positive and finally confirmed by HIV-1/2 STAT-PAK (Chembio Diagnostic Systems, Inc., USA). The HBsAg was tested using an ELISA kit manufactured by the LabACON (Hangzhou Biotest Biotech Company, Ltd.).

#### Study endpoints

The primary endpoint was the awareness and seroprevalence of HCV. The other measures of interest included patient demographics, risk factors for HCV, and medical history.

## Sample size determination

No formal sample size calculations were made because of the preliminary nature of the study.<sup>12</sup>

#### Statistical analysis

The data were entered into an Excel 2016 spreadsheet (Microsoft Corporation, Redmond, WA, USA) and subsequently was imported into Statistical Package for the Social Sciences (SPSS) Version 22.0 (IBM Corp., Armonk, NY, USA). Statistical analyses were performed with SPSS Version 22.0 (IBM Corp.). The bivariate analysis was performed using a Pearson's chi-square test or Fisher's exact test, whenever appropriate, to compare the demographic characteristics (e.g. age and marital status) as well as risk factors for HCV (e.g. occupational sex worker and multiple sex partner) and awareness and prevalence rates. Conditional logistic regression was employed in the multiple regression analysis to determine variables associated with awareness of HCV, while controlling for other confounding variables (such as religion, marital status, and ownership of housing). In this analysis, the odds ratio (ORs), adjusted odds ratios (aORs), and confidence interval (CI) set at 95% were determined, and p<0.05 was considered significant.



Figure 1. Flowchart of the study participants.

## Ethical consideration

Written consent was obtained from all participants before recruitment into the study. Ethical clearance for this study was obtained from the National Health Research Ethics Committee, with registration number: NHREC/01/01/2007-23/01/2020 (approval date: 23 January 2020) in accordance with the Helsinki code of conduct for biomedical research involving human subjects. Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) guidance was used for reporting. The full details of the study are in the protocol.<sup>11</sup>

#### Results

During the 1-month pilot study period, 159 participants were enrolled into the study. The data of the 159 participants

were reported. The flowchart is shown in Figure 1. The result of the bivariate analysis of the association between HCV infection awareness and respondents' socio-demographic characteristics is shown in Table 1, while Table 2 shows the association between hepatitis C virus awareness and respondents' socio-demographic characteristics based on bivariate test and multiple logistic regressions.

Of the 159 participants, 77 (48.4%; 95% CI=38.2%– 60.5%) were aware of HCV infection. HCV awareness was found to be associated with participants' age (p=0.018) and educational level (p=0.001), but not with religion, unmarried status, occupation, or residing in a rented apartment. Two participants were screened and confirmed positive for HCV, giving an HCV prevalence of 1.3% (95% CI=0.2%–4.5%). All the two HCV-positive participants (100.0%, 95% CI=12.1%–100.0%) and 99 (63.1%, 95%

Variables/subgroup	Aware of HCV (%) (N=77)ª	Not aware of HCV (%) (N=82)ª	OR (95% CI)	p-value
Age (years)				
21–30	39 (50.6)	26 (31.7)		
31–40	31 (40.3)	39 (47.6)	2.21 (1.16-4.21)	0.016*
41–50	7 (9.1)	17 (20.7)		
Religion				
Christianity	37 (48.1)	43 (52.4)		
Islam	40 (51.9)	39 (47.6)	0.84 (0.45-1.56)	0.580
Marital status			, , , , , , , , , , , , , , , , , , ,	
Married	66 (85.7)	71 (78.3)		
Not married	11 (14.3)	11 (13.7)	0.93 (0.38-2.29)	0.874
Educational level	× ,			
Post-secondary	61 (79.2)	44 (53.7)		
Pre post-secondary	16 (20.8)	38 (46.3)	3.29 (1.63-6.64)	0.001*
Participants' occupation			, ,	
Unemployed	22 (28.6)	36 (43.9)		
Formally employed	32 (41.6)	18 (22.0)	0.51 (0.26-0.98)	0.046*
Farming, trading, artisan (others)	23 (29.9)	28 (34.1)		
Ownership of housing				
Own house	27 (35.1)	28 (34.1)		
Rented apartment	50 (64.9)	54 (65.9)	1.04 (0.54–2.00)	0.903

 Table 1. Association between hepatitis C virus awareness and respondents' socio-demographic characteristics based on bivariate test.

HCV: hepatitis C virus; OR: odds ratios; CI: confidence interval.

Pre post-secondary includes secondary, primary, quoranic, and no education.

<sup>a</sup>Values are given as number (percentage). \*=statistically significant.

 Table 2. Association between hepatitis C awareness and respondents' socio-demographic characteristics based on bivariate test and multiple logistic regression.

Variables/subgroup	Aware of HCV (%) (N=77)ª	Not aware of HCV (%) (N=82)ª	Unadjusted OR (95% CI)	p-value	Adjusted OR (95% CI)	p-value
Age (years)						
21–30	39 (50.6)	26 (31.7)	Reference		Reference	
31–40	31 (40.3)	39 (47.6)	2.21 (1.16-4.21)	0.016*	1.60 (1.16–4.21)	0.018*
41–50	7 (9.1)	17 (20.7)	· · · · ·		· · · · · ·	
Educational level						
Post-secondary	61 (79.2)	44 (53.7)	Reference		Reference	
Pre post-secondary	16 (20.8)	38 (46.3)	3.29 (1.63-6.64)	0.001*	1.48 (1.17–1.86)	0.001*
Participant's occupation			· · · · ·		· · · · · ·	
Unemployed	22 (28.6)	36 (43.9)	Reference		Reference	
Formally employed	32 (41.6)	18 (22.0)	0.51 (0.26-0.98)	0.046*	0.65 (0.42-1.00)	0.051
Farming, trading, artisan (others)	23 (29.9)	28 (34.1)	```			

HCV: hepatitis C virus; CI: confidence interval; OR: odds ratio.

Pre post-secondary include secondary, primary, quoranic, and no education.

Conditional logistic regression was employed (p < 0.1) in the multiple regression analysis to control confounding variables: religion, marital status, and ownership of housing.

<sup>a</sup>Values are given as number (percentage). \*=statistically significant.

CI=51.3%–76.8%) HCV-negative participants had identifiable HCV risk factors (p > 0.05). Dual seropositivity of anti-HCV/anti-HIV and anti-HCV/HBsAg each accounted for 0.6% and none of the participants was triply infected or mono-infected with HCV. Table 3 shows the association between awareness of HCV and respondents' risk factors for HCV infection. The commonest identified risk factors included multiple sexual partners, 25 (15.7%), shared needles, 22 (13.8%), and blood transfusion, 18 (11.3%).

Risk factor	Aware of HCV (%) (N=77) <sup>a</sup>	Not aware of HCV (%) (N=82) <sup>a</sup>	OR (95% CI)	p-value
IV drug use	7 (9.1)	5 (6.1)	1.54 (0.47–5.07)	0.478
Occupational sex worker	0 (0.0)	I (I.2)	0.35 (0.01-8.74)	0.523
Multiple sex partner	9 (11.7)	16 (19.5)	0.55 (0.23–1.32)	0.180
Non-use of condom outside sex partner	7 (9.1)	3 (3.7)	2.63 (0.66-10.58)	0.172
Sexual contact of case of HIV, HBV, or HCV	1 (1.3)	3 (3.7)	0.35 (0.04–3.40)	0.363
Shared needles	11 (14.3)	11 (13.4)	1.08 (0.44–2.65)	0.874
Tattoo	2 (2.6)	3 (3.7)	0.70 (0.11–4.32)	0.703
Past history of HBV	3 (3.9)	1 (1.2)	3.28 (0.33–32.27)	0.308
History of blood transfusion	9 (11.7)	9 (10.9)	1.07 (0.40–2.86)	0.887
No risk factor	28 (36.3)	30 (36.6)	0.99 (0.52–1.89)	0.977

Table 3. Association between awareness of hepatitis C virus infection and respondents' risk factors for HCV based on bivariate test.

HCV: hepatitis C virus; OR: odds ratios; CI: confidence interval; HBV: hepatitis B virus; IV: intravenous. <sup>a</sup>Values are given as number (percentage).

Tabl	e 4.	Association	between	hepatitis	C vir	us status	and	respond	ents	risk	factors	for	HC	V
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Risk factor	HCV positive (%) (N=2) <sup>a</sup>	HCV negative (%) (N=157) <sup>a</sup>	OR (95% CI)	p-value
IV drug use	0 (0.0)	12 (7.6)	2.33 (0.11–51.21)	0.592
Occupational sex worker	0 (0.0)	I (0.7)	20.87 (0.67-648.11)	0.060
Multiple sex partner	I (50.0)	24 (15.3)	5.54 (0.34–91.65)	0.231
Non-use of condom outside wedding partner	0 (0.0)	10 (6.4)	2.81 (0.13-62.38)	0.514
Sexual contact of case of HIV, HBV, or HCV	0 (0.0)	4 (2.5)	6.82 (0.28-163.70)	0.236
Shared needles	I (50.0)	21 (13.4)	6.47 (0.39-107.53)	0.193
Tattoo	0 (0.0)	5 (3.2)	5.54 (0.24-129.84)	0.249
Past history of hepatitis B	0 (0.0)	4 (2.5)	6.82 (0.28-163.70)	0.236
History of blood transfusion	0 (0.0)	18 (11.5)	1.51 (0.07-32.65)	0.793
No risk factor	0 (0.0)	58 (36.9)	0.34 (0.02–7.21)	0.488

HCV: hepatitis C virus; OR: odds ratios; CI: confidence interval; HBV: hepatitis B virus; IV: intravenous.

<sup>a</sup>Values are given as number (percentage).

The association between HCV status and respondents' risk factors for HCV is shown in Table 4. The risk factors variables (histories of multiple sexual partnership (p=0.231), shared needle (p=0.193), blood transfusion (p=0.793), etc.) did not have significant association with HCV-positive status.

## Discussion

This study reveals the high awareness gaps and burden of HCV infection among pregnant women in Nigeria. In this study, only 48.4% of the participants were aware of HCV infection and such awareness was significantly influenced by young age (p=0.018) and high educational level (p=0.001). The prevalence of HCV infection among pregnant women was 1.3%, and all the HCV-positive participants and 63.1% of HCV-negative participants had identifiable HCV risk factors. All the HCV-positive cases were either dually infected with HIV or HBsAg. None of the participants was triply infected for HCV, HBV, and HIV or mono-infected with HCV.

Our findings identified awareness gaps for HCV infection among pregnant women in Nigeria with less than 50% being aware of HCV infection. This was in contrast with findings from a study in the United States conducted among pregnant women with opioid use disorder in which 97% were aware of HCV.<sup>13</sup> The high awareness gaps among this study's participants might be due to difference in study populations. While this study involved all pregnant women attending antenatal clinics, the quoted study from the United States<sup>13</sup> involved only pregnant women with opioid use disorder who may have been previously reached with messages regarding HCV by their healthcare providers in the course of discussing the implications of opioid use and infections that could be transmitted by intravenous drug users. The differences in the epidemiology of HCV in the study settings may be an additional reason for the observed differences in awareness levels. The low awareness of hepatitis C virus infection among the study participants is worrisome and calls for need for more health education on hepatitis C virus and other infections among pregnant women in Nigeria. The antenatal care provides a big opportunity to offer pregnant women health education including hepatitis C infection. The finding from this study makes the case for provider counseling and education at the time of antenatal booking and care for pregnancy.

This study has also revealed that the level of awareness was positively influenced by young age (p=0.018) and high educational level (p=0.001). These findings are understandable because younger women and women with high educational levels may be more likely to seek health-care-related information. In a recent Ethiopian study, young age (17–25 years) was a significant predictor of HCV infection,<sup>14</sup> while a previous report<sup>15</sup> from the same country revealed high HCV status among pregnant women with higher education level.

The seroprevalence of anti-HCV antibodies of 1.3% was higher than 0.5% reported previously in Anyigba, Nigeria, by Omatola et al.<sup>16</sup> and 0.3% reported in India, by Malhotra et al.<sup>17</sup> in an antenatal population in India, but similar to 1.6% reported in a recent Ethiopian study by Dagnew et al.<sup>14</sup> However, this rate was lower than 3.9% reported by Okusanya et al. in Irrua, Nigeria,<sup>18</sup> as well as findings of other epidemiological studies that revealed a prevalence of 2%–4%.<sup>16,19,20</sup> Our findings were also lower than 8.07% found in a study in Ethiopia.<sup>17</sup> In a systematic review involving 26 studies conducted by the Centers for Disease Control and Prevention (CDC) on HCV infection during pregnancy, the HCV-positive prevalence was 1.2%.<sup>21</sup>

As revealed in this study, HCV prevalence has no statistically significant association with any of the potential risk factors for HCV even though it was higher among participants with histories of multiple sexual partners and shared needles. This finding is analogous to a previous Nigerian study by Okusanya et al.<sup>18</sup> as well as an Ethiopian report by Dabsu and Ejeta.<sup>15</sup> Moreover, since both HIV and HCV or HBV and HCV share common risk factors and mode of transmission, it will be equally important to incorporate information on HCV risk factors into HIV/AIDS and HBV intervention strategies in Nigeria as an alternative. However, to drastically reduce the prevalence of this deadly disease in Nigeria, a nationwide health promotional campaign should be done to help create awareness regarding HCV infection.

Unlike findings from a previous study in Keffi, Nigeria, by Oti et al.<sup>22</sup> where no pregnant woman was co-infected with HCV and HBV, our report revealed that no woman was mono-infected with HCV and 0.6% had HCV/HBV or HCV/HIV. Our overall HCV–HIV co-infection of 0.6% was lower than 1.5% reported by Ezechi et al.<sup>23</sup> in Lagos, Nigeria, among HIV-positive pregnant population and 2.9% reported in Ethiopia.<sup>15</sup>

The Society of Gynecology and Obstetrics of Nigeria (SOGON) has not advised for routine screening of pregnant women for HCV infection in Nigeria. However, in 2018, universal hepatitis C virus screening during pregnancy was recommended by the American Association for the Study of Liver Diseases and the Infectious Diseases Society of America. Since that time, the CDC United States Preventive Task Force (USPSTF) have updated their previous guidance and specifically recommended screening for all pregnant women during each pregnancy, except in settings where the prevalence of HCV infection is <0.1%.<sup>21</sup> By extension, this guideline should apply to Nigerian settings since the prevalence of HCV infection among pregnant women in Nigeria in this study is 1.3%. These guidelines all acknowledge the cost-effectiveness of universal screening and the inadequacies of risk-based screening. However, in the absence of endorsement of these guidelines from the obstetrics societies, screening practices will likely remain non-uniform across obstetrics practice units in Nigeria.<sup>7</sup> Again, there are no agencies/groups in Nigeria that are recommending universal HCV screening.

The clinical importance of this study was that pregnancy is one of the few points of contact women of reproductive age have with their healthcare providers; therefore, pregnancy provides a crucial time for targeting this population for HCV screening. Children also benefit from maternal screening, because the primary route of infection in children is vertical transmission during pregnancy, and children are not routinely assessed for liver disease.<sup>1</sup> The lack of disease awareness certainly can contribute to lack of awareness of risk behaviors and preventive strategies to mitigate risk, as well as the need of diagnostic testing. Universal screening offers several advantages that position us for a future where HCV treatment in pregnancy can happen and offers us progress toward the elimination of HCV.<sup>24</sup> Our experience underscores the importance of obtaining information about at-risk behavior. Although universal HCV screening will identify patients who fail to acknowledge risk behaviors, discussing HCV and the medical complications associated with acquiring HCV in pregnancy and its MTCT should also be part of prenatal counseling. Future research should evaluate the impact of this prenatal screening guideline on clinical practice.

The strength of this study is the multicenter study site design and appears to be the first national study in Nigeria to evaluate the HCV awareness rate among pregnant women population to the best of the authors knowledge. In addition, studies have shown that after using a screening assay, a confirmatory test to confirm anti-HCV status was needed; this was provided by the HCV RNA testing carried out to confirm the HCV infection in this study.<sup>1,3</sup> Since the study design was cross-sectional and the data were preliminary, it shares the shortcoming of constructing cause and effect relationship. There are some limitations. For instance, although our present protocol included determination of rates of seroconversion for these viral infections in pregnancy, we could not evaluate for seroconversion or birth outcomes in our present preliminary findings, which is an important reminder that women can remain at risk for new HCV infections even during pregnancy.<sup>11</sup> Awareness of HCV could be quite challenging in some settings where people may confuse HCV, hepatitis A virus (HAV), and HBV. However, we avoided this confusion by interpreting the questionnaires in vernacular language. This study

appears inadequately powered to demonstrate an association between risk factors for HCV and HCV-positive status. If this association had been demonstrated, it could be used to argue that risk-based screening is preferable to universal screening.

## Conclusion

The awareness of HCV infection among pregnant women in Nigeria is less than 50%, and it is positively influenced by younger age and higher educational level. HCV prevalence rate was 1.3% although it appears inadequately powered to demonstrate an association between risk factors for HCV and HCV-positive status. The high prevalence of HCV provides preliminary evidence that routine screening may be a viable option. However, the results of the completed survey with a larger study sample would provide more clarity.

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#### **Author contributions**

G.U.E., I.I.M., A.R., G.O.A., P.O.F., O.M.L., H.A.U., C.U.O., U.C.O., and S.O.K. contributed to the study conceptualization and methodology; R.O.E., C.H.J., P.O.A., C.P.C., H.S.I., F.E.A., E.O.I., B.A.A., A.I.N., O.D.O., S.A.O., I.C.O., U.I.A., S.O.I., and A.A. conducted the clinic study, ensured completion of the participants data, and documented the required data; C.H.N. and G.U.E. analyzed the data and drafted the original manuscript; C.E.U., S.I.N., S.N.C., O.S.U., R.C.C., E.O.U., E.A.E., C.C.O., I.K.N., A.A.O., E.P.I., and C.N.O. worked on formal analysis; N.N.J.-I., I.C.A., O.C.E., I.A.Y., and J.I.I. contributed to the project administration, writing (review and editing), data visualization, and supervision. All authors have seen and approved their contributions and the final version of the manuscript.

#### Data availability

The data used to support the findings of this study are available from the site publicly.

#### **Declaration of conflicting interests**

The author(s) declared no potential conflicts of interest with respect to the research, authorship, and/or publication of this article.

#### **Disclosure statement for publication**

All authors have made substantial contributions to: conception and design of the study, or acquisition of data, or analysis and interpretation of data; drafting the article or revising it critically for important intellectual content; and final approval of the version submitted. This manuscript has not been submitted for publication in another journal.

#### Ethical approval and consent to participate

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#### References

- Saab S, Kullar R and Gounder P. The urgent need for hepatitis C screening in pregnant women: a call to action. *Obstet Gynecol* 2020; 135(4): 773–777.
- Bigna JJ, Kenne AM, Hamroun A, et al. Gender development and hepatitis B and C infections among pregnant women in Africa: a systematic review and meta-analysis. *Infect Dis Poverty* 2019; 8(1): 16.
- Ragusa R, Corsaro LS, Frazzetto E, et al. Hepatitis C virus infection in children and pregnant women: an updated review of the literature on screening and treatments. *AJP Rep* 2020; 10(1): e121–e127.
- Freriksen JJM, van Seyen M, Judd A, et al. Review article: direct—acting antivirals for the treatment of HCV during pregnancy and lactation—implications for maternal dosing, foetal exposure, and safety for mother and child. *Aliment Pharmacol Ther* 2019; 50(7): 738–750.
- Hong J, Kushner T, Dieterich D, et al. Reducing mother-tochild transmission of HCV: Is it attainable with a multidisciplinary approach. *J Hepatol* 2019; 71(1): 229–230.
- El-Shabrawi MHF, Kamal NM, Mogahed EA, et al. Perinatal transmission of hepatitis C virus: an update. *Arch Med Sci* 2019; 16(6): 1360–1369.
- Kushner T, Park C, Masand D, et al. Hepatitis C seroprevalence among consecutive labor and delivery admissions in two New York City Hospitals. *Open Forum Infect Dis* 2020; 7(11): ofaa514.
- US Preventive Services Task Force; Owens DK, Davidson KW, et al. Screening for Hepatitis C Virus Infection in Adolescents and Adults: US Preventive Services Task Force Recommendation Statement. JAMA 2020; 323: 970–975.
- Sheffield JS and Argani CH. Hepatitis C virus infection in the pregnant woman: is it time for universal screening. *Obstet Gynecol* 2020; 135(4): 770–772.

- Spera AM, Eldin TK, Tosone G, et al. Antiviral therapy for hepatitis C: has anything changed for pregnant/lactating women? *World J Hepatol* 2016; 8(12): 557–565.
- 11. Eleje GU, Mbachu II, Ogwaluonye UC, et al. Prevalence, seroconversion and mother-to-child transmission of dual and triplex infections of HIV, hepatitis B and C viruses among pregnant women in Nigeria: study protocol. *Reprod Health* 2020; 17(1): 144.
- Noordzij M, Tripepi G, Dekker FW, et al. Sample size calculations: basic principles and common pitfalls. *Nephrol Dial Transplant* 2010; 25(5): 1388–1393.
- Krans EE, Rothenberger SD, Morrison PK, et al. Hepatitis C virus knowledge among pregnant women with opioid use disorder. *Matern Child Health J* 2018; 22(8): 1208–1216.
- Dagnew M, Million Y, Gizachew M, et al. Hepatitis B and C viruses' infection and associated factors among pregnant women attending antenatal care in hospitals in the Amhara National Regional State, Ethiopia. *Int J Microbiol* 2020; 2020: 8848561.
- Dabsu R and Ejeta E. Seroepidemiology of hepatitis B and C virus infections among pregnant women attending antenatal clinic in selected health facilities in East Wollega Zone, West Oromia, Ethiopia. *Biomed Res Int* 2018; 2018: 4792584.
- Omatola CA, Okolo MO and Abraham JO. Hepatitis C virus coinfection in human immunodeficiency virus infected pregnant women in Anyigba, Kogi State, Nigeria. *Nat Sci* 2018; 16: 62–68.
- 17. Malhotra P, Nanda S, Malhotra V, et al. Prevalence of HIV, hepatitis B, hepatitis C in pregnancy at tertiary care center

of Northern India. *Adv Res Gastroentero Hepatol* 2016; 1(4): 80–82.

- Okusanya BO, Aigere EO, Eigbefoh JO, et al. Seroprevalence and clinico-epidemiological correlates of hepatitis C viral antibodies at an antenatal booking clinic of a tertiary hospital in Nigeria. *Arch Gynecol Obstet* 2013; 288(3): 495–500.
- Kabinda JM, Akilimali TS, Miyanga AS, et al. Hepatitis B, hepatitis C and HIV in pregnant women in the community in the Democratic Republic of Congo. *World J AIDS* 2015; 5(2): 124–130.
- Opaleye OO, Igboama MC, Ojo JA, et al. Seroprevalence of HIV, HBV, HCV, and HTLV among Pregnant Women in Southwestern Nigeria. *J Immunoassay Immunochem* 2016; 37(1): 29–42.
- Schillie S, Wester C, Osborne M, et al. CDC recommendations for hepatitis C screening among adults—United States, 2020. MMWR Recomm Rep 2020; 69(2): 1–17.
- Oti BV, Pennap GR and Ngari HR. HBsAg and anti-HCV prevalence among pregnant women accessing antenatal care in a tertiary healthcare facility in Central Nigeria. *Hepatol Pancreat Sci* 2018; 2(1): 1–4.
- Ezechi OC, Kalejaiye OO, Gab-Okafor CV, et al. Seroprevalence and factors associated with Hepatitis B and C co-infection in pregnant Nigerian women living with HIV infection. *Pan Afr Med J* 2014; 17: 197.
- Jhaveri R, Broder T, Bhattacharya D, et al. Universal screening of pregnant women for hepatitis C: the time is now. *Clin Infect Dis* 2018; 67(10): 1493–1497.