

## Sexual behavioral correlates with HSV-2 seroprevalence among pregnant women in Nigeria

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### Abstract

**Introduction:** The burden of HSV-2 infection, the cause of most cases of genital herpes in Nigeria, varies from region to region; and so are the associated factors. This infection is known to be responsible for several negative pregnancy outcomes. There is currently no documented data on sexual behavioral factors associated with the occurrence of HSV-2 infection or seroprevalence among pregnant women in Nigeria. This study aimed at identifying the sexual behavioral correlates of HSV-2 seroprevalence among pregnant women in Benin City, Nigeria.

**Methodology:** The cross-sectional study design was adopted and the study took place between November 2011 and June 2012. Four hundred and ten consenting ante-natal clinic patients in two major tertiary hospitals in Benin City were consecutively and prospectively included. Data sources were represented by questionnaires, the patient's case records and laboratory investigations. Each patient's serum was analyzed for HSV-2 antibody detection. Data analysis was performed using SPSS version 16.

**Results:** Four hundred and ten patients were enrolled with average age 30.6 years. Seroprevalence of HSV-2 antibody was 47.3%. Sexual behavioral factors that were significantly associated with HSV-2 seroprevalence included early exposure to sexual intercourse, number of sex partners, involvement in polygamous marriages, involvement of husband in extra-marital affairs and hormonal contraceptive use.

**Conclusion:** The prevalence of HSV-2 among pregnant women in Benin City is high. Public health campaigns aimed at: delaying onset of sexual activity; encouraging monogamous relationships; and emphasizing that hormonal contraceptives do not protect from STI's, are recommended.

**Key words:** Herpes simplex Virus 2; seroprevalence; sexually transmitted infections; STI's risk factors; pregnancy.

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### Introduction

Genital herpes, an infection commonly caused by human herpes simplex virus 2 (HSV-2) in Nigeria and in most regions of the world, has been described as a silent pandemic with different countries being at different stages of the epidemic [1,2]. Humans are the only natural host [3]. The fact that transmission through viral shedding continues, also in the frequent asymptomatic cases, has sustained the worsening of the pandemic [4]. The HSV-2 pandemic is further reinforced by the HIV pandemic and *vice versa*; and it is becoming clear that the efforts at controlling the spread of HIV may remain ineffective if control of HSV-2 infection, along with other sexually transmitted infections (STI) is not integrated [4,5]. Furthermore, HSV-2 infection has been reported to produce a wide range of deleterious obstetric effects, ranging from

neonatal herpes, neonatal death, neuronal deficits, mental handicap, congenital malformations, intrauterine fetal death, preterm labour, low birth weight babies, and spontaneous abortions [6,7].

Since the HSV-2 infection is life-long and has no known cure, primary prevention remains the mainstay of its control. Hence it is important to understand the factors that are associated with the occurrence of infection [8]. There is yet no documented behavioural correlate of this infection or seroprevalence in Nigeria. This study is aimed at determining the behavioral correlates of HSV-2 seroprevalence among pregnant women in Benin, Nigeria.

## Methodology

### *Study location, design and duration*

Participants were ante-natal clinic patients of the two major tertiary hospitals in Benin City, Nigeria: University of Benin Teaching Hospital and Central Hospital, Benin-City. The cross-sectional study took place between November 2011 and June 2012. Participants were consecutively and prospectively recruited.

### *Ethical issues*

Informed written consents were sought and consecutively consenting pregnant women were recruited into the study. The study obtained ethical approval (ADM/E22/A/VOL.VII/740) from the Ethical committee of the University of Benin Teaching Hospital.

### *Inclusion and exclusion criteria*

Consenting pregnant women attending antenatal clinic in both University of Benin Hospital and Central Hospital between November 2011 and June 2012 were included in the study. Exclusion criteria were absence of pregnancy and failure to sign the consent form. No clinical diagnostic criteria were used in the study.

### *Data collection*

Sources of data were represented by questionnaires, hospital records, and laboratory investigations. Blood samples were obtained from participants and the serum was separated and used for the HSV-2 IgG antibody assay. A structured questionnaire was used to collect data on specific participants' sociodemographic and behavioural profiles.

### *Laboratory procedures*

Blood samples were collected in 5 ml plain vacutainer tubes and allowed to clot and sera separated by centrifugation at room temperature. Samples were stored in cryovials at  $-20^{\circ}\text{C}$ . The HSV-2 IgG assay procedure was utilized the enzyme-linked immunosorbent assay (ELISA) kit (Dia. Pro. Diagnostic Bioprobes srl, Milano, Italy). This is a glycoprotein G-based type-specific ELISA technique and test result was qualitative. All specimens and kit reagents were brought to room temperature and gently mixed. Procedures were performed in accordance with manufacturer's instructions. Quality control and test validation was included into the test protocols.

### *Data analysis*

Collected data was analyzed using the SPSS version 16.0 software. Chi Square was used to test associations. Statistical significance was ascribed based on  $p$  values  $< 0.05$ .

## Results

### *Patients general and obstetric characteristics*

The mean age of the 410 pregnant women studied was  $30.6 \pm 5.2$  years (range, 18 – 44 years). The majority of them (96.6%) were in the 21-40 years age bracket. Most of the participants (85.1%) were married and belonged to the Christian faith (81.7%) as shown in Table 1. A total of 382 (93.4%) of the study population completed at least secondary education, while the remaining 16.4% either did not have secondary education at all or had an incomplete secondary education (Table 1).

Most of the respondents (43.4%) were nulliparous, 28.3% primiparous; while a total of 27.5% of them were multiparous. Only two (0.7%) of the participants were grand-multiparae (Table 2). Regarding the gestational age at which participants were recruited into the study, 45.3%, 39.8% and 14.6% of the women were at the third, second and first trimesters respectively as shown in Table 2.

Of the 410 pregnant women studied 194 tested positive for HSV-2 antibody giving a prevalence rate of 47.3% (Table 3). The socio-demographic characteristics of the seropositive women were essentially similar to those of the study population. Their ages ranged from 16-45 years with a mean of  $32.4 \pm 4.9$  years. The majority of seropositive patients (184/194; 94.8%) fell into the 21-40 years age bracket, were married (161/194; 83.0%) and of Christian faith (164/194; 84.5%). Also 94.8% (184/194) of seropositive women had completed at least secondary education. Their obstetric characteristics are also similar to that of the study population with the majority comprising of nulliparous women (95/194; 49.0%) and women in their third trimester (91/194; 46.9%).

### *Seroprevalence and behavioral characteristics*

The majority of participants, 371 (90.5%), had only one sexual partner during the previous 12 months. The prevalence of HSV-2 was lowest in this group (45.8%) when compared to those with two or three recent partners (59.4% and 71.4% respectively).

**Table 1.** General characteristics of respondents

Characteristics	Frequency	Percentage %
<b>Age group (years)</b>		
15-20	4	1.0
21-25	62	15.1
26-30	146	35.6
31-35	135	32.9
36-40	53	12.9
41-45	10	2.4
<b>Religion</b>		
Christianity	335	81.7
Islam	75	18.3
<b>Marital status</b>		
Married	349	85.1
Single	32	7.8
Divorced	13	3.2
Widowed	16	3.9
<b>Level of Education</b>		
Graduate & Above	129	31.5
Post secondary	175	42.7
Secondary completed	78	19.2
Secondary uncompleted	9	2.2
Primary completed	17	4.1
Primary uncompleted	2	0.5

**Table 2.** Obstetric characteristics of respondents

Obstetric characteristics	Frequency	%
<b>Trimester</b>		
1 <sup>st</sup> Trimester	60	14.6
2 <sup>nd</sup> Trimester	163	39.8
3 <sup>rd</sup> Trimester	187	45.3
<b>Parity</b>		
Nullipara	178	43.4
Primipara	116	28.3
Para-2	74	18.0
Para-3	27	6.6
Para-4	12	2.9
Para-5 or more	3	0.7

**Table 3.** Seroprevalence of HSV<sub>2</sub>-antibody among participants

Participants	Frequency (%)
HSV <sub>2</sub> -Ab Positive	194 (47.3)
HSV <sub>2</sub> -Ab Negative	216 (52.7)
Total	410 (100.0)

The observed association between number of sex partners in the previous 12 months and the prevalence of HSV-2 was statistically significant ( $p = 0.023$ ) (Table 4). Table 4 also shows an increase in prevalence of HSV-2 correlated to the number of lifetime sex partners from 33.2% in participants with only one lifetime sex partner to 90.1% in those with five. This observed trend was also statistically significant ( $p = 0.031$ ).

Prevalence of HSV-2 decreased as the age of first sexual exposure increased. This association was also found to be statistically significant ( $p = 0.043$ ) (Table 4). No particular trend in prevalence was followed regarding the total number of years of sexual exposure.

Only a small proportion of the participants reported involvement in polygamous marriages and the involvement of their husbands in extramarital affairs: 25/410 (6.1%) and 64/410 (15.6%)

respectively. Nevertheless, there were statistically significant increases in prevalence of HSV-2 among these categories of participants (Table 4). Although there was an increased HSV-2 prevalence (55%) among participants that practiced sex under the influence of alcohol compared to their counterpart who did not (46.5%), there was no statistical significance ( $p = 0.072$ ). On the contrary, use of hormonal contraceptives for at least more than one year in the past is associated with statistically significant increase in HSV-2 prevalence ( $p = 0.021$ ) (Table 4).

## Discussion

We detected 47.3% HSV-2 seroprevalence rate but since there is no data in our country we could not compare this rate with other studies. This figure however seems close to the values found among high risk groups: 59% among commercial sex workers in

**Table 4.** Behavioural factors and HSV-2 infection

Behavioural Factor	Number tested	HSV-2 Ab Positive (%)	$\chi^2$	P-value
<b>Number of sexual partners in the past one year</b>				
1	371	170 (45.8)	5.69	0.023
2	32	19 (59.4)		
3	7	5 (71.4)		
<b>Number of life time sex partners</b>				
1	199	66 (33.2)	4.57	0.031
2	113	37 (37.6)		
3	60	45 (75.0)		
4	27	23 (85.2)		
5	11	10 (90.1)		
<b>Age at first sexual exposure (years)</b>				
≤ 15	86	59 (68.6)	2.96	0.043
16-20	163	98 (60.1)		
21-25	151	35 (23.2)		
26-30	10	2 (20.0)		
<b>Total number of years of sexual exposure (years)</b>				
≤ 5	183	81 (44.3)	0.01	0.529
6-10	106	49 (46.2)		
11-15	65	36 (55.4)		
16-20	48	25 (52.1)		
>20	8	3 (37.5)		
<b>Involvement in polygamous marriage</b>				
Yes	25	20 (80.0)	11.8	0.001
No	385	174 (45.2)		
<b>Husband's involvement in extramarital affairs.</b>				
Yes	64	39 (60.9)	10.3	0.001
No	346	155 (44.8)		
<b>Sex under influence of alcohol</b>				
Yes	40	22 (55.0)	3.64	0.072
No	370	172 (46.5)		
<b>Use of hormonal contraceptives in the past (&gt; 1 year)</b>				
Yes	270	143 (53.0)	9.45	0.021
No	130	51 (39.2)		

Nigeria [9]; and 56.3% among hotel/bar workers in Tanzania [10]. Although this seroprevalence result falls within the range of 30%-80% found among women of reproductive age in sub-Saharan Africa [11], it is higher than HSV-2 seroprevalences found in several studies among populations of pregnant women in the sub-Saharan Africa region: 20.7% and 33.6% from different locations in Tanzania [12,13]; and 26% in Senegal [14].

This figure is still remarkably higher than findings from studies on pregnant populations outside Africa: 7.5% in India [15], 11.5% in Australia [16], 14.5% in Mexico [17], and 22% in USA [18]. The implication of this finding is that HSV-2 transmission in Benin, Nigeria is efficient; and it indicates that there could be underlying peculiar factors in the environment that enable the sustenance of such a high level of transmission efficiency.

Several sexual behavioral factors were observed, in this study, to be significantly contributory to the high HSV-2 prevalence, in keeping with the assertion that the presence of HSV-2 antibodies is a marker of indiscrete sexual behavior [28, 29]. It is therefore instructive that the following considerations should be included in individual case management and primary prevention strategies: more than one sex partner in the past 12 months; more than one lifetime sex partner; early sexual exposure; involvement in polygamous marriages; and use of hormonal contraceptives.

Early exposure to sexual intercourse has therefore been linked to increased risk of sexually transmitted infections (STIs) during adolescence [30, 31]. The increased STI risk is due, in part, to a biologic predisposition of the immature cervix to infection if exposed [32] and to the increased likelihood of engaging in riskier sexual behaviors among persons who initiate sexual intercourse at younger ages [31, 34]. Although, early sexual exposure as found in this study was significantly associated with HSV-2 seroprevalence, the total number of years of exposure was not significantly associated. This posits that the effect of early sexual exposure derives from the associated loose morals and inefficient parental control during adolescence. Primary preventive strategies, such as encouraging the use of condom, should address this gap in parenting.

In a previous report from Kenya [27], hormonal contraceptive use was found to reduce the risk of HSV-2 and HIV infection; this is contrary to our finding in this study where use of hormonal contraceptives was found to be significantly associated with higher prevalence of HSV-2. Our finding is

however in agreement with another study report from Kenya [35]. Misguided confidence in the protective role of hormonal contraceptives may explain the significantly increased prevalence of HSV-2 among hormonal contraceptive users in this study. There ought to be serious public enlightenment campaign about the limitations of contraceptive use.

Assessment of these sexual behavioral factors in individual case management could help in raising the index of suspicion of HSV-2 seroprevalence and are potentially useful screening criteria, if validated. These factors also ought to be taken into consideration while designing primary prevention strategies for STIs.

## Conclusion

The prevalence of HSV-2 among pregnant women in Benin City, Nigeria, is high. Number of sexual partners in the last 12 months, number of lifetime sexual partners, early sexual exposure and involvement in polygamy are associated sexual behavioral factors. Primary prevention remains the most effective strategy to control HSV-2 infection and associated complications. Public health campaigns aimed at delaying onset of sexual activity and encouraging monogamous relationships are recommended. Also there ought to be serious public enlightenment on the limitations of hormonal contraceptives use for protection against STIs.

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