The prevalence and determinants of female serodiscordance among HIV positive pregnant women attending PMTCT clinic in Owerri, Imo state, Nigeria

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INTRODUCTION

Human Immunodeficiency Virus and Acquired Immunodeficiency Syndrome (HIV/ AIDS) pandemic has remained one of the most challenging health/social issues in recent few decades. According to 2014 National HIV Sero-prevalence Sentinel Survey among Pregnant Women Attending Antenatal Clinics in Nigeria the prevalence of HIV was 3.0%.¹

In Africa, more than 85% of adult HIV infections are due to heterosexual transmission.² In response to growing...
Evidence that married or cohabitating sero-discordant couples are important source of new HIV infections in sub-Saharan Africa, programs are being urged to target prevention efforts to married or cohabitating sero-discordant couples.\footnote{METHODS}{3} HIV sero-negative partners in discordant cohabiting couples (one partner is HIV positive and the other negative) are at a 10\% (or more) annual risk of acquiring HIV infection.\footnote{METHODS}{3} Hence, sero-negative partners in discordant cohabiting relationships are substantially exposed to risk of acquiring HIV.

Nevertheless, what continues to baffle many including medical personnel is the increasing incidence/prevalence of discordant relationship. Some studies especially in Africa have shown that female sero-discordance (where the female partner is positive and male negative) is more prevalent than male sero-discordance (where the male partner is positive and female negative).\footnote{Methods}{4,5} Researchers are yet to come up with an explanation for this situation, although a couple of indicators have been observed for this occurrence.\footnote{Methods}{9,13} Analysis of HIV data in five African countries (Burkina Faso, Ghana, Cameroon, Kenya and Tanzania) showed that extra-marital sexual activity among co-habiting women was a substantial source of vulnerability to HIV infection.\footnote{Methods}{13} This was at odds with levels of self-reported marital infidelity by females and with the common perception that unfaithful male are the main link between high risk group and the general population.\footnote{Methods}{13} According to Sagay et al, socio demographic factors including regional, ethnic and religious settings play significant role in the dynamics of HIV transmission among couples in Nigeria.\footnote{Methods}{5}

Some studies have suggested that male-to-female transmission of HIV is higher than female-to-male transmission.\footnote{Methods}{14,15} The possible causes have been traced to the interrelation of the several factors ranging from transmission rate due to anatomical structure of male and female genitals where men are less vulnerable to contracting the virus than women.\footnote{Methods}{15} However, other studies especially in African settings have reported a very similar transmission rate between the sexes.\footnote{Methods}{10}

Apart from few prevalence studies (non-in FMC Owerri) on female HIV sero-discordance in the South Eastern Nigeria, authors did not come across any studies that assessed the possible socio-demographic and behavioral factors that may be significantly associated with female sero-discordance. This study, therefore, was undertaken to determine the prevalence of female sero-discordance and assess the possible socio-demographic and behavioral factors that may be significantly associated with female sero-discordance among HIV positive pregnant women attending PMTCT clinic in Federal Medical Center Owerri, Imo state Nigeria.

**METHODS**

This was a cross sectional study carried out from December 2015 to May 2016 on consecutively recruited pregnant women (with their partners) who tested positive to HIV I and II at the PMTCT clinic Federal medical Centre (FMC) Owerri.

FMC is a tertiary health institution situated in Owerri South East zone of the country and the capital city of Imo State,. The population was predominantly Igbo and Christians. Federal Medical Center Owerri serves mainly Imo, parts of Anambra, Enugu, Abia, and Rivers states. The FMC Owerri has an established Obstetrics and Gynecology department with four firms. One of the four firms oversee the PMTCT program supported by Center for Clinical Care and Clinical Research of Nigeria (CCCRN) project. The study was approved by the research and ethical committee of FMC Owerri. Informed written consents were obtained from the subjects.

Data were collected using a structured self-administered and validated questionnaire with the assistance of the midwives who were trained in voluntary counseling and testing (VCT). The laboratory investigations were done at the Center for Clinical Care and Clinical Research of Nigeria (CCCRN) laboratory in FMC Owerri. Screening for HIV I and II was carried out by laboratory staff with serial algorithm using Determine and Stat-pak (and Unigold as tie-breaker where necessary).

Those included in the study were guided to fill out questionnaire on bio data, duration of marriage/cohabitation, retroviral status before marriage/cohabitation, sexual behavior (frequency, condom/other barrier use). The participants’ male partners also presented to the clinic to provide information bordering on their (male partner) sexual behaviors. To ensure individual partner’s confidentiality in sensitive information, each partner was interviewed separately. Social classes were assigned to the women using partner’s occupation and educational status of the women according to the scoring system designed by Olusanya et al. for Nigeria and other African countries.\footnote{Methods}{16} The CD4+ Count of the female partner at booking was determined from the laboratory using a power-operated quality-controlled Cy Flow Counter 1 (made in Germany). The use of / and duration on HAART by the female partner was also determined for those already on HAART.

**Statistical analysis**

Data were computed and analyzed using computer software package for social sciences version 20 (SPSS, Inc. USA). Descriptive statistics (mean, standard deviation, percentages etc.) was determined for continuous variables. Pearson Chi square tests, a non-parametric inferential statistic were used to determine significant association for dichotomous variables. Multiple logistic regression analysis was used to identify the factors that were independently associated with female sero-discordance. P-values <0.05 at 95\% confidence interval were considered statistically significant.
RESULTS

A total of 106 HIV positive pregnant women were studied. The mean age was 31.5±4.1 years with a range of 21-42 years. The mean years of cohabitation were 5.1±3.4 years with a range of 1-15 years. Average sexual frequency per week among couples ranges from 1-5 with a mean of 1.67±1.3. The mean parity was 2.0±1.5 with a range of 0-7.

Sixty-seven (67) of these women had sero-negative male partners, while 39 had sero-positive male partners, giving a female sero-discordance prevalence of 63.2% and concordance rate of 36.8%. Of the study population 52.8% (56/106) was lower class, middle class 43.4% (46/106) and upper class 3.8% (4/106;) of the HIV positive pregnant women. Majority of the women (97/106; 91.5%) knew their HIV statuses before marriage/cohabitation. Out of these, fifty-six (56/106; 52.8%) were seropositive for HIV while (41/106; 38.7%) were negative before marriage. Nine (9/106; 8.5%) did not know their status before marriage/cohabitation. Out of the 17 male partners that seroconverted after marriage/cohabitation, 12 (70.6%) were of couple that used condom (usage ranging from “occasionally” to “always except for conception”) for intercourse and 37.3% among those that never used condom (usage ranging from “occasionally” to “always except for conception”) for intercourse while only 5 (29.4%) used condom.

Only (17/84; 20.2%) of the 84 male partners that were sero-negative before marriage were infected or seroconverted after marriage, while 41 of the female seroconverted after marriage/cohabitation. Pre-exposure and/or post-exposure prophylaxes were very uncommon. Only 2 (1.9%) practices it among the participants. Condom use was significantly associated with female sero-discordance (P=0.002). Female sero-discordance rate was 62.7% among couple that used condom (usage ranging from “occasionally” to “always except for conception”) for intercourse and 37.3% among those that never used condom (Figure 1).

Table 1: Participants’ characteristics.

| Variable          | Frequency (%) Male | Female |
|-------------------|-------------------|--------|
| Age group         |                   |        |
| 20-24             | 7 (6.6)           | 0 (0.0) |
| 25-29             | 24 (22.6)         | 11 (10.4) |
| 30-34             | 48 (45.3)         | 22 (20.8) |
| 35-39             | 25 (23.6)         | 53 (50.0) |
| ≥40               | 2 (1.9)           | 20 (18.9) |
| Total             | 106 (100.0)       | 106 (100.0) |
| Education         |                   |        |
| Primary           | 5 (4.7)           | 1 (0.9) |
| Secondary         | 65 (61.3)         | 73 (69.0) |
| Tertiary          | 36 (34.0)         | 32 (30.1) |
| Social class      |                   |        |
| Lower             | 56 (52.8)         |        |
| Middle            | 46 (43.4)         |        |
| Upper             | 4 (3.8)           |        |
| HIV status before marriage |     |        |
| Positive          | 56 (52.8)         | 21 (19.8) |
| Negative          | 41 (38.7)         | 80 (75.5) |
| Unknown           | 9 (8.5)           | 5 (4.7) |
| HIV status at the time of study |         |        |
| Positive          | 106 (100)         | 39 (36.8) |
| Negative          | 0 (0.0)           | 67 (63.2) |

Female sero-discordance rate for the lower, middle, and upper classes were (28/56; 50%), (35/46; 76.1%) and (4/4; 100%) respectively and was statistically significant (P<0.01). There were no significant relationships between age, parity, duration of marriage/cohabitation, average sexual frequency per week and female sero-discordance (Table 2).

Table 2: Association of variables with female sero-discordance.

| Variable       | Discordant female (%) | P-value | Crude OR (95% CI) |
|----------------|-----------------------|---------|------------------|
| Age ≤34        | 57/85 (67.1)          | 0.140   |                  |
| Age ≥35        | 10/21 (47.6)          |         |                  |
| Parity 0       | 28/38 (73.7)          |         |                  |
| Parity 1-5     | 38/66 (57.6)          | 0.429   |                  |
| Parity >5      | 1/2 (50.0)            |         |                  |
| Social class   |                       |         |                  |
| Lower          | 28/56 (50.0)          | <0.01   | (0.041-2.200)    |
| Middle         | 35/46 (76.1)          |         | (ref)            |
| Upper          | 4/4 (100)             |         | (0.041-2.200)    |

Figure 1: Condom use and female sero-discordance.

Out of the 17 male partners that seroconverted after marriage/cohabitation, 12 (70.6%) were of couple that never used condom for intercourse while only 5 (29.4%) used condom.
Majority (86; 81%) of the female participants were on HAART at booking and no statistically significant relationship was found between use of HAART and female sero-discordance (P=0.12). Duration of use of HAART was also not found statistically significant (p=0.08).

CD4+ count of the female partner at booking was found statistically significantly associated with female sero-discordance (P=0.009). Female sero-discordance rate was 26.9% among couples whose female partners CD4+ was less than 200/uL and 73.1% among couple with female CD4+ count of 200/uL and above (Figure 2).

Figure 2: Female CD4+ count and female sero-discordance.

Binary Logistic regression between partners status, CD4+ count and non-use of condom showed that partners of females with CD4+ <200/uL had about 5.5 times higher odds of seroconversion than those with CD4+ of 200/uL and above (crude odds ratio (OR)=5.505, CI 1.365-22.203: P=0.02). Male partners in couples who never used condom for intercourse had about 4.3 times higher odds of seroconversion than those who used condom (crude odds ratio (OR)=4.276, CI 1.818-10.058: P=0.001). Multiple logistic regression between partners’ status, CD4+ count of the female and non-use of condom strongly suggests both were independent risk factors for sero-conversion of HIV negative partners.

DISCUSSION

The finding of 63.2% female discordance rate in this study is close to the rate of 54.7% found in Nnewi (south east Nigeria) but much lower than 89% found in Cross River (south-south Nigeria), and 78.8% found in Benin city (south west Nigeria). However, it is higher than the rate of 7.7% and 48.4% found in Kano and Jos respectively, all in Northern Nigeria. This finding fits into the earlier survey by Sagay et al, which revealed a significant difference in female sero-discordance across sociocultural and geopolitical zones in Nigeria in which it showed a higher female sero-discordance rate in the Southern than Northern Nigeria. Compared to other studies from Africa, present finding is much higher than the population-based estimate of sero-discordance of 2% and 13% found in Rwanda and Zimbabwe and Lesotho respectively. This difference in the finding could be due to varying HIV prevalence across the zones or a reflection of mixed epidemic reported in some studies. It is also possible such differences could be due to varying cultural practices, literacy and socioeconomic levels across the geopolitical zones and countries.

The finding of higher percentage (38.7%) of infection or seroconversion after marriage/cohabitation among female partners compared to 21.3% among male partners may be explained by the hypothesis that women are more at risk to HIV infection (in heterosexual intercourse) than men. The possible causes have been traced to the interrelation of the several factors including differences in anatomical structure of male and female genitals that make men less vulnerable to contracting the virus than women. However, the study could not ascertain whether the HIV infection after marriage in both sexes was due to intra- or extra-marital sexual relationship or other negligible risk factors. To establish that will require a well-controlled prospective study, if ethically permissible.

Majority of the participants had pre-marital/cohabitation knowledge of their HIV status. This is a common practice these days. It can be deduced from Table 1 that about 35 HIV positive women got married to HIV negative partners. It is possible that couples don’t know the implication of their partner’s positive status before marriage and/ or intentionally go into marriage/cohabitation with sero-discordant partners or at worst the sero-positive partner did not disclose their real status to their partner before marriage. Either of these will pose a great problem in the successful prevention of HIV infection. In this study, lower class was 52.8%, middle class 43.4%, and upper class 3.8% of the HIV positive pregnant women. This may imply that HIV infection was more prevalent among women of lower and middle class compared to the upper class in Owerri. It could also be that, for socioeconomic reasons, the upper class was less likely to access health care in a public health facility than the middle and lower classes. Sero-discordance rate for the lower, middle, and upper classes were 50%, 76.1% and 100% respectively, and were statistically significant. This may be explained by the assumption (if most couple had discordant female partners at onset) that sero-discordant couples of higher social class were more likely to be aware and adhere to measures for prevention of transmission to the seronegative partners than those of the lower and middle classes and hence more likely to remain sero-discordant than the later. It could also be that discordant marriages were commoner among the upper than the middle and lower classes. However, the very small population of the upper class in the study is a limitation that requires caution in interpretation of findings.
There were no significant relationships between age, parity, duration of marriage/cohabitation, sexual frequency and female sero-discordance. All things being equal, increased exposure (occasioned by longer duration of marriage/cohabitation and higher sexual frequency) should be associated with increased risk of infection\textsuperscript{11-13} and affect the discordance rate. However, other co-variables like CD4+ count, use of HAART and sexual practice (condom use etc.) can confound the effect of increased exposure.

Condom use was significantly associated with female sero-discordance. Female sero-discordance rate was 62.7\% among couple that used condom (usage ranging from “occasionally” to “always except for conception”) for intercourse compared to 37.3\% among those that never used condom. Although direct causality cannot be established here considering the cross sectional nature of study, and one may argue that sero-discordant couples are more likely to use condom (to protect the uninfected partner) than sero-concordant couples who may feel have no need for protection, this finding supports the fact that correct and consistent use of condom is protective against HIV infection.\textsuperscript{17,18} Also, it is interesting that when the 17 male partners that seroconverted after marriage/cohabitation were cross-tabulated with condom use, 12 (70.6\%) were of couples that never used condom for intercourse compared to only 5 (29.4\%) who used condom. This may imply that, all things being equal, condom use may have reduced male sero-conversion rate from 21\% (17/80) to 6.3\% (5/80) in this study. This finding is tandem with that of Allen et al, where condom use was found to reduce the annual risk of HIV transmission among sero-discordant couple from 20-25\% to 3-7\%.\textsuperscript{17} One can assume that even the 5 that seroconverted despite condom use may be due to improper and inconsistent use.

CD4+ count of the female partner at booking was found statistically significantly associated with female sero-discordance. This can be explained by the fact that CD4+ count is a major disease marker in HIV positive patients as it reflects the immune status of the patients (used as AIDS-defining immunological condition when value is less than 200/\mu{}l\textsuperscript{3}). CD4+ count is also known to have inverse relationship with viral load which has been shown as the most powerful predictor for HIV transmission to an uninfected partner.\textsuperscript{19} However, one of the limitations of this study was inability to assess the viral load of the participants. Again, the power of this finding is weakened by the cross-sectional nature of the study as the CD4+ count at booking may not represent the count at the time of infection of the partner, all things being equal.

Use of HAART and duration of use were not found statistically significantly associated with female sero-discordance. However, this can be explained by the fact that majority (81\%) were already on HAART at booking. It was not ethically permissible to keep some HIV positive pregnant women from HAART for purpose of study considering the already known benefit of HAART to mother, fetus, and partner. The knowledge and practice of pre-exposure prophylaxis were very uncommon (<2\%) among the discordant participants.

**CONCLUSION**

In conclusion, this study has shown a high prevalence of female sero-discordance among HIV positive pregnant women in FMC Owerri. Higher social class, condom use and higher female-partner CD4+ count are significantly associated with female sero-discordance. The knowledge and practice of pre-exposure prophylaxis were very uncommon (<2\%) among the discordant participants.

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**Ethical approval:** The study was approved by the Research and Ethical Committee of FMC Owerri

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