# **BMJ Open** Dyspareunia in HIV-positive and HIV-negative middle-aged women: a cross-sectional study

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#### ABSTRACT

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Dr Ana Lúcia Ribeiro Valadares; anarvaladares@gmail.com **Objectives:** To evaluate whether dyspareunia is associated with HIV status in menopausal women and also to assess which factors are associated with dyspareunia in a group of HIV-positive menopausal women.

**Methods:** A cross-sectional study was conducted with 178 HIV-negative and 128 HIV-positive women aged 40–60 years. The Short Personal Experiences Questionnaire (SPEQ) was used to collect data. Sociodemographic, clinical, behavioural and reproductive factors were evaluated, as well as factors related to the HIV infection. Dyspareunia was defined as pain during intercourse. A bivariate analysis and Poisson multiple regression analysis were performed.

**Results:** Overall, 41.4% of the HIV-positive women reported dyspareunia compared with 34.8% of the HIVnegative women (p=0.242). In the HIV-positive women, bivariate analysis revealed an association between dyspareunia and having a steady partner (p=0.047); the woman's partner having undergone HIV testing (p=0.020); vaginal dryness (p<0.001); muscle/joint pain (p=0.021); physical/emotional violence (p=0.049); urinary incontinence (p=0.004); and the use of lamivudine/zidovudine (p=0.048). The Poisson multiple regression analysis found an association between dyspareunia and vaginal dryness (prevalence ratio (PR) =1.96, 95% CI 1.10 to 3.50, p=0.023) and urinary incontinence (PR=1.86, 95% CI 1.06 to 3.27, p=0.031). **Conclusions:** Dyspareunia was common in this group of HIV-positive women and was associated principally with vaginal dryness and urinary incontinence. The importance of treating dyspareunia within the context of sexual health in this group of women should be emphasised and appropriate management of this issue may reduce the likelihood of lesions on the vaginal wall, which may act as a portal of entry for other infections.

#### **INTRODUCTION**

Dyspareunia is defined as persistent or recurrent genital pain that occurs just before, during or after intercourse. It is one of the most common problems reported by menopausal

# Strengths and limitations of this study

- We have not found studies on dyspareunia in HIV-positive women.
- We have highlighted the importance of vulvovaginal atrophy and its association with dyspareunia in middle-aged HIV-positive women.
- We have shown that HIV infection was not significantly associated with dyspareunia, probably because HIV-positive women had few HIV-related symptoms.
- The results of this study may help physicians to focus attention on vulvovaginal atrophy and its consequences in this group of HIV-positive women.
- It is a cross-sectional design study.
- There were some differences in the clinical characteristics of the HIV-positive and HIV-negative women.

women. The variation in the frequency of dyspareunia probably reflects many issues including sociocultural aspects, the period of observation during which the condition was evaluated (ever, the past year) and the duration or design of the study under discussion (questionnaire wording, participants).<sup>1</sup>

For women of all ages, the pain caused by dyspareunia often results in distress, impaired sexual functioning and poor sexual enjoyment, difficulty in relationships and a poorer quality of life. In postmenopausal women, dyspareunia may also intensify personal issues related to ageing, body image and health.<sup>2</sup>

As with most of the sexual difficulties faced by women in midlife and beyond, dyspareunia is typically considered a consequence of declining ovarian hormone levels and is usually attributed to vaginal atrophy;<sup>3</sup> however, other factors may also be involved.<sup>4</sup> In fact, psychosexual and biological factors (including muscular, endocrine, immune, neurological, vascular and iatrogenic factors) that predispose to, precipitate and perpetuate the condition may interact with different degrees in the individual woman, contributing to a continuum of symptoms of increasing severity, with the potential to impair sexual intercourse.<sup>5</sup> Age,<sup>6</sup> depression, anxiety and sexual dysfunction in the partner<sup>4 5</sup> are some of the other factors associated with dyspareunia. It seems that cognitive– emotional variables (catastrophisation, depression, anxiety) are significant predictors of dyspareunia and relationship adjustment variables were inversely associated with pain severity.<sup>7</sup> Findings also suggest that dyspareunia impacts the psychosexual adjustment of affected women as well as of their partners.<sup>8</sup>

Menopausal women who are HIV positive may present a unique set of issues that could affect their sexuality. These issues may include the meaning of their illness, their quality of life, HIV transmissibility, and the dilemma of whether or not to disclose the condition to their partner. Florence *et al*<sup>p</sup> reported sexual dysfunction to be common in HIV-positive women, principally as a result of their HIV status and of psychological factors that included depression, irritability and anxiety. On the other hand, women with better mental health after HIV diagnosis, a more positive attitude towards living with HIV, a better quality of life, fewer HIV-related symptoms and who had never used injectable drugs were found to have better sexual functioning.<sup>10</sup> A possible role of antiretroviral drugs in causing sexual dysfunction has been a matter of debate. While some studies have suggested that antiretroviral therapy (ART) indeed plays a role in sexual function, others have failed to find any such association.<sup>11</sup>

The majority of studies on dyspareunia have failed to deal with factors associated with HIV infection, a topic yet to be fully investigated in HIV-positive women during the ageing process. Therefore, the objectives of the present study were to evaluate whether dyspareunia is associated with HIV status in middle-aged women and to assess the factors associated with dyspareunia in HIV-positive middle-aged women.

### METHODS Study design

## Study design

A cross-sectional study was conducted in 537 women aged 40–60 years, of whom 273 were HIV-positive and 264 were HIV-negative and screened for inclusion. Patients were recruited at the infectious diseases and HIV outpatient clinics (HIV-positive women) and at the menopausal ambulatory care (HIV-negative women), both at the Teaching Hospital of the University of Campinas (UNICAMP). Patients were also invited to participate at the infectious diseases outpatient public clinic (HIV-positive women) in Belo Horizonte. Of these, 178 HIV-negative women and 128 HIV-positive women had had vaginal intercourse in the previous month and were willing to answer a questionnaire on dyspareunia. These women were then admitted to the study.

For inclusion in the HIV-positive group, laboratory confirmation of the women's seropositive status by one of the recommended tests (ELISA or Western Blot) was required (all of them had it), while the women recruited to the HIV-negative group had to have tested negative. The blood sample tests of HIV-negative and HIV-positive women were collected at the moment of admission in this study (follicle stimulating hormone (FSH), luteinising hormone (LH) and thyroid stimulating hormone for all; ELISA or Western Blot HIV tests for HIV-negative women; and Viral load and CD4 cells for HIV-positive women). Exclusion criteria consisted of nursing mothers, bilaterally oophorectomised women and those unable to answer the questionnaire. The evaluation instrument was the Short Personal Experiences Questionnaire (SPEQ).<sup>12 13</sup> Sociodemographic, clinical, behavioural and reproductive characteristics were assessed as well as issues relating to the HIV infection and partner-related factors.

## **Dependent variable**

The dependent variable dyspareunia, defined as pain during sexual intercourse, was graded from 1 to 6, where 1 referred to the absence of pain and 6 to maximum pain. A score of less than two was considered to represent the absence of dyspareunia and a score of two or more to represent the presence of dyspareunia.<sup>4 12 13</sup>

## **Independent variables**

The independent variables were dichotomised as follows: HIV status (positive/negative); skin colour (white/other); marital status (has a stable partner/no stable partner); schooling ( $\leq 7$  years/ $\geq 8$  years); employment (yes/no); monthly family income (≤US\$750/>US \$750; receives pension (ves/no); smokes (ves/never or ex-smoker); alcohol use (currently drinks or used to drink/never drank); hot flushes (yes/no); depression (yes/no); vaginal dryness (yes/no); urinary incontinence (yes/no); weight gain (yes/no); muscle and joint pain (yes/no); self-perception of health (excellent or good/poor or very poor); suffers or has already suffered some form of physical or emotional violence (yes/no); has been forced to have intercourse (yes/no); uses (yes/no); chronic disease: hypothyroidism statins (yes/no); FSH (<40/≥40), LH (<25.7/≥25.7); age at first sexual intercourse ( $\leq 19$  years/ $\geq 20$  years); other type of sexual intercourse in the preceding month: giving oral sex (yes/no), receiving oral sex (yes/no); woman lives with sexual partner (yes/no); number of sexual partners in the previous year (none/ $\geq 1$ ); partner underwent HIV testing (yes/no); quality of life following diagnosis (changed/unchanged); CD4 cell count  $(<350/\geq350)$ ; CD4 cell count nadir  $(<199/\geq200)$ ; use of antiretroviral drug 3TC (lamivudine, Epivir; yes/no); use of antiretroviral drug tenofovir (yes/no); use of antiretroviral drug lamivudine/zidovudine (yes/no); use of antiretroviral drug efavirenz (yes/no); antiretroviral drug used in the past: lamivudine/zidovudine (yes/no); and antiretroviral drug used in the past: efavirenz (yes/ no). Menopausal status was classified as premenopausal, perimenopausal or postmenopausal. Women were considered premenopausal if they continued to have

regular menstrual cycles similar to those present during the woman's reproductive life. They were considered to be in the perimenopause if their menstrual cycles were irregular and they had been amenorrhoeic for less than 12 months. Finally, women were classified as postmenopausal if they had been amenorrhoeic for 12 months or more.<sup>14</sup> Data on physical activity was obtained through two questions: Do you practise physical exercise or participate in sports every week? How often in a week do you practise physical exercise or participate in sports? It was classified in up to two times a week or three or more times a week. Vaginal lubrication during sexual activity was graded from 1 to 6, where 1 referred to the absence of lubrication and 6 to maximum lubrication. This was dichotomised into four or less or more than four.

#### **Statistical analysis**

A bivariate analysis was performed in which dyspareunia was considered the dependent variable (dyspareunia) and analysed as a function of the independent variables. Pearson's  $\chi^2$  test and the Yates correction were used to compare the groups.<sup>15</sup> The Poisson multiple regression analysis<sup>16</sup> was adjusted in the various models for each one of the independent variables to evaluate the factors associated with the presence of dyspareunia.

### Ethics

The study was approved by the internal review board of CAISM/UNICAMP and was conducted in compliance with the current version of the Declaration of Helsinki and with Resolution 196/96 of the Brazilian National Committee for Ethics in Research (CONEP) and its subsequent revisions. This study forms part of a larger study evaluating menopausal symptoms, bone mass, sexual function and metabolic markers. Process: CEP: 407/2010, CAAE 0313.0.146.000-10.

Women who agreed to participate in the study after receiving instructions from the researchers and who signed a free informed consent form were included.

#### RESULTS

The HIV-positive women were younger and less likely to have a steady partner, to be employed or to have a formal education compared with the HIV-negative women. More than half the HIV-positive women were premenopausal or perimenopausal. The characteristics of the women interviewed are shown in table 1.

Overall, 41.4% (n=53) of the HIV-positive women and 34.8% (n=62) of the HIV-negative women reported dyspareunia. There was no association between HIV status and dyspareunia (p=0.242). Furthermore, in the multiple regression analysis of the entire sample of HIV-positive and HIV-negative women taken together (n=306), dyspareunia was not associated with HIV status, but was associated with vaginal dryness (prevalence ratio (PR)=2.06, 95% CI 1.37 to 3.10, p=0.001) and urinary incontinence (PR=1.68, 95% CI 1.14 to 2.46, p=0.008). In the HIV-positive group, 91.4% of the women were currently using ART, and of these 87% reported using ART regularly (data not presented as a table). Approximately 77% of the HIV-positive women had a CD4 cell count nadir >200. The most common way in which HIV had been acquired was by heterosexual transmission, and the average duration of the HIV infection was 9.5 ±5.6 years (mean±SD), with a mean duration of therapy of 8.7 years±4.5 (mean±SD). A more detailed description of the HIV-infected women is provided in table 2.

Bivariate analysis revealed an association between dyspareunia in the HIV-positive women and having a steady partner (p=0.047); the woman's partner having undergone HIV testing (p=0.020); vaginal dryness (p<0.001); muscle/joint pain (p=0.021); physical/emotional violence (p=0.049); urinary incontinence (p=0.004); and the use of lamivudine/zidovudine (p=0.048), table 3.

According to the Poisson multiple regression analysis, the principal factors associated with dyspareunia in the group of HIV-positive women were: vaginal dryness (PR=1.96; 95% CI 1.10 to 3.50; p=0.023) and urinary incontinence (PR=1.86; 95% CI 1.06 to 3.27; p=0.031; table 4).

## DISCUSSION

The objectives of this study were to evaluate whether HIV status was associated with dyspareunia and to assess the factors associated with pain during sexual intercourse in HIV-positive women aged 40–60 years.

The calculated number of women required for the sample size was 74 per group;<sup>4</sup> however, to enable analysis of the HIV-positive group alone, the required number was 188 women.<sup>17</sup> Since the actual sample size achieved was 128, the absolute difference was 8.5%, acceptable since it is less than 10%.

Information on dyspareunia in HIV-positive women is scarce, especially in middle-aged women. To the best of our knowledge, no other studies have been conducted on dyspareunia in HIV-positive women. It has been reported that sexual function in HIV-positive women may be driven principally by psychological factors and other problems related to HIV infection.<sup>17 18</sup> This study, however, found that in the overall sample of HIV-positive and HIV-negative women dyspareunia was not affected by HIV status. This finding is in agreement with the results of another author, who also reported that few women believed HIV in itself to be the cause of any decline in their sexual functioning, since those women had good immunovirological status.<sup>10</sup> One supposes that results would be different in a sample of women without good HIV control. In this study, more than threequarters of the HIV-positive patients had a CD4 cell count nadir >200 and CD4 cell counts >500 in their last evaluation, thus reflecting adequate control of the disease. This may partially explain why no association was found between HIV status and dyspareunia. In line with this, another study showed that women with CD4

Table 1         Some characteristics of women according to HIV status				
	Group			
Characteristic	HIV-infected (%; n=128)	HIV-uninfected (%; n=178)	p Value	
Age (years)				
40–44	43.8	24.7	<0.002*	
45–49	28.9	29.2		
50–54	15.6	23.1		
≥55	11.7	23.0		
Skin colour				
White	35.2	47.2	0.047†	
Non-white	64.8	52.8		
Number of deliveries				
Up to 1	25.0	25.4	>0.999†	
≥2	75.0	74.6		
Marital status				
With partner	58.6	87.1	<0.001†	
Without partner	41.4	12.9		
Schooling (years)				
≤7	62.5	40.4	<0.002*	
8–11	23.4	37.1		
≥12	14.1	22.5		
Employment status				
Yes	59.4	71.9	0.030†	
No	40.6	28.1		
Menopausal status				
Premenopausal	39.8	24.7	<0.002*	
Perimenopausal	28.1	21.4		
Postmenopausal	32.1	53.9		
Current smoking status				
Yes/former	28.1	15.2	0.009†	
No	71.9	84.8		
Physical activity				
Up to 2 times/week	77.3	74.2	0.614†	
≥3 times/week	22.7	25.8		
*Pearson's $\chi^2$ ; and Yates's $\chi^2$ .				

counts  $\leq 199$  cells/µL reported poorer sexual functioning compared with those whose cell count was  $\geq 200$ .<sup>19</sup> Other studies have shown that the CD4 cell count nadir may also have long-term consequences in terms of prognosis and mortality.<sup>20</sup>

Nevertheless, the CD4 cell count nadir and the last CD4 evaluation were not associated with dyspareunia in this study, probably because of the small number of women with these low values.

The most important factors associated with dyspareunia in the logistic regression analysis, in HIV-positive and HIV-negative groups analysed together, and in the HIV group analysis were vaginal dryness and urinary incontinence, both of which are urogenital disorders associated with oestrogen deficiency. The association between vaginal dryness and pain during sexual intercourse has been well documented in the literature, in addition to its consequence on vulvovaginal health.<sup>21–23</sup> With respect to the association between urinary incontinence and dyspareunia, the findings of this study are in agreement with the results published by Salonia *et al*,<sup>24</sup> who evaluated 216 women with urinary incontinence and found 44% of dyspareunia in these women. The type of urinary incontinence was not evaluated in this study. Nevertheless, there is good evidence that the effects of urinary incontinence on sexual functioning are similar irrespective of whether the condition has been classified as stress, urge, mixed incontinence<sup>25</sup> or even interstitial cystitis.<sup>26</sup> Urinary incontinence is associated with feelings of embarrassment and inadequacy as well as low self-esteem. It may also be associated with dyspareunia.<sup>24</sup>

Factors associated with dyspareunia in HIV-positive women:

In the bivariate analysis, the fact that the woman's partner had not been tested for HIV was associated with less dyspareunia. It is reasonable to speculate that not knowing her partner's HIV status may in some way 'minimise' a woman's concerns regarding transmission and reduce the probability of tension and dyspareunia.<sup>27</sup> Another factor related to the sexual partner that was associated with an increase in dyspareunia in the bivariate analysis was the woman having a steady partner, although this association was borderline. One

 Table 2
 Characteristics associated to HIV status

 associated with dyspareunia in women with a sexual
 partner in the month before the interview (n=128)

Characteristics	Ν	%
HIV duration of infection	Mean: 9.5	SD: 5.06
(n=125; years)		
Duration of HIV therapy	Mean: 8.7	SD: 4.47
(n=93; years)		
Nadir CD4 levels*		
0–100	18	14.5
101–200	10	8.1
201–500	62	50.0
Last CD4 levels*		
0–100	5	4.0
101–200	1	0.8
201–500	43	34.7
>500	75	60.5
Total	124	100.0
HIV risk factor for acquisition		
Heterosexual acquisition	97	75.8
Illicit drug use	3	2.3
Blood transfusion	2	1.6
	26	20.3
Total	128	100.0
Use of TARV		
Yes	117	91.4
No	11	8.6
Total	128	100.0

**Table 3** Factors associated with dyspareunia (score  $\geq$ 2) in middle-aged HIV-positive women: bivariate analysis

	Dyspareunia %			
Variable	n	Score≥2	Score<2	p Value
Marital status				0.047
Married/live together	75	49.3	50.7	
Do not live together	53	30.2	69.8	
Did partner have an				0.020
HIV test?				
Yes	88	50.0	50.0	
No	27	22.2	77.8	
Vaginal dryness				<0.001
Yes	53	58.5	41.5	
No	71	26.8	73.2	
Muscular/articular pain				0.021
Yes	83	49.4	50.6	
No	45	26.7	73.3	
Physical/emotional				0.049
violence				
Yes	128	55.5	44.5	
No	174	32.8	67.2	
Urinary incontinence				0.004
Yes	41	61.0	39.0	
No	87	32.2	67.8	
Use of biovir				0.048
Yes	57	29.8	70.2	
No	63	49.2	50.8	

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Table 4 Variables associated	d with dyspareunia in
HIV-positive women with a set	xual partner in the month
before the interview	

Variable	PR	95% CI	p Value		
Vaginal dryness	1.96	1.10 to 3.50	0.023		
Urinary incontinence	1.86	1.06 to 3.27	0.031		
Poisson multiple regression (n=124). PR, prevalence ratio.					

explanation for this finding may lie in the psychological problems generated by the infection itself, which may arise more frequently in stable relationships.<sup>27 28</sup> As one has not controlled for frequency of intercourse, one thought is that the dyspareunia is probably due a lower frequency of intercourse rather than an inferior quality of the relationship. Results of the bivariate analysis revealed an association between physical/emotional violence and dyspareunia. Violence is known to be associated with poorer psychological adjustment and adverse sexual health outcomes in women.<sup>29 30</sup> In addition, having muscle pain was associated with dyspareunia in the bivariate analysis. This finding is in line with another study showing that musculoskeletal pain often interferes with sex and may be associated with dyspareunia.<sup>31</sup> A borderline association was found between the use of lamivudine/zidovudine and dvspareunia; however, no explanation for this association was found in the literature. One may hypothesise that dyspareunia in these women could be due to the side effects of these drugs on depression.

There are some limitations to this study that must be taken into account. First, its cross-sectional design does not permit any conclusions to be drawn with respect to causality. It is also important to note that it was a clinical sample. So the results found in this study may not be extrapolated to the general population. Furthermore, there were some differences in the clinical characteristics of the HIV-positive and HIV-negative women. These differences could be attributed to the fact that the HIV-negative women were selected at specialist outpatient clinics providing care to menopausal women. By selecting HIV-positive women also in menopausal outpatient care, maybe groups would be similar. Nevertheless, multivariate analysis, conducted in a sufficiently large sample of women after controlling for confounding factors, confirmed that HIV infection was not significantly associated with dyspareunia. Good control of the HIV infection and the regular use of ART by the majority of the women may have brought this group of women closer to the HIV-negative group in terms of their characteristics.

## CONCLUSIONS

In this study population, HIV infection was not associated with the presence of dyspareunia. The principal factors associated with dyspareunia in HIV-positive

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women were vaginal dryness and urinary incontinence. These data indicate a need for multidisciplinary care for HIV-positive menopausal women, paving particular attention to ensuring the women's compliance with ART and offering improved care when these two clinical situations are present to ensure that these women come as close as possible in this respect to HIV-negative women. Greater attention to dyspareunia as a potential component of women's general HIV and sexual care is warranted. A proactive approach to conversations about vulvovaginal atrophy would improve the management of dyspareunia and vaginal dryness. In addition to improving the quality of these women's sexual lives, we hypothesise that appropriate management of this issue may reduce the likelihood of lesions on the vaginal wall, which may act as a portal of entry for other infections.

#### Collaborators Lívia Akl helped in the collection of data.

**Contributors** ALRV, AMP-N and LC-P contributed to the conception or design. ALRV, DdCG, WCD and ASM contributed to the acquisition of data. MHdS, ALRV, AMP-N and LC-P contributed in the analysis of data. ALRV, AMP-N, LC-P and MHdS contributed in the interpretation of data. All the authors were involved in the drafting of the manuscript or revising it critically for intellectual content. All the authors gave final approval of the version to be published.

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#### Competing interests None.

#### Patient consent Obtained.

Ethics approval The project was approved by the internal review board of CAISM/UNICAMP and was conducted in compliance with the current version of the Declaration of Helsinki and with Resolution 196/96 of the Brazilian National Committee for Ethics in Research (CONEP) and its subsequent revisions.

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Data sharing statement We have used a questionnaire to collect data for this study. The instrument used to collect data is available by emailing anarvaladares@gmail.com

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