Sex Work as an Emerging Risk Factor for HIV Seroconversion among People who inject drugs in the SurvUDI Network

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Short summary: Among Central Eastern Canada PWID, sex work was
 independently associated with HIV incidence. Participants who engaged in sex work
 were compared with other participants with respect to HIV risks (word count: 29).

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46 **ABSTRACT**

Background: Recent analyses have shown an emerging positive association
between sex work and HIV incidence among people who inject drugs (PWIDs) in the
SurvUDI network.

Methods: Participants who had injected in the past 6 months were recruited across 50 the Province of Quebec and in the city of Ottawa, mainly in harm reduction 51 programs. They completed a questionnaire and provided gingival exudate for HIV 52 antibody testing. The associations with HIV seroconversion were tested with a Cox 53 proportional hazard model using time-dependent covariables including the main 54 55 variable of interest, sexual activity (sex work; no sex work; sexually inactive). The final model included significant variables and confounders of the associations with 56 sexual activity. 57

Results: Seventy-two HIV seroconversions were observed during 5 239.2 personyears of follow-up (Incidence rates: total=1.4/100 person-years (py), [95%CI: 1.1-1.7]; sex work=2.5/100 py [1.5-3.6]; no sex work=0.8/100 py [0.5-1.2]; sexually inactive=1.8/100 py [1.1-2.5]). In the final multivariate model, HIV incidence was significantly associated with sexual activity (sex work: adjusted hazard ratio (AHR)=2.19 [1.13-4.25],; sexually inactive: AHR=1.62 [0.92-2.88]), and injection with a needle/syringe used by someone else (AHR=2.84, [1.73-4.66]).

65 Conclusions: Sex work is independently associated with HIV incidence among 66 PWIDs. At the other end of the spectrum of sexual activity, sexually inactive PWIDs 67 have a higher HIV incidence rate, likely due to more profound dependence leading to 68 increased vulnerabilities, which may include mental illness, poverty and social exclusion. Further studies are needed to understand whether the association
between sex work and HIV is related to sexual transmission or other vulnerability
factors.

Keywords : people who inject drugs, HIV incidence, sex work, sexual activity,
injection behaviours

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77 **INTRODUCTION**

The Joint United Nations Programme on HIV/AIDS and World Health Organization 78 (UNAIDS/WHO) working group on global HIV/AIDS and STI surveillance provide 79 recent updated guidelines for HIV surveillance activities in several epidemiological 80 contexts¹. In concentrated HIV epidemics, it is recommended to carry out regular 81 biobehavioural surveys in key populations at higher risk for HIV infection in order to 82 understand the local HIV epidemic and how it is changing, as well as to identify 83 84 opportunities to control the epidemic. These key populations include people who inject drugs (PWIDs), men who have sex with men (MSM) and commercial sex 85 workers as well as their clients, because behaviours that increase the HIV risk are 86 frequent among these populations, i.e. unprotected sex with multiple partners, 87 injection drug use (needle/syringe-sharing) and unprotected anal sex ¹. 88

People who inject drugs are considered at increased risk for HIV infection mainly 89 because of sharing contaminated syringes and other injection paraphernalia. Sexual 90 transmission is possible, but its extent is generally difficult to estimate ². It has been 91 92 suggested that in PWIDs, sexual risks are present but may be masked or overshadowed by parenteral risks ³. This would be rather likely considering the much 93 higher risk of HIV transmission of needle/syringe-sharing (0.7% per exposure) 94 compared to oral (maximum of 0.06% per exposure) or vaginal sex (maximum of 95 0.15% per exposure), with the exception of anal sex (maximum of 3% per 96 exposure)⁴⁻⁶. Thus, sexual transmission is more likely to occur in specific situations 97 such as in the context of commercial sex trade or male homosexual intercourse i.e. 98 high number of sex partners or frequent anal sex. This in turn may lead to bridging of 99 HIV and other sexually transmitted and blood-borne infection epidemics between 100

101 PWIDs, who cumulate risks, and individuals who do not, as for example, persons in 102 the general population ^{7,8}.

Important sex differences exist for sex work and should be taken into account ⁹⁻¹¹. 103 For example, sex work is frequent in PWIDs, but generally much more frequent in 104 female than in male PWIDs¹¹. It occurs generally in the context of heterosexual sex 105 for women (i.e. very low HIV prevalence in clients in most developed countries) and 106 107 women are particularly vulnerable to sexual violence and abuse including negotiation of condom use ¹⁰. On the other hand, sex work in men takes place generally in the 108 109 context of sex with other men (i.e. very high HIV prevalence in clients in most developed countries) ¹². 110

111 The SurvUDI network is a biobehavioural survey among PWIDs of Central Eastern Canada, ongoing since 1995. Recent analyses of SurvUDI data have shown an 112 emerging positive association between sex work and HIV incidence among PWIDs 113 ¹³. In this previous analysis, a time period interaction variable was included (1995-114 2002 vs. 2003-2009) in a multivariate model and the association of sex work with 115 HIV incidence was significant for 2003-2009 but not for 1995-2002. Considering that 116 the questionnaire for the period 2004-2014 includes much more variables than for 117 the period 1995-2002, repeating this analysis for 2004-2014 has allowed a better 118 characterization of participants according to the main variable of interest. The 119 hypothesis was that the association with sex work would be detected for the period 120 2004-2014. The objectives of the present study were to examine the association 121 122 between sex work and HIV seroconversion among PWIDs in the SurvUDI network between 2004 and 2014 and to compare participants engaging in sex work with 123 124 other participants with respect to HIV risks.

126 **METHODS**

Study design and subjects. The complete methodology of the SurvUDI study has 127 been described elsewhere ¹⁴. Briefly, the SurvUDI network is an ongoing 128 biobehavioural survey for HIV, HCV and associated risk behaviours among PWIDs in 129 Eastern Central Canada. The network was implemented in 1995 and targets hard-to-130 reach, mostly out-of-treatment PWIDs. Eligibility criteria include being aged 14 and 131 older, injecting at least once within the past 6 months, speaking French or English 132 and being able to provide informed consent. Participants are recruited in urban 133 areas, including Montréal and neighbouring South Shore, Québec City, the Hull-134 Ottawa region, and five semi-urban areas of the province of Québec. Overall, since 135 2004, 94.6% of participants were recruited in harm reduction programs. Others were 136 recruited in drop-in centres, detention centres, detoxification clinics, and 137 rehabilitation programmes. Participation includes an interviewer-administered 138 139 questionnaire and collection of gingival exudate using the Orasure device (Bethlehem, Pennsylvania, US) for HIV and HCV antibody testing. The study design 140 is an open cohort of services where participants who attend harm reduction 141 programs more than once at times of study enrollment are followed longitudinally. 142 The present sample includes participants recruited from March 2004 to March 31 143 2014 who were initially HIV seronegative and with at least one follow-up visit. 144 Participants are identified using an encrypted code based on their initials, birth date 145 and sex, and they are given a stipend ranging from CAN\$5.00 to \$10.00 at the end 146 of each study visit. All procedures have been approved by the ethics committee of 147 the Centre de recherche du CHU de Québec. 148

149 Study variables. The dependent variable was HIV incidence. HIV infection was set at the midpoint between the last HIV negative follow-up visit and the visit when the 150 HIV positive result was first detected. Potential confounders were identified based on 151 152 a literature review and on previous analyses in this cohort. Covariates considered in the multivariate analyses as potential confounders included age (<25; ≥25 153 years)^{13,15,16}, sex ^{13,15,16}, the region where the interview took place (urban or semi-154 urban/rural)¹³, high school completion¹⁷, homelessness^{18,19}, recent incarceration 155 ^{15,19}, injected and non injected drug use ^{13,20}, cocaine as the most often injected drug 156 ¹³, injection with strangers (unknown people) ^{14,18}, injection with needles/syringes 157 used by someone else ^{13,17}, daily injection ^{13,21}, the number of sexual partners ¹⁷, 158 consistent condom use for vaginal and anal sex ²², always injecting alone and 159 injecting in public places (previous analyses, unpublished data). Sexual activity, the 160 main exposure of interest, was categorized as a three-level variable, namely, being 161 sexually active without engaging in sex work (no sex work, NSW), being sexually 162 active and engaging into sex work (sex work, SW) or being sexually inactive. Sex 163 work was defined as having client sex partners in the past six months, i.e. partners 164 giving money, drugs, goods or other things in exchange for sex. The choice for the 165 reference category (no sex work) was based on the hypothesis that this group would 166 show the lowest risk compared to the other two groups. This assumption was based 167 168 on previous analyses in our population showing that sexual inactivity in male IDUs was associated with a higher HIV prevalence ¹⁴. In the descriptive analyses of 169 participants according to sexual activity, several variables considered relevant were 170 presented even if not retained as confounders in the multivariate analysis. Sexual 171 orientation was added to the questionnaire in February 2011. Questions on 172

behaviours referred to the 6 months prior to the interview, except for daily injectionwhich refers to the last month.

175 Laboratory procedures. Collected oral fluid samples were kept at 4°C and shipped within 2 weeks to the Laboratoire de santé publique du Québec (LSPQ; Institut 176 national de santé publique du Québec), where they were centrifuged upon reception. 177 The extracted liquid was kept at -20°C for a maximum of 6 weeks until analysis. The 178 presence of HIV antibodies was assessed by enzyme immunoassay (EIA) using 179 HIV-1 Vironostika Microelisa System (bioMérieux, Durham, North Carolina, USA) 180 from 2004 to 2009 and GS HIV-1/HIV-2 PLUS O EIA (Bio-Rad Laboratories 181 (Canada) Ltd., Montréal, Qc, Canada) thereafter. The presence of HCV antibodies 182 was assessed using ORTHO® HCV 3.0 ELISA Test System (Bio-Rad Laboratories 183 (Canada) Ltd., Montréal, Qc, Canada) according to a modified method developed by 184 Judd et al.²³. Samples were considered negative if results were less than 75% of the 185 cut-off value. Sample results that were greater than 75% of the cut-off value were 186 retested in duplicate. A sample was deemed positive if at least two out of three 187 results were greater than the cutoff value. 188

Statistical analyses. HIV seronegative participants at baseline who had at least one 189 190 follow-up visit for the period from 2004 to 2014 were considered when assessing HIV incidence rates. Baseline characteristics and behaviours are those reported at the 191 second visit, i.e. exposure during the first time interval in the Cox proportional hazard 192 model. Using the Kaplan-Meier method, we estimated the cumulative probability of 193 seroconversion during follow-up. The cumulative HIV incidence rates were plotted 194 and compared for sexual activity reported at baseline using the log-rank test. 195 Bivariate and multivariate Cox proportional hazard regression analyses were carried 196

197 out to evaluate the association between HIV incidence and sexual activity. Behavioural variables and age were treated as time-dependent variables. The 198 exposure used for a given time interval corresponds to the exposure measured at 199 200 the visit at the end of this interval, i.e. exposure reported for the preceding 6 months. The final model included significant (p<0.05) variables and confounders i.e. variables 201 202 changing other AHR by 10% or more when removed of the complete model. Effect modification by sex was also tested for the association between sexual activity and 203 204 HIV incidence, and was considered statistically significant at a p value of 0.15 or less 205 (Wald chi-squared test). To compare risk profiles between participants engaged in sex work and other participants, cross-sectional sex-stratified descriptive analyses 206 207 were performed using the last visit for non-seroconverters and the visit when HIV 208 infection was first detected for seroconverters. Cross-sectional comparisons were performed using Pearson's chi-squared tests or ANOVA. All analyses were carried 209 210 out using the SAS statistical suite software version 9.4 (SAS Institute Inc., Cary, 211 North Carolina, USA).

214 **RESULTS**

Table 1 shows baseline characteristics of participants. The sample included 1 528 participants cumulating 5 239.2 person-years of follow-up, with a majority of men and of subjects from urban sites. Only a small proportion of PWIDs were less than 25 years old. Homelessness as well as cocaine as the most often injected drug were frequently reported. Sex work was more frequently observed in women whereas sexual inactivity was more frequent in men.

Figure 1 shows cumulative hazard of HIV seroconversion and HIV incidence 221 according to baseline sexual activity for the ten-year period (2004-2014). Occurrence 222 223 of seroconversions was proportionally distributed throughout the follow-up period in 224 all groups. A statistically significant difference was observed between groups, with the highest cumulative hazard observed in participants who reported sex work 225 226 (p<0.0001). The highest HIV incidence rate was observed in participants who reported sex work (more than three-fold higher compared to sexually active 227 participants who did not report sex work), followed by HIV incidence rate in sexually 228 229 inactive individuals.

In the final multivariate model (Table 2), the association of HIV incidence with sex work is the second strongest with an adjusted incidence rate ratio of 2.19 [1.13-4.25]. The strongest association was observed for injection with a needle/syringe used by someone else (AHR=2.84, [1.73-4.66]). Effect modification analyses showed that the association between HIV incidence and sexual activity did not vary significantly according to sex (p-value for effect modification=0.5729). Sex-stratified rate ratios are presented in table 2. 237 Tables 3 and 4 show cross-sectional, sex-stratified analyses of injection and sex behaviours according to sexual activity. Male SW (Table 3) were significantly 238 239 younger and more likely to be homeless and to report a history of incarceration in the 240 past 6 months compared to other groups. Male SW were also significantly more likely to use heroin by injection, to report cocaine as the most often injected drug and 241 242 to use non-injected crack/freebase, amphetamines and ecstasy. Male SW were significantly more likely to inject with strangers and to inject with needles/syringes 243 244 used by someone else obtained mainly from strangers compared to male NSW. A 245 large proportion of male SW had sex with at least 6 male sex partners in the past 6 months (41.2%) and reported at least one female sex partner (58.8%). Among male 246 247 SW who had multiple male sex partners, 72% reported heterosexual or bisexual 248 orientation (data not shown). The proportion of men who reported inconsistent condom use for vaginal or anal sex is high (sex work=42.6%, no sex work=62.7%). 249 250 Sexually inactive men were significantly more likely to report daily injection and to 251 always inject alone compared to other groups. Male SW and sexually inactive men were more likely to report injection with needles/syringes used by someone else 252 obtained mainly from strangers compared to male NSW. HCV prevalence (positivity 253 254 for HCV antibodies) was significantly higher in male SW compared to other groups (intermediate in sexually inactive men). 255

Female SW (Table 4) were more likely to be homeless and to report a history of incarceration in the past 6 months compared to other groups. They were also significantly more likely to use injected cocaine, to report cocaine as their most often injected drug and to use non-injected crack/freebase and ecstasy. Female SW were more likely to report injection with strangers and injection with a needle/syringe used by someone else obtained mainly from strangers. The majority of women were sexually active. Among female NSW, 25.3% reported at least two male sex partners in the past six months. Approximately half of female SW had sex with at least 21 male partners in the past six months (48.1%). A high proportion of women reported inconsistent condom use for vaginal or anal sex (sex work=60.9%, no sex work=84.1%). Sexually inactive women were significantly more likely to always inject alone. HCV prevalence (positivity for HCV antibodies) was significantly higher in female SW compared to the other groups.

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271 **DISCUSSION**

The objectives of the present study were 1) to characterize the association between 272 273 sex work (i.e reporting client sex partners) and HIV seroconversion among PWIDs in the SurvUDI network between 2004 and 2014 and 2) to describe the characteristics 274 and risk profile of participants who reported sex work in the past 6 months. The 275 276 highest HIV incidence rate was observed in participants who reported sex work. In the multivariate analysis, sex work and injection with a needle used by someone else 277 were significantly and independently associated with HIV incidence. Both male and 278 female PWIDs who reported sex work were more frequently in situation of 279 homelessness and reported more unsafe injecting as well as sexual behaviours. 280

In the present study, the association between HIV incidence and sexual activity did 281 not show significant effect modification by sex. Although the direction of the 282 association is the same for men and women, the association observed in men is not 283 significant and is weaker than in women. This is not surprising given the small 284 number of men reporting client sex partners. Thus, it is not clear whether the 285 absence of a significant modifying effect by sex, as well as the absence of a 286 significant association among men in the sex-specific analyses, are due to a lack of 287 power (inability to detect an existing effect modification that could show an 288 association among women only) or to the absence of modifying effect. 289

Recently, Kerr *et al.* ¹⁵ also observed a significant association between sex work and HIV incidence in Vancouver PWIDs (men and women combined), but it did not remain significant in multivariate analysis. A methodological issue that may explain differences with the present study is the definition of the reference category for the

sex work variable. Kerr et al. 15 defined sex work as a dichotomous variable where 294 «no sex work» presumably included sexually inactive individuals. As observed in the 295 present study, sexually inactive PWIDs may differ from other participants in their 296 297 characteristics as well as their drug use risk patterns. In the present study, compared to sexually active male who did not report sex work, sexually inactive male were 298 299 significantly more likely to report daily injection and injection with needles/syringes used by someone else obtained mainly from strangers. Sexually inactive participants 300 may have a more profound dependence associated to increased vulnerabilities, 301 which may include mental illness, poverty and social exclusion. This could have 302 confounded the association observed in the above-mentioned study. Several other 303 304 studies have examined the link between sex work or sexual risk profile and HIV transmission in PWIDs of developed countries ^{16,17,24-29}. Most of these studies 305 reported very similar findings i.e associations with homelessness, incarceration, 306 cocaine injection and crack use. 307

Unprotected sex with multiple partners, including unprotected anal sex ³⁰, are common among the participants of our study. It is difficult to estimate the relative contribution of unsafe injection and sexual behaviours to the transmission of HIV. Vickerman *et al.* ² have recently proposed a deterministic mathematical model to estimate the proportion of HIV infections due to sexual transmission in PWIDs populations. In this model, the HCV prevalence in HIV-infected PWIDs and the HIV/HCV prevalence ratio are used as markers of sexual transmission.

Several authors have suggested that in PWIDs, sexual risks are present but may be masked by parenteral risks ³. This may be an explanation why sex work was not significantly associated with HIV incidence in PWIDs of the SurvUDI network for the

period 1995-2002 while a positive association emerged afterward ¹³. Similar findings 318 were obtained in at least another recent study performed in MSM-PWIDs in San 319 Francisco¹⁷. In that study, it was suggested that an independent association with 320 sex work had emerged as a result of an "unmasking" effect, with the prevalence of 321 needle-sharing declining. This may also be a plausible hypothesis in the present 322 situation as needle/syringe-sharing in the SurvUDI network significantly decreased 323 from 1995 to 2014 (data not shown), with large-scale implementation of harm 324 reduction programs on the whole territory during that period. 325

326 In some studies, authors suggested that the association frequently observed between HIV transmission and crack use may be explained by higher-risk sexual 327 behaviours in crack users ^{28,31}. This is consistent with the present data where 328 participants of both sexes who reported sex work were also significantly more likely 329 to use non injected crack. Interestingly, men who reported sex work were 330 331 significantly more likely to report non injected use of amphetamines (similar observation in women, but not significant) and women who reported sex work were 332 more likely to use ecstasy. Regardless of whether they reported sex work or not, 333 334 sexually active men also reported frequent use of ecstasy. Men who reported sex work were slightly more likely to report methamphetamine use, but this was not 335 significant. The complex relationships between drug use and sexual behaviours have 336 been discussed previously ³². Amphetamine-like drugs, including ecstasy, are known 337 to be typically used during sexual activities and to be associated with unsafe sexual 338 activities ³². 339

340 The present study may provide several indications to target local public health 341 interventions. First, most participants who reported sex work are women, more likely 342 homeless and reporting frequent unsafe injecting as well as sexual behaviours. Several authors ^{9,33} have suggested that public health interventions should be 343 developed to reduce vulnerabilities among female PWIDs who engage in sex work. 344 A multilevel, combined approach has been proposed ⁹, which includes individual and 345 social harm reduction initiatives, biomedical, as well as structural interventions i.e 346 addressing homelessness, mental health and poverty issues as well as supporting 347 legal reform for sex workers to reduce the risk of bloodborne infections, violence and 348 homicide death ²⁹. Some authors also reported that HIV prevention programs may be 349 inadequate for MSM/PWIDs ^{26,27}. In the present study, men who reported client sex 350 partners had multiple male sex partners, but most of them said that their sexual 351 orientation was heterosexual or bisexual. Consequently, these men are very unlikely 352 353 to be reached by interventions targeting MSM whereas interventions targeting PWIDs may not be adapted to their high risk sexual exposure ^{26,27}. Despite the fact 354 that PWIDs/MSM/sex workers represent a very small population, targeting 355 356 interventions to this population should be a priority considering their potential for a high level of effectiveness and the possibility of bridging HIV and HCV epidemics 357 between PWIDs and MSM, as well as between male PWIDs reporting sex work and 358 their female sex partners ^{34,35}. 359

The data obtained through the SurvUDI network have some limitations. First, participants are not representative of all PWIDs. They are probably more representative of those who frequent harm reduction programs, where approximately 90% of participants were recruited. Moreover, participants who returned differ slightly from those with a single visit ¹³. Participants who returned may have more at risk behaviours, and this could have overestimated the association with sex work. Second, self reporting of behaviours may involve social desirability and recall biases that may lead to over-reporting of protective behaviours, like condom use, and underreporting of high risk behaviours, like syringe-sharing, possibly reducing the strength of the observed associations with HIV incidence.

In conclusion, further studies are needed to understand whether the independent association between HIV incidence and sex work is related to sexual transmission or other vulnerability factors, which may include mental illness, poverty and social exclusion, but increased risk due to sexual transmission cannot be excluded.

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FIGURE HEADINGS

Figure 1: Cumulative hazard of HIV seroconversion and HIV incidence according to baseline sexual activity, 2004-2014. Incidence rate calculation based on the sexual activity status reported at the second visit (baseline), i.e. exposure during the first time interval in the Cox proportional hazard model; Sexual activity is missing for 5 participants; Sexually active who did not report sex work; py: person-years.

Characteristics and behaviours n (%)	Men (n=1 147)	Women (n=378)	Total (n=1 528) ²	
Number of visits, median (min-max)	3 (2-18)	3 (2-14)	3 (2-18)	
Age, mean ± SD	37.8 ± 10.1	32.9 ± 9.5	36.6 ± 10.1	
Age (< 25 years)	113 (9.9)	88 (23.3)	201 (13.2)	
High school completed	563 (49.4)	203 (54.0)	769 (50.7)	
Urban recruitment region	1 027 (89.5)	322 (85.2)	1 352 (88.5)	
Homelessness	506 (44.2)	134 (35.6)	640 (42.0)	
History of incarceration	168 (14.7)	31 (8.2)	200 (13.1)	
Cocaine as the most often injected drug	608 (53.3)	173 (46.0)	783 (51.6)	
Injection with a needle/syringe used by someone else	234 (20.6)	110 (29.4)	344 (22.8)	
Daily injection ³	405 (35.4)	145 (38.5)	550 (36.1)	
Injection with strangers	402 (35.1)	134 (35.6)	536 (35.2)	
Time since first injection (\geq 6 years)	873 (76.2)	249 (66.4)	1 125 (73.9)	
Sexual activity				
No sex work ⁴	709 (61.9)	181 (48.3)	891 (58.5)	
Sex work	76 (6.6)	151 (40.3)	229 (15.0)	
Sexually inactive	360 (31.4)	43 (11.5)	403 (26.5)	

Table 1: Baseline¹ characteristics and behaviours of participants, 2004-2014

¹ Baseline characteristics based on information reported at the second visit, i.e. exposure during the first time interval in the Cox proportional hazard model.
 ² Sex is missing for 3 participants
 ³ In the last month
 ⁴ Sexually active who did not report sex work

	Crude HR	95% Cľ²	AHR	95% Cl ²
Model without effect modification				
Socio-demographic variables				
Sex (men vs women)	0.61	[0.37-1.02]	0.83	[0.45-1.51]
Injection behaviours				
Injection with a needle/syringe used by someone else	3.04	[1.87-4.93]	2.84	[1.73-4.66]
Sexual activity				
No sex work ³	1.00		1.00	
Sex work	2.81	[1.55-5.10]	2.19	[1.13-4.25]
Sexually inactive	1.48	[0.84-2.60]	1.62	[0.92-2.88]
Model with effect modification by sex ⁴				
Injection behaviours				
Injection with a needle/syringe used by someone else	3.04	[1.87-4.93]	2.91	[1.77-4.77]
Sexual activity				
No sex work	1.00		1.00	
Sex work, women	3.50	[1.26-9.72]	3.31	[1.19-9.21]
Sexually inactive, women	2.40	[0.57-10.04]	2.86	[0.67-12.01]
Sex work, men	1.87	[0.64-5.41]	1.64	[0.56-4.77]
Sexually inactive, men	1.37	[0.74-2.53]	1.47	[0.80-2.73]

Table 2: Multivariate analysis¹ of the associations between HIV incidence and sexual activity, 2004-2014

¹ Cox proportional hazard regression model; analysis performed using n=1 362 participants, excluding those with missing values for any of the independent variables ² 95 % confidence intervals ³ Sexually active who did not report sex work ⁴ p-value=0.5729 for effect modification, Wald chi-square test

HR: hazard ratio; AHR: adjusted hazard ratio

Characteristics and behaviours	No sex	Sex work	Sexually	p-value
n (%)	work ²	(n=51)	inactive	-
	(n=656)		(n=438)	
Age, mean ± SD	38.3 ± 10.1	35.6 ± 10.0	42.4 ± 9.7	<0.0001
Age (<25 years)	56 (8.5)	6 (11.8)	16 (3.7)	0.0030
Urban recruitment region	582 (88.7)	47 (92.2)	397 (90.6)	0.4936
Homelessness	263 (40.2)	30 (60.0)	175 (40.1)	0.0200
History of incarceration	87 (13.3)	14 (28.0)	59 (13.5)	0.0142
Drugs used by injection				
Cocaine	518 (79.1)	44 (86.3)	328 (75.1)	0.0975
Heroine	206 (31.5)	19 (37.3)	110 (25.2)	0.0366
Dilaudid® (prescribed or not)	296 (45.2)	27 (52.9)	207 (47.6)	0.4707
Cocaine as the most often injected drug	325 (49.9)	34 (68.0)	224 (52.0)	0.0450
Non-injected drugs				
Crack/freebase	420 (64.0)	43 (84.3)	255 (58.2)	0.0007
Amphetamines	241 (36.7)	28 (54.9)	102 (23.3)	<0.0001
Methamphetamine	41 (6.3)	5 (9.8)	18 (4.1)	0.1302
Ecstasy	127 (19.4)	11 (21.6)	29 (6.6)	<0.0001
Dilaudid® (prescribed or not)	163 (24.9)	12 (23.5)	74 (17.0)	0.0083
Daily injection ³	190 (29.1)	17 (34.7)	184 (42.2)	<0.0001
Time since first injection (≥ 6 years)	542 (82.9)	44 (86.3)	371 (84.9)	0.5975
Injection with strangers	178 (27.3)	24 (47.1)	110 (25.2)	0.0042
Always injected alone	188 (28.8)	13 (25.2)	185 (42.4)	<0.0001
Injection in public places	368 (56.2)	35 (68.6)	248 (56.6)	0.2219
Injection with a needle/syringe used by	119 (18.3)	12 (23.5)	62 (14.3)	0.0960
someone else	. ,	. ,	. ,	
Injection with needles/syringes used by	20 (3.1)	3 (6.0)	28 (6.5)	0.0291
someone else obtained mainly from		. ,	. ,	
strangers				
Number of female sex partners				
0	32 (4.9)	21 (41.2)	438 (100)	-
1	363 (55.3)	8 (15.7)	0	
2-5	224 (34.2)	15 (29.4)	0	
6-20	33 (5.0)	5 (9.8)	0	
≥21	4 (0.6)	2 (3.9)	0	
Number of male sex partners				
0	607 (92.5)	11 (21.6)	438 (100)	-
1	30 (4.6)	8 (15.7)	Ò	
2-5	14 (2.1)	11 (21.6)	0	
6-20	4 (0.6)	7 (13.7)	0	
≥21	1 (0.2)	14 (27.5)	0	
Condom use				
Consistent use for vaginal or anal sex	207 (32.2)	8 (17.0)	-	-
Inconsistent use for vaginal or anal sex	403 (62.7)	20 (42.6)	-	
Oral sex only	33 (5.1)	19 (40.4)	-	
Positivity for HCV antibodies	460 (70.1)	45 (88.2)	340 (77.6)	0.0012

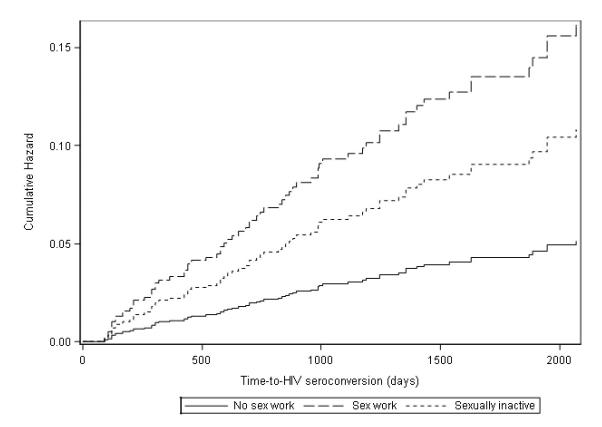
Table 3: Characteristics and, injection and sex behaviours according to reported sexual activity in men¹, 2004-2014

¹ Analyses performed using the last visit for non-seroconverters and the visit when HIV infection was first detected for seroconverters ² Sexually active who did not report sex work ³ In the last month NS : non-significant

Characteristics and behaviours n (%)	No sex work ²	Sex work (n=129)	Sexually inactive	p-value
	(n=198)		(n=46)	
Age, mean ± SD	33.5 ± 10.0	34.4 ± 8.6	40.5 ± 10.7	<0.0001
Age (< 25 years)	46 (23.2)	15 (11.6)	3 (6.5)	0.0031
Urban recruitment region	168 (84.9)	106 (82.2)	44 (95.7)	0.0838
Homelessness	59 (30.0)	53 (41.4)	11 (23.9)	0.0365
History of incarceration	11 (5.6)	18 (14.1)	1 (2.2)	0.0068
Drugs used by injection				
Cocaine	132 (66.7)	112 (87.5)	25 (54.4)	<0.0001
Heroine	75 (37.9)	49 (38.3)	10 (21.7)	0.0977
Dilaudid® (prescribed or not)	98 (49.8)	66 (51.6)	21 (46.7)	0.8479
Cocaine as the most often injected drug	80 (40.8)	71 (55.5)	15 (33.3)	0.0085
Non-injected drugs				
Crack/freebase	113 (57.1)	92 (71.3)	23 (50.0)	0.0091
Amphetamines	64 (32.3)	51 (39.5)	14 (30.4)	0.3337
Methamphetamine	8 (4.0)	3 (2.3)	3 (6.5)	0.4171
Ecstasy	29 (14.7)	29 (22.5)	4 (8.7)	0.0540
Dilaudid® (prescribed or not)	31 (15.7)	30 (23.6)	6 (13.3)	0.1289
Daily injection ³	76 (38.8)	55 (42.6)	15 (32.6)	0.4756
Time since first injection (≥ 6 years)	144 (72.7)	109 (84.5)	39 (84.8)	0.0216
Injection with strangers	51 (25.9)	49 (38.6)	3 (6.5)	0.0001
Always injected alone	38 (19.3)	23 (18.1)	18 (39.1)	0.0069
Injection in public places	98 (49.5)	74 (57.8)	18 (39.1)	0.0762
Injection with a needle/syringe used by	57 (28.9)	44 (34.9)	3 (6.5)	0.0011
someone else				
Injection with needles/syringes used by	6 (3.1)	11 (8.8)	0 (0.0)	0.0166
someone else obtained mainly from				
strangers				
Number of female sex partners				
0	169 (85.4)	105 (82.0)	46 (100)	-
1	20 (10.1)	11 (8.6)	-	
2-5	9 (4.6)	8 (6.3)	-	
6-20	0 (0.0)	4 (3.1)	-	
≥21	0 (0.0)	0 (0.0)	-	
Number of male sex partners				
0	11 (5.6)	0 (0.0)	46 (100)	-
1	136 (69.0)	4 (3.1)	-	
2-5	44 (22.3)	29 (22.5)	-	
6-20	5 (2.5)	34 (26.4)	-	
≥21	1 (0.5)	62 (48.1)́	-	
Condom use				
Consistent use for vaginal or anal sex	26 (13.3)	42 (32.8)	-	
Inconsistent use for vaginal or anal sex	164 (84.1́)	78 (60.9)	-	
Oral sex only	5 (2.6)	8 (6.3)	-	
Positivity for HCV antibodies	139 (70.2)	107 (83.0)	32 (69.6)	0.0252

Table 4: Characteristics and, injection and sex behaviours according to reported sexual activity in women¹, 2004-2014

¹ Analyses performed using the last visit for non-seroconverters and the visit when HIV infection was first detected for seroconverters ² Sexually active who did not report sex work; ³ In the last month NS : non-significant



	Number of participants	Seroconversions	Follow-up (py)	HIV incidence rate per 100 py [95 % confidence intervals]
Total	1 528	72	5 239.2	1.4 [1.1-1.7]
Sexual activity				
No sex work	891	25	2 965.1	0.8 [0.5-1.2]
Sex work	229	23	902.5	2.5 [1.5-3.6]
Sexually inactive	403	24	1 349.3	1.8 [1.1-2.5]