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Original Article

The Association of HIV and Syphilis among Men Who Have Sex with Men: An Integrated Biological and Behavioral Survey

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ABSTRACT

Increasing rates of syphilis in diverse geographical settings demand that people should have a heightened awareness of syphilis, especially in high-risk patients, including those who are MSM contracting HIV-positive. This study aimed to assess the association of HIV and syphilis among men who have sex with men (MSM) in Depok and Bogor using an Integrated Biological and Behavioral Survey 2018/2019. A cross-sectional study was conducted involving 750 eligible participants. Inclusion criteria: MSM living in study location, aged over 15, and had sexual relations with men at least once in the previous year. A prevalence ratio was obtained through multivariate Cox regression analysis. This study found that 11.3% of subjects were identified as having syphilis, and 20.1% were declared positive for HIV. The final model evidence a PR value of 2.46; 1.59-3.81 with a p-value of 0.00 for the risk of MSM infected with HIV to develop syphilis. Overall, HIV has a statistically significant relationship with the incidence of syphilis after being adjusted by the use of condoms with non-permanent partners and STI screening. Integrated intervention techniques, HIV testing, and enhanced awareness are required to reduce HIV, syphilis, and co-infection; targeting MSM and other vulnerable groups is in urge.

Keywords: HIV, Syphilis, MSM

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INTRODUCTION

Sexually transmitted infections (STIs) have significant implications for sexual and reproductive health, including stigmatization, infertility, cancer. and pregnancy complications, and an increased risk of contracting HIV. The World Health Organization (WHO) estimates that in 2020, there were approximately 374 million new STI cases worldwide, with 156 million cases of Trichomoniasis, 129 million cases of Chlamydia, 82 million cases of Gonorrhea, and 7.1 million cases of syphilis ¹. These infections pose a serious threat to public health and require effective preventative measures, such as comprehensive sex education, access to STI testing and treatment, and the use of barrier methods during sexual activity. Failure to address this issue could result in significant long-term health consequences for individuals and communities.

Syphilis is caused by the bacterium *Treponema pallidum*, which commonly spreads by direct contact with a syphilitic sore ². It developed to the secondary stage to include symptoms such as skin rashes and mucous membrane lesions. The disease rates are climbing rapidly, especially among gays, bisexuals, and men who have sex with men (MSM) ^{3–5}. Between 2020 and 2021,

reported primary and secondary cases increased around 29% in the United States, while the majority (36%) of these cases occurred only in MSM and men who have sex with men and women. Notably, the highest cases of syphilis among MSM occur in areas with HIV prevalence of more than 5% and lower-middleincome countries ⁵.

Numerous studies from diverse geographic locations documented that the clinical course of HIV was associated with increasing syphilis cases^{6,7}. A significant increase followed the increase in HIV rates in infections⁸. **Syphilis** syphilis causes inflammatory genital ulcers and lesions, which can increase the risk of HIV transmission by increasing HIV shedding and acquisition by providing a portal of entry to HIV⁹.

In Indonesia, the highest HIV prevalence recorded as of September 2022 was among men (71%), while the number of HIV cases in the MSM group reached 27.5% ¹⁰. Also, a previous study confirmed MSM with HIV infection are more likely to develop syphilis with a HR= 4.53; 2.24-9.17 ¹¹. A similar trend was also found in China, where the prevalence of syphilis is relatively high among HIV-positive MSM (14.9%; 12.7–17.2). These figures alert the increasing number of vulnerable groups against STIs, mostly syphilis.

Given the continued risk of syphilis transmission and its close association with HIV infection among MSM, this study addresses the issue using a large Integrated Biological and Behavioral Survey. Few studies have been conducted to determine the prevalence and risk factors of syphilis among MSM in Indonesia; however, a smaller sample size may be the limitation. This study aims to evaluate the prevalence of syphilis and determine the risk factors of syphilis among MSM in Depok and Bogor.

METHOD

A cross-sectional study was designed using a large Integrated Biological and Behavioral Survey (STBP 2018/2019). STBP study participants were 750 individuals identified as MSM living in Bogor and Depok. Men aged over 15 and who had sexual relations with men at least once in the last year were eligible for the study. Study participants completed a structured questionnaire about their sexual and health behaviors over time and provided biological samples at each study visit. The participants provided written informed consent before enrolling.

The purpose of this research is to examine the correlation between HIV status (as an independent variable) and the occurrence of syphilis in MSM (dependent variable). We also assessed the other associated factors, including age, education, employment, marital status, symptoms, Hepatitis B and Hepatitis C status, number of partners, having a permanent partner, consistency of condom use, circumcision, STI screening, STI treatment, information exposure, gathering behavior, and residence. All data was compiled from STBP, with HIV and syphilis variables specifically being obtained from the rapid test results.

Univariate analysis was used to summarize the descriptive frequency for each variable. Stratification analysis was used to determine confounding variables. Multivariate analyses were used in Cox regression to identify the estimation of HIV against syphilis infections after adjusting confounders. Factors that were determined as confounders (Prevalence ratio discrepancy $\geq 10\%$) in the stratification analysis were considered when computing the final model.

RESULTS

The demographic characteristics of STBP participants at baseline are described in Table 1 below.

Table 1.	Participants'	Demographic
Characteristic	S	
Characteristics	Frequency	y %
Age		
≤35	653	3 87,07
>35	97	7 12,93
Occupation		
Employed	550) 73,33
Unemployed	200) 26,67
Marital status		
Not married	659	9 87,87
Married	60) 8,00
Divorced	31	4,13
Education		
High	575	5 76,67
Low	175	5 23,33
The	overall partic	ipants were

The overall participants were predominantly aged ≤ 35 years, employed (73.33%), not married (87.87%), and had a higher educational background (76.67%).

The biological examinations (syphilis

status, HIV status, Hepatitis B, and Hepatitis C), and social behavior was displayed on table 2.

Table 2. Dist	tribution of beha	vioural
characteristics,	health access,	and
environment	,	
Variable	Frequency	%
Syphilis	Trequency	/0
Positive	85	11,33
Negative	665	88,67
HIV		
Positive	207	27,64
Negative	542	72,36
Syphilis symptoms		,
Detected	90	12,00
Undetected	660	88,00
Hepatitis B	000	88,00
Positive	38	5,07
Negative	712	94,93
Hepatitis C	/12	94,95
Positive	3	0,40
Negative	747	99,60
Number of partners	/ + /	<u> </u>
>1	232	31,18
$\frac{21}{0-1}$	512	68,82
Permanent partner	512	00,02
Yes	343	45,73
No	407	54,27
Selling sex to men	+07	57,27
Yes	261	34,80
No	489	65,20
Buying sex from men		05,20
Yes	75	10,00
No	675	90,00
Use of condoms	075	70,00
In a permanent partner	r	
Consistent	158	21,07
Inconsistent	185	24,67
N/A	407	54,27
Use of condoms in not		<u> </u>
permanent couples	-	
Consistent	136	18,13
Inconsistent	160	21,33
N/A	454	60,53
Use of condoms in		
transgender couples		
Consistent	7	0,93
Inconsistent	11	1,47
N/A	732	97,60
Circumcision		/
Yes	730	97,33

No	20	2,67
STI screening		
Never	591	78,80
Ever	159	21,20
STI treatment		
Never	90	56,60
Ever	69	43,40
Information exposure		
Unexposed	526	70,13
Exposed	224	29,87
Gathering behavior		
Yes	735	98,00
No	15	2,00
Residence		
With family/female	476	65,56
partner		
With a male	73	10,06
friend/partner/transvestite		
Alone	177	4,38

The results showed that 11% of the total 750 respondents were identified as having syphilis and 27% were declared HIV positive and based on the history of the symptoms experienced, most admitted that they did not show any symptoms of syphilis. In the results of hepatitis tests, both hepatitis B and C, <5% positive cases were found. From personal confessions, 68% have 0-1 partners, 54% do not have a permanent partner, more than 30% of respondents admit to selling commercial sexual services to men, and only 10% buy services from other men.

The results of assessing the consistency of condom use in different categories of partners show that only 21% consistently use condoms, both in regular and casual partners. Meanwhile, <1% of transgender sexual partners consistently use condoms. Moreover, almost 80% of respondents underwent circumcision. From the results of data collection on screening and treatment history, only around 21% had ever had an STI examination, and 43% had had STI treatment. Only a third of respondents had ever been exposed to information related to syphilis.

In terms of social behaviour, as many as 98% of respondents have the habit of gathering in various public facilities, and most of them admit to living with their family/partner. Woman/wife (65%).

		Syphilis (+) Syphilis (-)							- Prevalence Ratio (PR)			Homogonoite
Covariates	HIV	HIV (+) HIV (-)			HIV (+) HIV							Homogeneity – test
	n	%	n	%	n	%	n	%	Crude	Adjusted	$\Delta PR(\%)$	test
Age												
≤35	35	19,13	34	7,25	148	80,87	435	92,75	2,81	2,83	0,71	0,43
>35	9	37,50	7	9,59	15	62,50	66	90,41	(1,98-4,17)	(1,91-4,19)		
Occupation												
Employed	35	20,59	30	7,92	135	79,41	349	92,08	2,81	2,78	1,08	0,49
Unemployed	9	24,32	11	6,75	28	75,68	152	93,25	(1,89-4,17)	(1,87-4,13)		
Marital status												
Not married	38	20,11	39	8,32	151	79,89	430	91,68	2,81	2,75	2,18	0,23
Married	3	27,27	2	4,08	8	72,73	47	95,92	(1,89-4,17)	(1,86-4,07)		
Divorced	3	42,86	0	0,00	4	57,14	24	100				
Education												
High	40	23,12	35	8,73	133	76,88	366	91,27	2,81	2,66	5,6	0,95
Low	4	11,76	6	4,26	30	88,24	135	95,74	(1,89-4,17)	(1,79-3,95)		
Syphilis's symptoms												
Detected	8	22,22	3	5,66	28	77,78	50	94,34	2,81	2,84	1,06	0,58
Undetected	36	21,05	38	7,77	135	78,95	451	92,23	(1,89-4,17)	(1,90-4,24)		
Hepatitis B		,		,		,		,				
Positive	6	28,57	2	11,76	15	71,43	15	88,24	2,81	2,72	3,31	0,87
Negative	38	20,43	39	7,43	148	79,57	486	92,57	(1,89-4,17)	(1,82-4,05)		
Hepatitis C												
Positive	0	0,00	1	50,00	1	100	1	50,00	2,81	2,80	0,36	0,88
Negative	44	21,36	40	7,41	162	78,64	500	92,59	(1,89-4,17)	(1,89-4,15)	,	
No. of partners		,		,		,		,				
>1	11	25,58	10	5,29	32	74,42	179	94,71	2,81	2,68	4,85	0,10
0-1	33	20,25	31	8,91	130	79,75	317	91,09	(1,89-4,17)	(1,81-3,95)		
Permanent partner		,		,		,		,				
Yes	19	19,39	17	6,97	79	80,61	227	93,03	2,81	2,82	0,36	0,95
No	25	22,94	24	8,05	84	77,06	274	91,95	(1,89-4,17)	(1,90-4,18)	,	,
Selling sex to men		· · ·		- ,		,	-	- ,				
Yes	15	22,06	15	7,77	53	77,94	178	92,23	2,81	2,81	0,00	0,97
No	29	20,86	26	7,45	110	79,14	323	92,55	(1,89-4,17)	(1,89-4,17)	-,	
Buying sex from men		,-0		.,				. =,				
Yes	6	35,29	4	6,90	11	64,71	54	93,10	2,81	2,82	0,36	0,28
No	38	20,00	37	7,64	152	80,00	447	92,36	(1,89-4,17)	(1,90-4,17)	0,20	0,20

Table 3. The Association of HIV and Syphilis Stratified by Covariates

Use of condom in permanent partner												
Consistent	13	20,63	9	9,47	50	79,37	86	90,53	2,81	2,69	4,46	0,80
Inconsistent	6	17,14	8	5,37	29	82,86	141	94,63	(1,89-4,17)	(1,81-3,99)		
N/A	25	22,94	24	8,05	84	77,06	274	91,95				
Use of condom in non-permanent par	tner											
Consistent	8	21,05	9	9,18	30	78,95	89	90,82	2,81	2,58	8,9*	0,01*
Inconsistent	9	32,14	1	0,76	19	67,86	131	99,24	(1,89-4,17)	(1,76-3,77)		
N/A	27	19,15	31	9,94	114	80,85	281	90,06				
Use of condom in transgender couple	;											
Consistent	0	0,00	1	20,00	2	100	4	80,00	2,81	2,81	0,00	0,98
Inconsistent	0	0,00	1	11,11	2	100	8	88,89	(1,89-4,17)	(1,89-4,18)		
N/A	44	21,67	39	7,39	159	78,33	489	92,61				
Circumcision												
Yes	44	21,78	40	7,59	158	78,22	487	92,41	2,81	2,81	0,00	0,91
No	0	0,00	1	6,67	5	100	14	93,33	(1,89-4,17)	(1,89-4,16)		
STI screening												
Never	24	17,52	31	6,84	113	82,48	422	93,16	2,81	2,55	10,19*	0,99
Ever	20	28,57	10	11,24	50	71,43	79	88,76	(1,89-4,17)	(1,70-3,83)		
STI treatment												
Never	10	27,78	8	14,81	26	72,22	46	85,19	2,81	2,65	6,04	0,22
Ever	10	29,41	2	5,71	24	70,59	33	94,29	(1,89-4,17)	(1,29-5,39)		
Information exposure												
Unexposed	26	20,80	28	6,98	99	79,20	373	93,02	2,81	2,73	2,93	0,59
Exposed	18	21,95	13	9,22	64	78,05	128	90,78	(1,89-4,17)	(1,84-4,05)		
Gathering behavior												
Yes	43	21,50	40	7,49	157	78,50	494	92,51	2,81	2,80	0,36	0,49
No	1	14,29	1	12,50	6	85,71	7	87,50	(1,89-4,17)	(1,89-4,15)		
Residence												
With family/female partner	23	17,83	20	5,76	106	82,17	327	94,24	2,81	2,80	0,36	0,54
With a male	3	13,64	5	10,00	19	86,36	45	90,00	(1,89-4,17)	(1,88-4,08		
friend/partner/transvestite												
Alon	18	36,00	16	12,60	32	64,00	111	87,40				

*Confounders: $\triangle PR \ge 10\%$ or substantial justification

STI screening was considered a confounder in assessing the association of HIV with the incidence of syphilis. However, by considering the substance assumption, the variable consistency of the use of condoms in non-permanent partners is still included in the modelling even though the PR difference is <10%. It was also identified that there was a potential interaction between HIV and consistent use of condoms in casual partners (p-value <0.05) on syphilis.

After being included in multivariate modeling, it appears that the interaction variable is not significant enough to explain the difference in the risk of syphilis due to HIV between those who are consistent and inconsistent in using condoms in casual or non-permanent partners (p value> 0.05), so the interaction is ignored and the final model is obtained (Table 4).

Table 4. Final Adjusted Model of theAssociation of HIV with Syphilis

Variable	Adj.PR	95%	p-value		
		Lower	Upper		
HIV					
Positive	2,46	1,59	3,81	0,000	
Negative	Ref.				
Use of condo	oms in non	-permane	nt partner		
Consistent	Ref.				
Inconsistent	0,62	0,28	1,36	0,231	
N/A	1,06	0,61	1,83	0,837	
STI					
screening					
Never	0,62	0,39	0,98	0,042	
Ever	Ref.				

Table 4 shows an assessment of the relationship between HIV and the incidence of syphilis after controlling for the variables of condom use in casual partners and STI examination. Respondents who were identified as HIV positive had more than double the risk of experiencing syphilis compared to those who were HIV-negative.

DISCUSSION

This cross-sectional population-based study using an Integrated Biological and Behavioral Survey sample showed that the overall prevalence of syphilis among MSM in Depok and Bogor City was 11.3%. This prevalence of syphilis is lower than that reported in a previous study among MSM in Southeast Asia studies, which ranged between 12-14% ^{12,13}. However, our studies found various factors associated with an increased probability of syphilis infection. This information will be critical for the prevention of syphilis in Indonesia.

Syphilis is a growing public health concern, and the prevalence of syphilis is increasing among MSM globally ^{14,15}. This study addresses the tendency of a large number of HIV-positive MSM and explores the risk estimate by considering other covariates. The present study suggests that HIV-positive participants were associated with syphilis (PR= 2.46; 1.59-3.81) with a prevalence of 20.1% cases. The prevalence ratio was obtained after being adjusted by consistency of condom use with sexual non-permanent partners and STI screening. It is also important to highlight that these findings were slightly higher than the rate reported in a recent study in Jiangsu Province, which reported 13.64% of active syphilis infection 16.

Compared with heterosexual men, MSM is more likely to have a history of multiple STIs, including HIV, syphilis, gonorrhea. lymphogranuloma venereum (LGV), enteric STIs, HPV, human herpes virus (HHV-8), hepatitis B, and possibly hepatitis A and C. As the main focus of the study, the prevalence of active syphilis infection is even higher in HIV-positive MSM than in the general MSM population, and it ranges from 1.8 to 9.7%. This might be attributable to the social behavior reason. It is well established that the MSM is the most vulnerable group of people who are being at a high level of risk of HIV and STIs to unprotected anal due sex. heterosexuality, and multiple sex partners ^{17,18}.

Risky behaviors that put people at greater risk of contracting HIV include having anal or vaginal sex without a condom and suffering from other sexually transmitted infections (STIs) such as syphilis, herpes, chlamydia, gonorrhea, and bacterial vaginosis ¹⁹. Recently, in the United States, about half of MSM who suffer from primary and secondary syphilis are also living with HIV ²⁰. Skin sores caused by STDs such as syphilis allow HIV to enter the body more easily. In addition, it is possible to contract HIV because behavior and circumstances that put you at risk of contracting other STDs can also provide a greater risk of contracting HIV ²⁰.

Although some studies evidence the linkage between HIV infection and syphilis

incidence among gays, there is no definitive explanation for the pathway. Refugio and Klausner proposed a few hypotheses from the literature review ²¹. First, the increasing depletion of CD4 T-cells in persons with HIV reduces the host's ability to protect itself against infections ²². Second, it's also possible that HIV infection is simply a cover for engaging in syphilis-promoting behaviors. It has been claimed that the availability of highly active antiretroviral treatment (HAART) may have indirectly boosted syphilis incidence by increasing sexual risk behavior. Furthermore, HIV-infected MSM may engage in other riskreduction behaviors such as serosorting (i.e., sex partners are only HIV-infected men) or strategic positioning, in which the HIV-infected person assumes the receptive role during anal sex rather than the insertive role, which is more likely to transmit HIV²³.

Although this study prevents the risk of bias by controlling association estimation by several covariates, we acknowledged its limitations. Since it was a cross-sectional study, a chronological event between independent and dependent variables exists. This study cannot explain which cases were infected first, whether HIV infection or syphilis infection. Moreover, bias due to participants' responses is possible owing to the fact that this secondary data is collected by someone else, and we could not control over the data collection process.

CONCLUSION

Based on the results of this study, HIV is related to the incidence of syphilis in MSM. In this variable, STI testing is a confounding variable, and there is a potential interaction between HIV and consistent condom use in irregular partners. In the final model assessing the relationship between HIV and the incidence of syphilis after controlling for the variables of condom use in casual partners and STI examination, it was found that the risk of syphilis in HIV-positive respondents was 2.4 times. Risk factor screening in key populations as a way to prevent and control HIV and STIs, it is hoped that government institutions (Ministry of Health, District/City Health Services, Health Service Facilities, NGOs related to HIV and STIs) can work together to increase outreach to risk groups such as MSM.

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CONFLICTS OF INTEREST

The authors declare no conflict of interest

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