1 2	<u>Original Research Article</u> Seroprevalence of Syphilis among Human Immunodeficiency Virus Positive Individuals
3	Attending Immune Suppressed Syndrome Clinic at International Hospital, Kampala
4	Uganda
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12	Abstract
13	Background: Syphilis continues to be a persistent public health challenge and gaining renewed attention
14	against the back drop of HIV pandemic especially in the less developed high HIV stricken countries like
15	Uganda. This study enrolled 150 HIV infected individuals. The objective was to determine the syphilis
16	sero-prevalence and factors associated with syphilis infection among HIV positive individuals attending
17	immune suppressed syndrome (ISS) clinic at International Hospital Kampala -Touch Namuwongo Project
18	(TNP).
19	Methods/Design: This was a cross sectional study that recruited participants between January and May,
20	2014. Rapid plasma reagin test (RPR; BD Diagnostics) was used to screen for syphilis and if positive was
21	confirmed by the Treponema pallidum haemagglutination assay (TPHA; Biotec Laboratories Limited
22	Ipswich Suffolk, UK).

**Results**: We found 10% (n=15) seroprevalence of syphilis in the HIV positive individuals. Further; gender, age, occupation, marital status, polygamous relationship and education level attained did not show statistically significance association with syphilis infection (p>0.05). This prevalence was slightly higher in males (10.9%) than females (9.6%). The age groups 15-30 and 31-63 years had the highest (73%) and lowest (27%) seroprevalence of syphilis respectively.

28 Conclusion: Syphilis appears to be common amongst HIV infected individuals studied. We recommend
29 an urgent need to sensitize, screen and treat reproductively and sexually critical age group.

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Key words: *Treponema pallidum* Hemagglutination Assay, Human Immunodeficiency Virus, Rapid
 plasma reagin, Immune Suppressed Syndrome, Prevalence, Syphilis

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#### 34 **1. Introduction**

35 Syphilis is a sexually transmitted infection (STI) caused by a spirochete bacterium *Treponema pallidum*. It 36 is a multistage disease characterized by localized, disseminated and chronic forms of infection. An 37 estimated 12 million people globally have been infected; of which, almost two-thirds are in sub-Saharan Africa and south/southeast Asia [1]. In Uganda a report in 2009, human immunodeficiency virus (HIV) 38 39 and active syphilis prevalences were 28.8% and 4.3 %, respectively, and a high risk sexual behaviour was 40 frequently reported [2]. Some 35 million individuals globally are infected with HIV (0.8 % prevalence) 41 [3]. Sub-Saharan Africa remains at the epicenter of global HIV with prevalence of 25 million infected 42 individuals, and some 3.2 million children under age 15 are infected with HIV; Adult prevalence (15-49) 43 age group is 4.7% [3].

Because of a common their transmission route as well as the fact that they are mainly blood borne infections, syphilis and HIV co-infection continues to be a public health problem especially in the low income settings. More specifically, syphilis causes genital ulcer and facilitates HIV entry and shading. Besides, it induces immune activation and favor viral replication, which in turn accelerate HIV transmissibility [4, 5]. Syphilis has been a subject of intrigue and controversy since it was first recognized in the 15<sup>th</sup> century coined "the great imitator", it can manifest in a variety of ways depending on the host and stage of infection thus making diagnosis and management difficult [6].

In sub-Saharan Africa like Ethiopia, a combination of social stigma and associated underreporting, their asymptomatic nature, and lack of diagnostic facilities make the health and socioeconomic impacts of sexually transmitted infections unknown [7], this is no different in contexts like Uganda. The incidence of syphilis is rising all over the world, partly due to the increased transmission in HIV patients and other high risk groups such as men who have sex with men [7], the Ugandan scenario is predominantly due to the former.

57 This study was aimed at determining the seroprevalence and risk factors of syphilis among HIV–positive 58 individuals attending ISS clinic at International Hospital Kampala, Namuwongo Touch Project so that to 59 provide baseline data on the prevalence of syphilis in this population and guide their routine care and 60 management.

#### 61 **2. Materials and methods**

This was a cross-sectional study conducted at International Hospital Kampala (IHK), Touch Namuwongo Project (TNP), IHK is a private healthcare facility owned by the International Medical Group, the largest private healthcare group in Uganda and a teaching hospital for International Health Sciences University (IHSU), Kampala. Recruitment took place between January and May, 2014. The hospital is located in Namuwongo in Makindye Division, in southeast Kampala. Touch Namuwongo Project is a HIV/AIDS 67 prevention and treatment project based at International Hospital Kampala serving both urban and peri-68 urban HIV positive individuals.

The study population comprised all HIV–positive confirmed individuals attending the TNP. Being an exploratory study, our sample size was calculated to be 150 assuming syphilis prevalence of 11% amongst HIV positive individuals slightly higher than 9.8% syphilis prevalence in HIV–infected patients in Ethiopia [8], for our sample size estimation, we chose syphilis prevalence of 11% among HIV infected individuals because Uganda has more than three times HIV prevalence (7.4%) compared to Ethiopia (2.4%); we further assumed 95% level of confidence.

Individuals excluded from the study were children < 15 years of age, as they were fewer in number at</li>
the clinic as well as individuals already receiving syphilis treatment.

Sample testing was carried out according to the directions of the manufacturers and all tests were run against known positive and negative controls for quality assurance. Only those samples positive by both RPR and TPHA were considered to have syphilis [9]. A well trained certified counselor interviewed the study participants using structured questionnaires on offering written informed consent and other risk factors such as; gender, age, multiple sexual partner (more than one sexual partner), current condom use, marital status, education level, occupation, religion, excessive alcohol or drug use during sexual activity, and history of STIs were also collected.

Data was analyzed using SPSS Version–16, and results were summarized using descriptive statistics.
Bivariate cross-tabulations was performed on selected risk factors to find those significantly associated
with syphilis. A *p*-value < 0.05 was considered to be statistically significant. The study was approved by</li>
both Ethics Committee of the International Health Sciences University and International Hospital
Kampala.

## 90 3. HIV testing Algorithm 91 HIV diagnostic tests function either by detecting host antibodies made against different HIV proteins or by directly detecting the whole virus itself or components of the virus such as the HIV p24 antigen or HIV 92 93 RNA. 94 Figure 1. Algorithm for HIV screening and confirmatory test in Uganda 95 96 Determine <sup>™</sup> HIV 1/2 Ag/Ab 97 Positive **Negative (report)** 98 HIV 1/2 STAT-PAK<sup>®</sup> Assay 99 100 101 **Positive (report)** Negative **Uni-Gold HIV Rapid Test** 102 103 104 Negative Positive 105 Report Report

When screening for HIV, Unigold is used as a tie breaker between Determine<sup>®</sup> and Stat-Pak<sup>®</sup> since it is highly specific for the HIV specific antibodies. Inconclusive laboratory HIV results are always referred for Enzyme Linked Immunosorbent Assay (ELISA) and Polymerase Chain Reaction (PCR) at the reference

109 laboratory at Medical Research Council (MRC)/Uganda Virus Research Institute (UVRI) Uganda Research
110 Unit on AIDS, Entebbe, Uganda.

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## 112 **4.0 Results**

- 113 A total of 150 HIV patients who attended TNP-ISS clinic at International hospital Kampala were enrolled
- in the study. Table 1, our study found 15 individuals positive for syphilis amongst the HIV infected
- positive participants, 33% were males and 67% were females (Figure 3). We further found 73% of those
- 116 who tested positive for syphilis were 15-30 years age group and 27% were above 30 years. None of the
- selected risk factors was associated with syphilis infections, statistical significance at p < 0.05.

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## 119 **Table 1. Social demographic factors**

Variable	Frequency( <i>n</i> =150)	Percentage (%)	
Sex			
Female	104	69.33	
Male	46	30.67	

## Occupation

Business	30	20.00		
Driver	10	6.67		
House wife	21	14.00		
Teacher	11	7.33		
Maid	5	3.33		
No job	11	7.33		
Others	62	41.33		
Marital status				
Married	92	61.33		
Divorced	21	14.00		
Single	37	24.67		
Nature of Marriage				
Monogamy	73	48.66		
Polygamy	40	26.67		
None	37	24.67		

Education level			_
Secondary	41	27.33	
University/tertiary	87	58.00	
Primary	18	12.00	
None	4	2.67	

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121 Out of 150 study participants, 30.7% and 69.3% were male and females respectively. Further, Table 1.

122 Shows the distribution of the different social demographic factors of the participants. More than half

were married (61.33%), 48.66 % in monogamous relationships, and more than half (58.0%) had

124 university/tertiary education level.

125

## 126 Table 2. Laboratory Syphilis confirmatory results

Results	Frequency(n=150)	Percentage (%)	
Negative	135	90.00	
Positive	15	10.00	

## 129 We recruited a total of 150 HIV infected individuals and confirmed 15 (10%) syphilis seropositive cases

130 (co-infected).

## 131 Table 3. Shows bivariate analysis between sero-prevalence of syphilis and selected risk factors

Variable	Chi value	P value	
Gender	0.0557	0.813	
Occupation	8.7132	0.727	
Marital status	0.5805	0.748	
Nature of marriage	3.4523	0.178	
Education level	5.4106	0.248	

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133 We did not find significant statistical association between syphilis infection and gender, occupation,

134 marital status, and level of education (*p*>0.05), Table 3.

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141 Figure 2. A graph showing the age distribution of syphilis among the HIV patients attending ISS clinic



## 142 international hospital Kampala (Touch Namuwongo Project)

144 Our study participants were between 15 to 63 years. About seventy three percent of the syphilis



seropositive patients were below the age of 30 years.

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154 Figure 3. A pie-chart showing the gender distribution co-infected individuals

156 Fifteen patients tested positive for syphilis amongst the HIV infected individuals (coinfected), 33% were

157 males and 67% were females, twice the number of males (Figure 3).

167 Figure 4. A box plot showing the prevalence of syphilis among HIV patients by age

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# Sero-prevalence VS Age

Result

## 169

170 More than 50% of the HIV/Syphilis duo infected individuals were below the age of 30 years well as more

171 than 50% of syphilis sero-negative HIV infected individuals were above 30 years of age (Figure 4).

172

#### 174 Discussion

175 Syphilis remains an important STI in the era of HIV infection. Individuals infected with HIV are 176 vulnerable to many STIs amongst other opportunistic infections. Therefore identification of syphilis 177 cases in this population should be a priority addressed by HIV/AIDS control programs especially in low 178 income settings where HIV continues to be on the rise. Our study determined syphilis prevalence to be 179 10% amongst HIV –positive individuals (co-infection) at TNP-IHK. Further, none of the selected risk 180 factors was found to be statistically significantly associated with syphilis infections. In a systematic 181 review of literatures from several regions of the world, a median point-prevalence of syphilis among 182 HIV-infected patients was shown to be 9.5% [10], therefore our prevalence is slightly higher than the 183 global median.

From our study, n=150; there were more female (104) than male (46) participants; this would partially be due to that fact that TNP being an HIV clinic, more pregnant mothers were more likely to seek counseling so as to prevent mother to child transmission (PMTCT) of HIV infection therefore this could have led to a high number of female participants. The low number of male participants could be possibly explained by men's inherent behaviors and reluctance to seek medical care, this is further complicated by the fact that in many African societies men are the bread winners of the families.

The 15 who were positive for HIV and syphilis (co-infections) giving a prevalence of 10%. This is in agreement with the recent findings of Shimelis *et al* (2013) done in the Hospital of Ethiopia which gave a seroprevalence of 9.8% [8] probably this may be due to the fact that the two countries have different HIV prevalence levels 2.4% in Ethiopia as compared to Uganda 7.4% [11], three times higher than latter. However, this study disagrees with a previous one carried Asiki *et al* (2011) among fishing communities of Lake Victoria in Uganda which reported a prevalence of 4.3% [2]. This may be partially explained by the fact that fishing communities are characterized by complex socio-demographics. 197 The high prevalence of the co-infections in this study could have been due to the high sexual activity 198 with 60% of the participants between 23-30 years age group. Polygamous marriages accounted for 199 46.7% of the co-infected participants. The co-infection prevalence was greater in male (10.9%) than in 200 females (9.6%). This is in agreement with the findings of Shimelis et al (2013) where the prevalence in 201 males was 11% and females 8.9% in Ethiopia. However, we found no association between gender and 202 co-infections (P=0.813). The slightly high prevalence in males could have been due to polygamous 203 relationships and lack of awareness of syphilis, since 80% of the males positive for syphilis in this study 204 were in polygamous marriages.

Among other risk factors; none of these factors was statistically significant. Marital status (P=0.748),

Nature of marriage (P=0.178), Education level (P=0.248), Occupation (P=0.727). The non-significance of
 these factors could have been possibly due to fairly small sample size and the patients failing to admit
 some factors like commercial sex workers and having had unprotected sex which were deemed to have

been the main exposing factors to syphilis as stated by the world health organization in 2011.

210 Despite the fact that age was not significantly associated with syphilis (P=0.543), the prevalence was

highest in the age group (15-30) at 73 % and lowest in the age group (> 30). This is in agreement with a

study by Sarah in United States (2011) [12]. This could have been due to high sexual activity and

ignorance about syphilis in the youths where the prevalence was found to be high.

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#### 215 Conclusion

This draws an imperative need to sensitize adolescents about the risk of Syphilis infections and transmission since this age group coincidences with the onset of sexual activity and reproduction. The UNAIDS (2014) indicated that Uganda has an HIV infection prevalence of 8.0% among 15-49 years age group. This duo infection has a potential for vertical transmission therefore care and management of

220	HIV po:	sitive individuals should involve routine syphilis screening with emphasis on pregnant mothers.
221	Sensitiz	zation efforts should be carried out by HIV/AIDS control programs among the youths to ensure
222	that th	ey understand the risk factors of the co-infections and proper treatment of all individuals testing
223	positive	e for syphilis. The counselors and other health care workers in the different HIV clinics need to be
224	knowle	dgeable regarding syphilis symptoms, this can be part of continued medical education at
225	differe	nt levels to increase case detection of syphilis amongst this vulnerable population.
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227	Compe	ting interests
228	Author	s declare no competing interests
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231	Refere	nces
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