RESEARCH ARTICLE

**Rectal gonorrhoea and chlamydia among men who have sex with men in coastal Kenya [version 2; peer review: 1 approved, 1 approved with reservations]**

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**Abstract**

**Background:** Men who have sex with men (MSM) have a higher prevalence of *Chlamydia trachomatis* (CT) and *Neisseria gonorrhoeae* (NG) infections compared to the rest of the population, often remaining undiagnosed. In Kenya, prevalence of rectal CT and NG infection and NG antimicrobial sensitivity are poorly described.

**Methods:** MSM who reported receptive anal intercourse (RAI) were recruited from an ongoing human immunodeficiency virus acquisition and treatment study in coastal Kenya in 2016-2017. Rectal swabs were collected at two time points 6 months apart to estimate prevalence and incidence of CT/NG infection using a molecular point-of-care assay. Participants positive for CT or NG were treated according to national guidelines. NG culture and antimicrobial susceptibility testing was performed. Participant and risk behaviour characteristics were collected and association with baseline CT/NG prevalence assessed by multivariable regression analysis.

**Results:** Prevalence of CT/NG in 104 MSM was 21.2% (CT 13.5%, NG 9.6%, dual infection 1.9%) at baseline and 25.9% in 81 MSM at follow-up (CT 14.8%, NG 14.8%, dual infection 3.7%). CT/NG incidence was estimated at 53.0 (95% CI, 34.5-81.3) per 100 person-years. Most CT/NG positive participants were asymptomatic: 95.5% at baseline and 100% at follow-up. CT/NG infection was associated with being paid for sex [adjusted

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**Open Peer Review**

**Reviewer Status**

- Invited Reviewers
  - 1
  - 2

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odds ratio (aOR) = 6.2, 95% CI (1.7-22.9)] and being in formal employment [aOR = 7.5, 95% CI (1.1-49.2)]. Six NG isolates were obtained at follow-up; all were susceptible to ceftriaxone and cefixime and all were resistant to penicillin, tetracycline and ciprofloxacin.

**Conclusions:** There is a high prevalence and incidence of asymptomatic rectal CT and NG in MSM reporting RAI in coastal Kenya. MSM who were paid for sex or had formal employment were more likely to be infected with CT/NG suggesting increased risk behaviour during transactional sex. Antimicrobial susceptibility results suggest that current antibiotic choices in Kenya are appropriate for NG treatment.

**Keywords**
Chlamydia, Gonorrhoea, Men Who Have Sex With Men, Kenya, Cefixime, Azithromycin, Antimicrobial Susceptibility

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Introduction

*Chlamydia trachomatis* (CT) and *Neisseria gonorrhoeae* (NG) infections are curable sexually transmitted infections (STIs) that can have distressing symptoms and complications, though a large proportion remains asymptomatic. Global estimates of prevalence of genitourinary CT and NG infections from 2009 to 2016 were 2.7% and 0.7%, respectively, amongst men aged 15–49 years old. Men who have sex with men (MSM) are a key population with regards to transmission of STIs. Prevalence of rectal CT and NG infections has ranged as high as 21.7%–27.2% and 13.8%–25.4%, respectively in Italian and Thai MSM communities.

As a traditionally highly stigmatized population, the prevalence and incidence of STIs in MSM in sub-Saharan Africa has only been studied since 2005. Prevalence of rectal CT and NG has been estimated at 8% and 8%, respectively, in South African MSM above the age 18, 13.5% and 23.3%, respectively, in adult Nigerian MSM and 28.2% and 29.5%, respectively, in adult Tanzanian MSM (median age 23). In Kenya, we have estimated the prevalence of rectal CT and/or NG to be 11.6% in a small sample of 43 human immunodeficiency virus (HIV)-negative adult MSM in the coastal region and 5.2% in HIV-positive and HIV-negative MSM over the age of 18 in the western part of the country. In MSM, younger age, sex with men only (as opposed to sex with both men and women), transactional sexual intercourse, unprotected anal intercourse, and being HIV positive have been found to be associated with rectal CT/NG.

Over the years, the diagnosis and treatment of CT and NG infection has changed as a result of evolving technology and the development of antimicrobial resistance (AMR). Diagnosis is based on a combination of clinical suspicion, such as symptoms and signs suggestive of urethritis or proctitis, and laboratory investigations.

**Nucleic acid amplification testing (NAAT)**, both highly sensitive and specific in detecting CT and NG, has superseded the use of cultures for the detection of infection where available. In addition, with the advent of commercial point-of-care (POC) NAAT tests such as the GeneXpert® CT/NG assay (Cepheid), results can be obtained much more rapidly (90 minutes with the GeneXpert®).

The provision of same-day results is of particular importance in treating the MSM population, which is less likely to attend healthcare services. In the context of NG infection, however, bacterial culture is still recommended to enable detection of AMR.

Globally, AMR is a major threat to public health and resistance to the key antibiotic classes has been detected. In Kenya, plasmid-mediated resistance of NG to penicillin and tetracycline has been shown to be more prevalent than in other countries and is likely related to extensive use of doxycycline. The World Health Organization (WHO) currently recommends treating CT with either azithromycin or doxycycline. Where sensitivity is unknown, the WHO recommends treating NG with ceftriaxone or cefixime combined with azithromycin. In cases where sensitivity is known, a single agent can be used.

The WHO has recommended syndromic treatment of STIs as an answer to the low health-seeking behaviour and continuity of care typically seen in developing countries. In Kenya, first-line recommended treatment for proctitis is cefixime or ceftriaxone and azithromycin. Given higher prevalence amongst MSM, the WHO recommends presumptive treatment of MSM reporting unprotected receptive anal intercourse (RAI) in the previous six months and either a partner with an STI or multiple partners. Sensitivity of this approach has been estimated at 74.1% with a specificity of 45.8% in MSM in coastal Kenya. This study was performed to estimate the prevalence of asymptomatic CT and NG infections in MSM reporting RAI and to assess the susceptibility of NG to commonly used antimicrobials.

**Methods**

**Study design**

The study was conducted between April 2016 and January 2017 at the Kenya Medical Research Institute (KEMRI) Mtwapa Clinic, located in a town popular for its beach tourism and nightlife. A total of 104 MSM of 174 eligible participants were recruited from an ongoing cohort study of high-risk MSM. Participants not enrolled were more likely to report sex with men exclusively, transactional sex, RAI, group sex, and any unprotected sex (data in supplemental material). Men reporting RAI in the previous 6 months and aged between 18–49 years old were included for analysis. MSM were tested for CT/NG at baseline and at a follow-up visit 6 months later. Participants who tested positive were treated accordingly; CT infection was treated with azithromycin (2 g stat dose) and doxycycline (100 mg twice a day for 7 days), NG infection with cefixime (400 mg stat dose) and azithromycin (2 g stat dose) and dual infection treated with triple therapy (cefixime, azithromycin and doxycycline). A clinician observed the participant taking cefixime, azithromycin and the first dose of doxycycline.

**Collection of participant characteristics**

At baseline, counsellors conducted personal interviews with the participants. Men were considered to have sex with men exclusively (MSME) as opposed to both men and women.
(MSMW) based on answers they had given in previous clinic visits; if they had never reported having sex with women, they were considered MSME. Participants were asked if they had received money for or paid for sex in the 3 months prior to CT/NG sample collection. Condom use and number of sexual partners was also assessed. Clinicians recorded participant symptoms at the visit, and a participant was considered symptomatic of proctitis if he complained of rectal discharge, pain or an ulcer in the perianal region.

Sample collection and antimicrobial susceptibility testing for NG
At baseline and follow-up, a trained clinician collected rectal swabs in all participants using a proctoscope and dry GeneXpert® CT/NG swabs. The first swab was used to detect CT and NG infection by NAAT with the GeneXpert® CT/NG Assay (Cepheid AB, Sweden), and the second was inoculated on Thayer Martin Modified agar. Inoculated plates were transported to a reference laboratory 40 km away at the end of each day for culture and antimicrobial susceptibility testing of NG\(^2\). A case of NG or CT infection was defined as being a positive result on the GeneXpert® CT/NG assay.

The disc diffusion test was used to assess antimicrobial susceptibility of NG to ceftriaxone, penicillin and tetracycline\(^3\). The Etest was performed to obtain minimum inhibitory concentrations (MIC) for cefixime and ciprofloxacin; cut-off values, as follows, were obtained from the Clinical and Laboratory Standards Institute (CLSI)\(^3\). For cefixime, a MIC \(\leq 0.06 \mu g/ml\) indicated susceptibility. For ciprofloxacin, a MIC \(\leq 0.5 \mu g/ml\) indicated susceptibility, 0.12-0.5 \(\mu g/ml\) intermediate susceptibility, and \(\geq 1 \mu g/ml\) resistance.

Data analysis
Data were analysed using Stata 13.0 (StataCorp LP, College Station, TX). Baseline participant characteristics were described. Prevalence of NG and CT at baseline and follow-up was estimated, as were incidence rates of each infection. Univariable linear regression was used to estimate association between participant characteristics and prevalence of CT / NG infection at baseline. Any variable with a p-value of \(p<0.2\) was used in a multivariable regression analysis to determine adjusted odds ratios (aOR). Age was used a-priori in the multivariable regression, given that the population at the clinic is principally under 25 years of age and that STIs are more prevalent in younger age groups\(^3,31\).

Ethical approval
Ethical approval (protocol numbers 894 and 1224) was obtained from the KEMRI/Scientific and Ethics Review Unit (KEMRI/SERU). Written informed consent was obtained from every participant.

Results
Participant characteristics
Between April and July 2016, 104 participants were recruited and tested. Participant characteristics are described in full in Table 1. Approximately half (55.8%) of those recruited were between the ages of 25–34, approximately nine out of ten (86.5%) were never married, and approximately half (49.0%) received secondary education only. Half the participants (51.0%) were Christian one quarter were Muslim (26.0%) and 23.1% were either not religious or of a different religious background. One in four (25.0%) were unemployed, with the rest either self-employed (57.7%) or in formal employment (17.3%). One in four (25.0%) of the participants reported unprotected sex in the week preceding testing; with a similar number indicating they had sex with men exclusively (26.9%). Half the participants had been paid for sex in the 3 months prior to the study. Just over one in three (34.6%) MSM were HIV positive.

Prevalence of CT/NG
At baseline, 22 (21.2%, 95% CI 13.8-30.3%) of the 104 MSM who took part in the study tested positive for CT and/or NG; 10 had NG (9.6%) and 14 had CT (13.5%)- 2 (1.9%) had dual infection. Only one participant reported symptoms whilst the rest (95.5%) were asymptomatic. The symptomatic participant was one of the two participants co-infected with both CT and NG. At baseline, none of the 104 swabs taken for culture, including those from the ten GeneXpert® NG positive participants, grew NG.

The prevalence of CT and/or NG infection was associated with receiving payment for sex \(\) (adjusted odds ratio (aOR)=6.2, 95% CI (1.7-22.9]) and being in formal employment \([aOR=7.5, 95\% CI (1.1-49.2)]\) on multivariable logistic regression (Table 1), in a model controlling for age, having sex with men only or both men and women (MSME or MSMW, respectively), employment status and receiving payment for sex in the previous three months. MSME was also associated with greater odds of prevalent CT/NG, though this did not reach statistical significance \([aOR=2.7, 95\% CI (0.7-7.1)].\)

Of the 104 participants included in the baseline analysis, 20 (19.2%) participants were lost to follow-up; 3 (2.9%) were missing a test at 6 months. Baseline CT/NG prevalence in these 23 participants was 31.8% (30.4%) compared to 18.5% in those who attended follow-up, although this difference was not statistically significant \((p=0.217)\). Twenty-one (25.9%) of 81 participants at follow-up tested positive for CT and/or NG infection; 12 (14.8%) participants had CT, 12 (14.8%) had NG, and 3 (3.7%) had dual infection. Five of the twenty-one (23.8%) cases of CT or NG that tested positive for CT or NG at follow-up were also positive at baseline; 5 had CT and 1 NG (1 CT and NG co-infection). All 21 follow-up participants were asymptomatic. NG was isolated from 50% (six) of twelve GeneXpert® NG positive follow-up participants.

Incidence of CT/NG
The 81 MSM who contributed data towards incidence calculation were followed up for a median of 5.7 months (interquartile range, 5.4-6.2 months), which amounts to a total follow-up time of 39.6 person-years. Overall, 21 MSM tested positive for CT or NG for an incidence rate of 53.0 (95% CI, 34.5-81.3) per 100 person-years (Table 2). Of these, 12 tested
Table 1. Risk factors associated with *Chlamydia trachomatis* (CT) and/or *Neisseria gonorrhoeae* (NG) infection in 104 men who have sex with men (MSM) at baseline.

<table>
<thead>
<tr>
<th>Variable</th>
<th>All, n (% of total)</th>
<th>CT/NG, n (%)</th>
<th>OR, (95% CI)</th>
<th>p value</th>
<th>AOR (95% CI)</th>
<th>p value</th>
</tr>
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<td></td>
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<td>18–24</td>
<td>36 (34.6)</td>
<td>10 (27.8)</td>
<td>0.9 (0.2-4.2)</td>
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<td>0.8 (0.1 – 5.7)</td>
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<td>3 (30.0%)</td>
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<td>Higher/tertiary</td>
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<td>19 (21.1)</td>
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<td>Ever married</td>
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<td>3 (21.4)</td>
<td>1.0 (0.3-4.0)</td>
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<td>Muslim</td>
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<td>Self</td>
<td>60 (57.7)</td>
<td>13 (21.7)</td>
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<td>1.8 (0.4-7.8)</td>
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<td>18 (17.3)</td>
<td>6 (33.3)</td>
<td>3.8 (0.8-18.1)</td>
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<td>7.5 (1.1-49.2)</td>
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<td>Received payment for sex in past 3 months‡</td>
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<td>No</td>
<td>97 (93.3)</td>
<td>21 (21.7)</td>
<td>Reference</td>
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<td>Yes</td>
<td>7 (6.7)</td>
<td>1 (14.3)</td>
<td>0.6 (0.1-5.3)</td>
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<td>Paid for sex in past 3 months</td>
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<td>No</td>
<td>97 (93.3)</td>
<td>21 (21.7)</td>
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<td>Yes</td>
<td>7 (6.7)</td>
<td>1 (14.3)</td>
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<td>Alcohol use in past month</td>
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<td>12 (19.4)</td>
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<td>No activity</td>
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<td>6 (17.7)</td>
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<td>All protected</td>
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<td>8 (18.2)</td>
<td>1.0 (0.3-3.3)</td>
<td>0.951</td>
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<td>Any unprotected</td>
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<td>2.1 (0.6-7.0)</td>
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<td>Condom use for anal sex in past 3 months‡</td>
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<td>18 (24.0)</td>
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<td>75 (72.1)</td>
<td>18 (24.0)</td>
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<td>Yes</td>
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<td>4 (15.4)</td>
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<td>15 (14.4)</td>
<td>3 (20.0)</td>
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<td>One</td>
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<td>0.4 (0.1-2.9)</td>
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<td>Negative</td>
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<td>15 (22.1)</td>
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<td>0.9 (0.3-2.3)</td>
<td>0.756</td>
<td></td>
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</tr>
</tbody>
</table>

*Included in the multivariable analysis priori. †Only factors significant at p<0.2 in the univariable analysis were included in the multivariable model. ‡3 participants did not report anal sex in the last 3 months but reported within the past 6 months. MSME, MSM exclusively; MSMW, MSM and women. OR, odds ratio; aOR, adjusted odds ratio; CI, confidence interval.
positive for CT [incidence rate of 30.0, 95% CI (17.2-53.3)] including 5 who tested positive for CT at baseline [incidence rate of 109.4, 95% CI (45.5-262.7)] and 7 who tested negative for CT at baseline [incidence rate of 20.0, 95% CI (9.5-41.9)], p=0.009. From the 21 patients who tested positive for CT or NG at follow-up, 12 were positive for NG [incidence rate of 30.0, 95% CI (17.2-53.3)], including 1 who tested positive for NG at baseline [incidence rate of 25.5, 95% CI (3.6-181.1)] and 11 who tested negative for NG at baseline [30.8 (17.1-55.6)], p=0.950.

Antimicrobial susceptibility of N. gonorrhoeae
Out of the 185 cultures performed (104 at baseline, 81 at follow-up), only six (3.2%) isolated NG; all six had been detected by the GeneXpert® NAAT and were isolated from follow-up samples. All 6 NG isolates were found to be 100% resistant to tetracycline and ciprofloxacin. Of the 6, 2 were partially resistant and 4 had intermediate susceptibility to penicillin. All 6 were sensitive to ceftriaxone and cefixime (Table 3).

Discussion
This study assessed the prevalence and incidence of rectal CT and NG in 104 MSM, identifying one in five participants at baseline and one in four participants at follow-up who were infected with either or both infections, and an estimated CT/NG incidence of 53.0 per 100 person years. Similarly high prevalence and incidence of rectal CT/NG infections were reported from other parts of sub-Saharan Africa\(^3\)\(^-\)\(^5\). Prevalence at baseline was statistically associated with receiving money for sexual intercourse, a four-fold increase in odds, which is consistent with the findings from previous studies\(^1\)\(^-\)\(^3\)\(^,\)\(^12\). Men who engage in transactional sex may have increased vulnerability to STIs, which is compounded by an increased burden of psychosocial morbidities, such as stigmatization and discrimination\(^3\). Being in formal employment was also associated with prevalent infection with CT/NG, possibly indicating that those with stable employment and, presumably, financial means engage in more transactional sexual activity; although there was no statistically significant association between paying for sex and rectal CT/NG prevalence.

The WHO recommends that CT/NG infections should be detected and treated in a timely manner, by either presumptive or syndromic treatment\(^2\)\(^-\)\(^5\). This is particularly important as these infections can remain undiagnosed for a long time unless routinely screened for, and thus be easily transmitted to other sexual partners. In this study, 95.5% of infections at baseline and 100% of infections at follow-up were asymptomatic. There are more asymptomatic cases of anorectal CT/NG reported in this study compared to other studies on MSM from the USA (58% asymptomatic CT, and 69% asymptomatic NG)\(^3\) and Kenya (58.3% asymptomatic CT/NG infection)\(^1\). Larger research studies are needed to confirm these findings. Nevertheless, this does support the need for more structured screening programmes to detect asymptomatic infections in developing countries, particularly in high-risk populations such as MSM\(^3\)\(^-\)\(^5\).

No NG was isolated from baseline rectal swabs in GeneXpert® NG-positive participants. At follow-up, six NG isolates were successfully cultured from twelve GeneXpert® NG positive participants. The only methodological differences at follow-up was prior notification of the laboratory at the reference centre 40 km away that a participant was positive on the GeneXpert® assay and one extra day of incubation to give more time to the scanty colonies to grow. It is well

### Table 2. Incidence of rectal CT and/or NG infections among 81 MSM, Mtwapa, Kenya.

<table>
<thead>
<tr>
<th>Participants at baseline and follow up, N=81</th>
<th>Cases/person-years</th>
<th>Incidence per 100 person-years (95% CI)</th>
<th>P-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Any CT/NG</td>
<td>21/39.6</td>
<td>53.0 (34.5-81.3)</td>
<td>n/a</td>
</tr>
<tr>
<td>Infection at baseline</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>No, N=66</td>
<td>15/32.1</td>
<td>46.7 (28.2-77.5)</td>
<td>-</td>
</tr>
<tr>
<td>Yes, N=15</td>
<td>6/7.5</td>
<td>79.8 (35.9-177.6)</td>
<td>0.281</td>
</tr>
<tr>
<td>Any CT</td>
<td>12/39.6</td>
<td>30.3 (17.2-53.3)</td>
<td>n/a</td>
</tr>
<tr>
<td>Infection at baseline</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>No, N=72</td>
<td>7/35.1</td>
<td>20.0 (9.5-41.9)</td>
<td>-</td>
</tr>
<tr>
<td>Yes, N=9</td>
<td>5/4.6</td>
<td>109.4 (45.5-262.7)</td>
<td>0.009</td>
</tr>
<tr>
<td>Any NG</td>
<td>12/39.6</td>
<td>30.3 (17.2-53.3)</td>
<td>n/a</td>
</tr>
<tr>
<td>Infection at baseline</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>No, N=73</td>
<td>11/35.7</td>
<td>30.8 (17.1-55.6)</td>
<td>-</td>
</tr>
<tr>
<td>Yes, N=8</td>
<td>1/3.9</td>
<td>25.5 (3.6-181.1)</td>
<td>0.950</td>
</tr>
</tbody>
</table>

CT: Chlamydia trachomatis; NG: Neisseria gonorrhoeae; CI: Confidence interval
recognized that test methodology and technical expertise have an impact on the positivity rates of samples. Hence, antimicrobial susceptibility testing may be more reliably evaluated using rapid molecular POC tests in the future\(^6\). The low yield in this case may also be due to the practicalities of having to transport the culture plates to the reference laboratory and the order of swab collections. As the culture swab was taken last, the bacterial yield may have been insufficient to result in a positive culture.

Regardless of yield, antimicrobial susceptibility testing revealed that all cultured NG isolates were sensitive to cefixime (first line treatment for NG) and ceftriaxone, and that all isolates were resistant to tetracycline and ciprofloxacin (Table 3). Though azithromycin sensitivity was not assessed and baseline cultures were negative, the susceptibility of NG to cefixime at follow-up suggests that all treated participants were cured. Previous susceptibility studies of NG have shown that AMR is rising globally, and is a very real threat to healthcare systems, particularly for those caring for high-risk populations\(^2,5,21,22\). Genomic studies have demonstrated that resistant NG strains are transmitted between MSM partners\(^7\). This information calls for careful treatment of NG, and for routine antimicrobial susceptibility testing to be performed. With the given WHO recommendations of syndromic and presumptive treatment, it is possible that AMR may spread further as an increasing amount of studies highlight the ongoing high rates of resistant NG in MSM. It is therefore critical that future studies systematically review all existing data on prevalence, incidence and gonococcal AMR, and that global guidelines on STI treatment in MSM and other key populations are updated to take into account AMR and POC testing\(^2,23\).

Approximately a third of participants in this study (34.6%) were HIV positive, though this was not statistically associated with CT/NG prevalence. HIV infection has been found to be higher in participants with an underlying STI, presumably facilitated by inflammation of mucosal epithelium\(^6,8\). It is therefore critical to rapidly detect and treat STIs in order to minimize future HIV-related epidemics in the MSM populations\(^8\). Several studies have looked at the association between pre-exposure prophylaxis (PrEP) and the incidence of STIs; these studies demonstrated increased rates of CT/NG infections with PrEP use- potentially related to increased screening for STIs- increased condomless sexual intercourse and decreased serosorting between sexual contacts\(^1,4,12\). Several months after this study concluded, HIV-negative MSM in the same larger cohort started receiving PrEP; it will be interesting to assess the impact this has on CT/NG infection rates in this population.

This study has several limitations. First off, the sample size was small, rendering statistical analysis and interpretation of results for subgroups difficult. Due to the small sample size, we combined CT/NG infections to improve statistical power. As a result, we are unable to comment on predictors of CT and NG separately. A large number of eligible patients were not enrolled in the study, and from brief comparison those patients had higher reports of sexual risk behavior compared to enrolled participants (see supplemental data). Therefore, our CT and NG prevalence and incidence estimates may be underestimates of a true population prevalence and incidence. Additionally, though all cases of CT and NG were treated at baseline, a test of cure was not performed: it is not possible for us to ascertain whether all cases at follow-up were new cases (reinfection) or whether some may have been persistent cases. Furthermore, our study did not assess azithromycin resistance. Any future studies should assess this given the frequent use of azithromycin in syndromic treatment of rectal infections. Finally, the small yield of NG cultures highlights the difficulty of generalizing resistance data, and this calls for larger studies of antimicrobial resistance in Kenya.

### Conclusion

The prevalence of rectal CT/NG infections in MSM who report RAI in coastal Kenya is estimated at 21.2%, and the incidence rate at 53.0 per 100 person years. Baseline infections were associated with transactional sex and formal employment. All but one of the participants who tested positive for CT/NG infection were asymptomatic, supporting the use of a presumptive treatment approach. Gonococcal AMR is of serious concern, and thus there is a need for continued surveillance of NG antimicrobial susceptibility and for sharing of AMR data between research groups studying NG in Kenya.

<table>
<thead>
<tr>
<th>Isolate</th>
<th>Penicillin (inhibitory zone in mm)</th>
<th>Ceftriaxone (inhibitory zone in mm)</th>
<th>Tetracycline (inhibitory zone in mm)</th>
<th>Cefixime (MIC in µg/ml)(^a)</th>
<th>Ciprofloxacin (MIC in µg/ml)(^a)</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>32</td>
<td>45</td>
<td>14</td>
<td>0.016</td>
<td>4</td>
</tr>
<tr>
<td>2</td>
<td>8</td>
<td>36</td>
<td>18</td>
<td>0.016</td>
<td>3</td>
</tr>
<tr>
<td>3</td>
<td>0.19</td>
<td>45</td>
<td>13</td>
<td>0.016</td>
<td>16</td>
</tr>
<tr>
<td>4</td>
<td>0.25</td>
<td>40</td>
<td>14</td>
<td>0.016</td>
<td>4</td>
</tr>
<tr>
<td>5</td>
<td>0.19</td>
<td>45</td>
<td>19</td>
<td>0.016</td>
<td>2</td>
</tr>
<tr>
<td>6</td>
<td>0.024</td>
<td>45</td>
<td>16</td>
<td>0.016</td>
<td>4</td>
</tr>
</tbody>
</table>

\(\alpha\): MIC ≤0.5µg/ml indicates susceptibility

\(\beta\): MIC of ≤0.06µg/ml indicates susceptibility, 0.12-0.5µg/ml intermediate susceptibility, and ≥1µg/ml resistance

Table 3. Antimicrobial susceptibility of rectal *Neisseria gonorrhoeae* isolates.
**Data availability**

Underlying data

Figshare. Data Sharing Excel.xlsx. https://doi.org/10.6084/m9.figshare.7735001.v1. This file contains infection status for each participant, alongside details of risk factors and de-identified demographic information.

Extended data

Figshare. Data Sharing Worksheet_CT_NG.xlsx. https://doi.org/10.6084/m9.figshare.9896396.v1. This file contains information regarding variable names for above dataset.

This file contains supplemental data comparing characteristics of eligible patients not recruited versus those that were.

Data are available under the terms of the Creative Commons Attribution 4.0 International license (CC-BY 4.0).

**Acknowledgements**

The authors would like to acknowledge the International AIDS Vaccine Initiative for funding the project, KEMRI-Wellcome Trust Programme for allowing the project to be carried out at their facility, Kenyatta University for accepting and supervising this work as part of the MSc thesis and Dr Rohini Manuel, consultant clinical microbiologist, for expert input and review of the final version of this manuscript.

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**References**


for Prevention, Management and Control of Sexually Transmitted Infections. 2019. (September).


Open Peer Review

Current Peer Review Status: ✔️ ❓

Version 2

Reviewer Report 27 December 2019

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Nicola Low
Institute of Social and Preventive Medicine, University of Bern, Bern, Switzerland

Thank you for giving me the opportunity to review the revised version of this manuscript. The authors have done a good job of responding to almost all of my comments. There are a couple of outstanding issues, which I hope the authors will be able to address.

1. Reporting of response rate and comparison between responders and non-responders: thank you for describing the enrolment more completely. Unfortunately, the main text and the response are now contradictory. The main text says that 104 men were enrolled out of a possible 174 (60%). In the response, the authors say that “We were able to enrol 70 of the 174 eligible men (i.e. 40%). We have added a supplemental table…” The supplementary material is the Stata output and the definitions of the value labels for participation (0 or 1) aren’t given. It would help to summarise the Stata output in a table, with “Enrolled” or “Not enrolled”.

2. Antimicrobial treatment: please can you say which guidelines you followed for treatment (are they Kenyan guidelines) because the treatment given, whilst likely to be effective, is not the same as the WHO recommendations, which you cite (your refs 22 and 23).

3. Antimicrobial resistance: thank you for tabulating the findings of antimicrobial susceptibility testing. This is much more useful. Could you please now clarify the interpretation of the disc diffusion results:
   - The ref you give, Liu H et al. is for cephalosporins only. That paper says an inhibitory zone of >=35mm is susceptible – can you add that this is the cut-off?
   - The penicillin values look, to me, more like MIC value, e.g. ranging from 32 to 0.024 – how could 0.024 be read in mm? Please clarify.
   - Please give a ref and a cut-off for tetracycline susceptibility.

4. Data analysis and Table 1: I requested global p-values from a likelihood ratio test. I understand your reasons for wanting to present stratum specific results. My understanding is that, in the multivariable model, you want to compares the model fit for the whole variable. The likelihood ratio test gives this information. You can then show, using the point estimate and its confidence interval that the adjusted OR in one stratum is greater, or smaller, than the reference category. I am not
going to insist, but in any case, please remove the bold for p-values. It is unnecessary.

5. Abstract – you could remove the numbers about prevalence at follow up. It would be more useful to give the confidence intervals for the prevalences that you state.

**Competing Interests:** No competing interests were disclosed.

**Reviewer Expertise:** Epidemiology, bacterial sexually transmitted infections, antimicrobial resistance.

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard, however I have significant reservations, as outlined above.

Author Response 13 Feb 2020

Caroline Ngetsa, KEMRI-Wellcome Trust Research Programme, Kilifi, Kenya

Dear Reviewer,

Many thanks for your careful review of our revised manuscript and alerting us to few outstanding issues. Below, we have responded to your comments and indicated changes that we requested to be made to the manuscript.

- Reporting of response rate and comparison between responders and non-responders: thank you for describing the enrolment more completely. Unfortunately, the main text and the response are now contradictory. The main text says that 104 men were enrolled out of a possible 174 (60%). In the response, the authors say that “We were able to enrol 70 of the 174 eligible men (i.e. 40%). We have added a supplemental table…”

There was an error in our response letter. We confirm that the manuscript text is correct. The supplemental table also presented the correct numbers.

The supplementary material is the Stata output and the definitions of the value labels for participation (0 or 1) aren’t given. It would help to summarise the Stata output in a table, with “Enrolled” or “Not enrolled”.

We have changed the labels for participation (0 or 1) to (Not enrolled or Enrolled) on the supplemental table and the reference 45 has also been updated with these corrections.

- Antimicrobial treatment: please can you say which guidelines you followed for treatment (are they Kenyan guidelines) because the treatment given, whilst likely to be effective, is not the same as the WHO recommendations, which you cite (your refs 22 and 23).

Thank you for picking up on this omission. Our STD treatment followed from a 2015 Kenyan guidelines revision that initially proposed 2 gram azithromycin stat. The Kenyan guidelines have since aligned with WHO guidelines and changed to 1 gram azithromycin stat. We erroneously reported the old treatment regimen but have corrected it now in the manuscript (1 gram azithromycin as per Kenyan and WHO guidelines).

- Antimicrobial resistance: thank you for tabulating the findings of antimicrobial susceptibility testing. This is much more useful. Could you please now clarify the interpretation of the disc diffusion results:

We have updated the table with the antimicrobial sensitivity results with interpretation for both the disc diffusion and MIC results.

The ref you give, Liu H et al. is for cephalosporins only. That paper says an inhibitory zone of >=35mm is susceptible – can you add that this is the cut-off

We have added the cephalosporin cut off >=35mm in the table. We have also included cut offs for the other drugs listed in the table.
The penicillin values look, to me, more like MIC value, e.g. ranging from 32 to 0.024 – how could 0.024 be read in mm? Please clarify.

We acknowledge that this was an error in the column label. We have updated the column label to indicate ‘MIC in µg/ml’

Please give a ref and a cut-off for tetracycline susceptibility.

Ref: Performance Standards for antimicrobial susceptibility test; CLSI 2019 guidelines.

Cut-off for Tetracycline >=38mm is susceptible.

Data analysis and Table 1: I requested global p-values from a likelihood ratio test. I understand your reasons for wanting to present stratum specific results. My understanding is that, in the multivariable model, you want to compare the model fit for the whole variable. The likelihood ratio test gives this information. You can then show, using the point estimate and its confidence interval that the adjusted OR in one stratum is greater, or smaller, than the reference category. I am not going to insist, but in any case, please remove the bold for p-values. It is unnecessary.

Abstract – you could remove the numbers about prevalence at follow up. It would be more useful to give the confidence intervals for the prevalences that you state.

Thank you for your consideration with regards to Table 1, and the suggestion to remove the prevalences at time point 2 in the abstract. We consider removing the bolding of p-values a minor edit and proposed abstract edits of minorly importance and prefer to leave this as is.

We would like to thank you for making these excellent review points helping us to improve communication about our work. We trust that we have addressed these additional points and hope that our manuscript will meet your approval.

Competing Interests: No competing interests

Version 1

Reviewer Report 13 August 2019

https://doi.org/10.21956/wellcomeopenres.16606.r36001

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Nicola Low

Institute of Social and Preventive Medicine, University of Bern, Bern, Switzerland

Thank you for giving me the opportunity to review this manuscript, which addresses the important issue of extra-genital sexually transmitted infections (STI) in men who have sex with men (MSM) in Kenya. I have no concerns about the relevance and validity of the data. The study is small but, on the other hand, has detailed behavioural information, antimicrobial susceptibility data and a follow-up visit. The findings lack statistical precision, so the authors are restricted in the certainty of the conclusions that they can draw. In this situation, data from such studies are very important for future syntheses of evidence in systematic
In Wellcome Open Research, I think the authors have more flexibility about what to report, meaning that there is an opportunity for useful contextual information about the study and for full reporting that will make the findings reproducible and able to be used in a future systematic review. I have some suggestions that I hope the authors will find useful in revising their manuscript.

**Major comments**

1. **Study design:** this is a report of a cohort study, nested within a larger cohort study. Please make sure that all the items proposed in the STROBE (Strengthening the Reporting of Observational studies in Epidemiology) checklist for cohort studies (https://www.strobe-statement.org/index.php?id=available-checklists) are reported. In particular, I would like to see: a) the proportion enrolled and whether or not they differed from the men in the larger cohort study. The authors only report (p4, Results, Participant backgrounds, line 1), “104 participants were recruited and tested”; b) the follow-up time reported as a total number of months and average for each participant; 95% confidence intervals (CI) for the main percentages.

2. **Study design:** Can the authors say whether this study was pre-planned? If so, was there a target sample size with a power calculation? Was enrolment less than expected?

3. **Data analysis:** can the authors please calculate and report the incidence rate of *Chlamydia trachomatis* (CT) and *Neisseria gonorrhoeae* (NG) detection in anorectal samples? The availability of data from a follow-up visit is a strength of the study. Prevalence at follow-up (p4, Methods, Data analysis, line 3, and Table 2) isn’t very useful. It would be great to change the columns in Table 2 to report: numbers with positive results at baseline; the numbers of incident cases and follow-up time; incidence rate (with 95% CI) in those with a negative test result at baseline; repeat detection rate (with 95% CI) in those with a positive test at baseline who received treatment.

4. **Data analysis** (p4, Methods, Data analysis, line 11-13): Please delete the sentence with “…a cut-off p-value of p<0.05…” The p-value will be driven by the sample size and not clinical relevance. There are no pre-specified hypotheses, so null hypothesis significance testing does not make sense.

5. **Antimicrobial resistance:** thank you for reporting results of antimicrobial susceptibility testing even though culture at baseline was unsuccessful. The recovery rate at follow-up (6 of 12 GeneXpert positive results is as expected for rectal swabs). I didn’t find the presentation of the data in Figure 3 or percentages in the text very useful. A table of the results for each isolate would take up the same amount of space as Figure 3 and, for each antimicrobial tested, the MIC for each isolate could be shown. In the text, the absolute numbers are more useful than the percentages.

**Minor comments**

1. The causative agent of gonorrhoea is *Neisseria gonorrhoeae* (not gonorrhoea). Please check the spelling throughout.

2. “Burden” – this term is used several times to refer to the frequency of infection. “Burden of disease” usually includes a measurement of the consequences of the condition, e.g. a disability-adjusted life year. With asymptomatic infections for which the clinical complications are unknown or rare, burden is not really an appropriate term.
3. Introduction (p3, Para 1): Literature review. It would be useful to be a bit more precise in the reporting of the chosen studies:
   - Of interest, you may wish to update ref 2. WHO has just published its newest estimates for 2016 (https://www.who.int/bulletin/volumes/97/8/18-228486.pdf). The estimates of global prevalence for CT and NG for men are very similar to 2012. For men in the African region, 2016 estimates for urogenital infection in men are: CT 4.0% (95% CI 2.4–6.1), NG 1.6% (95% CI 0.9–2.6).
   - Line 8, “Worldwide” is not really accurate – these are two studies, one in Italy and one in Thailand is particularly high risk populations. Please rephrase.
   - Differences in reports of prevalence will differ depending on whether the study population is unselected or all report recent receptive anal intercourse. Your description should take these factors into account.
   - Is there a systematic review? If so, please cite. If not, maybe worth pointing out in Discussion as a research need.

4. Introduction (p3, ref 10): you could acknowledge that this estimate comes from your own group.

5. Introduction (p3, para 2): “…(NAAT)… has superseded the use of cultures…” NAAT are not available everywhere so you may want to qualify this statement with “where available”.

6. Methods, Sample collection (p4): Could you give more detail about the “standardized methodology” for collection of the swabs? Was a proctoscope used? How far inside the anal canal was the swab inserted? Was it moistened? These are important details for reproducibility. Technically, the swab is a rectal swab only if inserted past the dentate line, so swabs collected without proctoscopy, which sample mainly from the anal canal might be more accurately termed as “anorectal”.

7. Data analysis (p4, Methods, line 4): “bivariable” is ambiguous. When the model examines an association between an outcome and one other variable, it is usual to call this a univariable analysis.

8. Data analysis and Table 1: the p-values are given for each stratum (e.g. age, religion). In the multivariable statistical model, it is the contribution of the variable as a whole that is important. These ‘global’ p values should be derived from a likelihood ratio test that compares the model fit with the variable to the model that includes all variables other than the one of interest. For a binary variable the results are the same.

9. Table 1: is the ethnic origin of MSM in Kenya associated with their risk of rectal STIs? If so, ethnic group might be a useful descriptive variable to report.

10. Discussion: This could be better structured. It is important that you collected data about infections, behaviours and antimicrobial susceptibility. Both presumptive treatment and frequent screening and treatment impose selection pressure for the emergence of antimicrobial resistance. The discussion is an opportunity to bring together these lines of thought in thinking about how to address rectal STIs in MSM.

11. Discussion, Limitations (p7): It’s worth mentioning the limitation of having combined the data for CT and NG owing to the small sample size.
12. Conclusion (p7): “...between 21.2% and 25%.” These point estimates suggest spurious precision and lower 95% CI is much lower. I expect each estimate lies within the confidence interval of the other, given the sample size. Please rephrase in context of confidence intervals.

13. Conclusions: in Abstract (p1, and main text (p7) don’t match. In the Abstract, there is no support from the results that “The high prevalence of asymptomatic rectal CT and NG in MSM reporting RAI demonstrates the need for frequent screening”. The more cautious statements in the main text are more appropriate.

14. Dataset: could you add a worksheet with a key to the abbreviations for the variables?

Is the work clearly and accurately presented and does it cite the current literature?
Partly

Is the study design appropriate and is the work technically sound?
Partly

Are sufficient details of methods and analysis provided to allow replication by others?
Partly

If applicable, is the statistical analysis and its interpretation appropriate?
Partly

Are all the source data underlying the results available to ensure full reproducibility?
Yes

Are the conclusions drawn adequately supported by the results?
Partly

Competing Interests: No competing interests were disclosed.

Reviewer Expertise: Epidemiology, bacterial sexually transmitted infections, antimicrobial resistance.

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard, however I have significant reservations, as outlined above.

Author Response 17 Dec 2019

Caroline Ngetsa, KEMRI-Wellcome Trust Research Programme, Kilifi, Kenya

Dear Reviewers,

Many thanks for your comments on our manuscript. We have responded to each individually and reflect the changes in the amended manuscript.

Major comments

Study design: this is a report of a cohort study, nested within a larger cohort study. Please make sure that all the items proposed in the STROBE (Strengthening the Reporting of Observational studies in Epidemiology) checklist for cohort studies (https://www.strobe-statement.org/index.php?id=available-checklists) are reported.
Thank you for your review. We have ensured that all elements of the STROBE checklist are included, and have responded to each of your comments individually. We will make sure this is visible in the revised paper.

In particular, I would like to see: a) the proportion enrolled and whether or not they differed from the men in the larger cohort study. The authors only report (p4, Results, Participant backgrounds, line 1), “104 participants were recruited and tested”;

Between April 2016 and January 2017, eligible participants were identified at follow-up visits in an existing cohort study on HIV acquisition, based on self-report of any receptive anal intercourse (RAI) within the past 6 months. We were able to enrol 70 of 174 eligible men (i.e. 40%). We have added a supplemental table that can be accessed on figshare (link in References section) to compare characteristics of the 104 cohort participants not included with those of the 70 participants included in this study: “Participants not enrolled were more likely to report sex with men exclusively, transactional sex, RAI, group sex, and any unprotected sex (data in supplemental material).

b) the follow-up time reported as a total number of months and average for each participant; 95% confidence intervals (CI) for the main percentages.

Total follow-up was 5.7 months on average (range 5.4-6.2 months) for each participant, which amounted to a total follow-up time of 39.6 person-years.

Study design: Can the authors say whether this study was pre-planned? If so, was there a target sample size with a power calculation? Was enrolment less than expected?

This study was planned and done to estimate prevalence and incidence of CT/NG infections among MSM participants who reported RAI. We did not have a prespecified hypothesis, therefore a sample size was not calculated. The precision estimate of the confidence interval was based on exact binomial confidence intervals. The study was limited by a small budget to conduct the study.

Data analysis: can the authors please calculate and report the incidence rate of Chlamydia trachomatis (CT) and Neisseria gonorrhoeae (NG) detection in anorectal samples? The availability of data from a follow-up visit is a strength of the study. Prevalence at follow-up (p4, Methods, Data analysis, line 3, and Table 2) isn’t very useful.

It would be great to change the columns in Table 2 to report: numbers with positive results at baseline; the numbers of incident cases and follow-up time; incidence rate (with 95% CI) in those with a negative test result at baseline; repeat detection rate (with 95% CI) in those with a positive test at baseline who received treatment.

Thank you for the suggestion. Given that the data is available, we have used it to calculate incidence as well and revised our Table 2 to reflect this.

Data analysis (p4, Methods, Data analysis, line 11-13): Please delete the sentence with “…a cut-off p-value of p<0.05…” The p-value will be driven by the sample size and not clinical relevance. There are no pre-specified hypotheses, so null hypothesis significance testing does not make sense.

Thank you for highlighting this.

Antimicrobial resistance: thank you for reporting results of antimicrobial susceptibility testing even though culture at baseline was unsuccessful. The recovery rate at follow-up (6 of 12 GeneXpert positive results is as expected for rectal swabs). I didn’t find the presentation of the data in Figure 3
or percentages in the text very useful. A table of the results for each isolate would take up the same amount of space as Figure 3 and, for each antimicrobial tested, the MIC for each isolate could be shown. In the text, the absolute numbers are more useful than the percentages.

Many thanks for these comments. We have included a Table 3 instead of Figure 1, which presents the data you have suggested. In addition, we have written down whole numbers instead of percentages when describing the AMR data in the manuscript.

Minor comments
The causative agent of gonorrhoea is Neisseria gonorrhoeae (not gonorrhoea). Please check the spelling throughout.
Thank you for highlighting this.

“Burden” – this term is used several times to refer to the frequency of infection. “Burden of disease” usually includes a measurement of the consequences of the condition, e.g. a disability-adjusted life year. With asymptomatic infections for which the clinical complications are unknown or rare, burden is not really an appropriate term.
We have rethought our use of this word and ensured appropriate terminology is used throughout this manuscript.

Introduction (p3, para 1): Literature review. It would be useful to be a bit more precise in the reporting of the chosen studies:
Of interest, you may wish to update ref 2. WHO has just published its newest estimates for 2016 (https://www.who.int/bulletin/volumes/97/8/18-228486.pdf). The estimates of global prevalence for CT and NG for men are very similar to 2012. For men in the African region, 2016 estimates for urogenital infection in men are: CT 4.0% (95% CI 2.4–6.1), NG 1.6% (95% CI 0.9–2.6).
Thank you for pointing us to this updated piece of literature, we have updated our manuscript to reflect the most recent evidence.

Line 8, “Worldwide” is not really accurate – these are two studies, one in Italy and one in Thailand is particularly high risk populations. Please rephrase.
Thank you, we have been more specific in describing the results found.

Differences in reports of prevalence will differ depending on whether the study population is unselected or all report recent receptive anal intercourse. Your description should take these factors into account.
Is there a systematic review? If so, please cite. If not, maybe worth pointing out in Discussion as a research need.
Many thanks for pointing out that it may seem that way. It was not our intention to do a systematic review, but rather just present existing estimates of rectal STIs in MSM in other geographical regions than Kenya. This serves as a prelude to presenting our own data. We have included in our limitations the fact that we did not systematically review all existing literature.

Introduction (p3, ref 10): you could acknowledge that this estimate comes from your own group.
Thank you, we have now acknowledged our own results.

Introduction (p3, para 2): “…(NAAT)… has superseded the use of cultures…” NAAT are not available everywhere so you may want to qualify this statement with “where available”.
This has been amended.
Methods, Sample collection (p4): Could you give more detail about the “standardized methodology” for collection of the swabs? Was a proctoscope used? How far inside the anal canal was the swab inserted? Was it moistened? These are important details for reproducibility. Technically, the swab is a rectal swab only if inserted past the dentate line, so swabs collected without proctoscopy, which sample mainly from the anal canal might be more accurately termed as “anorectal”.

This has been detailed in the revised manuscript. We used proctoscopy and dry swab kits for sample collection.

Data analysis (p4, Methods, line 4): “bivariable” is ambiguous. When the model examines an association between an outcome and one other variable, it is usual to call this a univariable analysis.

Thank you for pointing this out, we have amended this.

Data analysis and Table 1: the p-values are given for each stratum (e.g. age, religion). In the multivariable statistical model, it is the contribution of the variable as a whole that is important. These ‘global’ p values should be derived from a likelihood ratio test that compares the model fit with the variable to the model that includes all variables other than the one of interest. For a binary variable the results are the same.

Thank you for this suggestion. When we apply the p-values derived from the likelihood ratio test, we lose individual characteristic of the variable with more than 2 strata. For example, formal employment was associated with prevalent infection (borderline) when we apply the Wald p-values. When we apply the likelihood ratio test, this association becomes concealed despite having an estimate with CI’s above 0. We therefore prefer to present the Wald P values for this analysis.

Table 1: is the ethnic origin of MSM in Kenya associated with their risk of rectal STIs? If so, ethnic group might be a useful descriptive variable to report.

Thank you, we looked at this value as 5 major ethnic categories and there is no significant difference by ethnicity.

Discussion: This could be better structured. It is important that you collected data about infections, behaviours and antimicrobial susceptibility. Both presumptive treatment and frequent screening and treatment impose selection pressure for the emergence of antimicrobial resistance. The discussion is an opportunity to bring together these lines of thought in thinking about how to address rectal STIs in MSM.

Many thanks for pointing this out. We have now better structured our conclusion and highlighted the very important fact that these methods could increase AMR.

Discussion, Limitations (p7): It’s worth mentioning the limitation of having combined the data for CT and NG owing to the small sample size.

Thank you for your suggestion, we have now added this as one of our limitations.

Conclusion (p7): “…between 21.2% and 25%.” These point estimates suggest spurious precision and lower 95% CI is much lower. I expect each estimate lies within the confidence interval of the other, given the sample size. Please rephrase in context of confidence intervals.

Thank you, we’ve done this now.
Conclusions: in Abstract (p1, and main text (p7) don’t match. In the Abstract, there is no support from the results that “The high prevalence of asymptomatic rectal CT and NG in MSM reporting RAI demonstrates the need for frequent screening”. The more cautious statements in the main text are more appropriate.

We have modified the abstract to reflect this, thank you.

Dataset: could you add a worksheet with a key to the abbreviations for the variables?

We have now included a worksheet with dataset abbreviations (it can be found on figshare, and there is a link to that in the References section of the manuscript).

Competing Interests: None
Are sufficient details of methods and analysis provided to allow replication by others?  
Yes

If applicable, is the statistical analysis and its interpretation appropriate?  
Yes

Are all the source data underlying the results available to ensure full reproducibility?  
Yes

Are the conclusions drawn adequately supported by the results?  
Yes

**Competing Interests:** No competing interests were disclosed.

**Reviewer Expertise:** Behavioral Epidemiology of STIs and HIV in MSM, Africa.

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard.

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**Author Response 17 Dec 2019**

Caroline Ngetsa, KEMRI-Wellcome Trust Research Programme, Kilifi, Kenya

Dear Reviewers,

Many thanks for your comments on our manuscript. We have responded to each individually and reflect the changes in the amended manuscript.

This is a useful report of rectal gonorrhea and chlamydia in MSM in coastal Kenya. The Introduction suffers from missing some crucial findings from a similar study of prevalence of rectal NG/CT in MSM in Tanga and Dar es Salaam, a few hundred miles south of where this study was carried out (Ross MW et al., BMJ Open 2014)1. Some of those interesting findings were replicated, including the lack of association between HIV and STI in MSM, plus similar high rates of rectal STIs. There may be lessons here about access to clinics that will perform rectal examinations/swabs, especially if the MSM are not "out" to health care workers (rectal STIs are a marker for MSM activity and likely to engender discriminatory behavior in many health workers). Religion was given for 77%, but what was the other quarter? Hindi? No religion?

Thank you for your review. We have now referred to the study in Dar Es Salaam and Tanga in the background and discussion sections of this manuscript. The point of discussion regarding access to clinics is also very relevant; we feel that we do raise this issue a bit in our introduction and discussion, by raising issues with stigma related to MSM in general; this should encompass healthcare access as well.

We have also written down the religious background of the remaining quarter of participants; this is also visible in Table 1.

This is an important study, with the gaps noted above. It would also have been helpful to briefly speculate as to what bias might have been associated with recruiting participants from an existing high-risk cohort study.
Thank you for noting this. We have included details of eligible patients who were not included in the study (and have included data available on figshare, a link is available in the References of the manuscript): importantly, it was noted that patients not included in the study had higher risk behaviour. This is unlikely to underestimate true infection prevalence and rate within this population.

*Competing Interests:* None