# Anorectal swabs as a marker of male-to-male sexual exposure in STI surveillance systems

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## SUMMARY

Identification of priority populations such as men who have sex with men (MSM) is important in surveillance systems to monitor trends of sexually transmitted infections (STIs). We explored using routinely collected non-behavioural data as a means to establish MSM status in surveillance by assessing anorectal swab as a marker of male-to-male sexual exposure. We used chlamydia testing data from a sexual health clinic, 2007–2012. Men reporting any male sexual partner(s) in the previous 12 months were considered MSM. The dataset was split into development and validation samples to develop a univariate predictive model and assess the model fit. The dataset included 30 358 individual men and 48 554 episodes of STI testing; 45% were among reported MSM and an anorectal swab was performed in 40% of testing episodes. Anorectal swabbing had good diagnostic performance as a marker for MSM status (sensitivity = 87%, specificity = 99%, positive predictive value =  $98 \cdot 6\%$ , negative predictive value =  $90 \cdot 3\%$ ). The model showed good fit against the internal validation sample (area under the curve = 0.93). Anorectal swabs are a valid marker of MSM behaviour in surveillance data from sexual health clinics, and they are likely to be particularly useful for monitoring STI trends among MSM with higher risk behaviour.

Key words: Chlamydia, men who have sex with men – MSM, sexually transmitted infections, surveillance.

### **INTRODUCTION**

Surveillance of sexually transmitted infections (STIs) allows trends in STI epidemiology to be identified

and the impacts of interventions and screening programs to be evaluated. Identification of key populations such as men who have sex with men (MSM), young people and Aboriginal and Torres Strait Islanders in surveillance systems is important to be able to monitor trends and epidemics of STIs in priority populations for STI control [1–3].

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In addition to passive surveillance, in which positive results are reported to a public health authority, some jurisdictions operate active surveillance systems. These have the advantage of capturing negative as well as positive testing data, and may target specific high-risk populations from sentinel sites, such as sexual health services. Sentinel surveillance systems have also been established to capture testing data from public and private laboratories (for example, the Australian Collaboration for Coordinated Enhanced Sentinel Surveillance of Blood Borne Viruses and Sexually Transmitted Infections (ACCESS), a multijurisdictional Australian system [4]). Laboratory-based surveillance can be used to estimate the prevalence of STIs (by monitoring the proportion of tests that are positive), as well as incidence and reinfection rates, among large populations from varied clinical settings. However, it is difficult to identify key populations within these datasets, due to the limited recording of behavioural information (for example, gender of sexual partners) in laboratory records and by clinicians on pathology request forms. Therefore, other means of identifying at-risk populations such as MSM are required in systems where sexual exposure data are not available.

Australian and international guidelines recommend that asymptomatic STI screening be conducted among MSM at least annually, and at multiple anatomical sites (urethral, pharyngeal and anorectal) [5–7]. As specimen site is captured by laboratory testing data, and it is unlikely that men would have an anorectal swab if not engaging in receptive anal intercourse, particularly since anorectal STIs are largely asymptomatic [8, 9], swab for anorectal chlamydia may be a valid marker of male-to-male sexual exposure.

Therefore, the objectives of this study were to determine the predictive value of anorectal swab site for male-to-male sexual exposure and whether having an anorectal swab can be used as a proxy to identify MSM in STI surveillance systems.

#### **METHODS**

#### Setting

The model was developed and internally validated using a dataset derived from the Victorian Primary Care Network for Sentinel Surveillance on Blood Borne Viruses and STIs (VCPNSS) [10]. VPCNSS links HIV and STI testing data to sexual behavioural data through a network of selected sentinel clinical sites and laboratories, including clinics that see a majority of MSM, sexual health clinics, family planning clinics and community health clinics that see both MSM and non-MSM populations. VPCNSS allows for uniquely identified individuals to be followed over time.

This analysis was limited to Melbourne Sexual Health Centre (MSHC), which recorded sexual exposure for all patients enabling MSM to be identified within the overall patient group; high caseload MSM clinics were excluded due to small number of non-MSM for comparison, and other clinics excluded due to small number of anorectal tests and insufficient behavioural data.

#### Study population

Data were limited to all chlamydia tests among men taken at any anatomical site at MSHC between 2007 and 2012. This provided a dataset with sufficient power for both model development and validation. Chlamydia testing data were used exclusively because it is the only STI with a generalised epidemic in Victoria (prevalent in both MSM and non-MSM) [11]. We included laboratory test records that matched to a completed questionnaire (see below), and we excluded test records from sex workers and known HIV-positive men receiving chronic disease management at MSHC, as these groups have specific testing patterns which may have introduced selection bias.

#### Data and definitions

A test collection episode was defined as one or more chlamydia tests in one individual within 1 week – this could include testing at multiple anatomical sites and sometimes re-testing of the same anatomical site, for example in the case of inadequate samples. An anorectal test collection was defined as any chlamydia test collection that included an anorectal swab.

Attendees at MSHC answer an electronic questionnaire on self-reported sexual behaviour, including the number and gender of their sexual partners in the previous 12 months. Men who reported having at least one male sexual partner in the 12 months prior to the test collection were considered to be MSM. Age at testing was calculated using the patient's date of birth and specimen collection date (first specimen collection within each collection episode). Additional demographic (age, country of birth) and sexual risk variables (number and gender of sex partners in previous 12 months, condom use during anal and vaginal sex in previous 12 months) were extracted from the questionnaire and laboratory data for assessing differences between MSM who had and had not had anorectal swabs, as detailed in Table 4.

#### Statistical analyses

A split-sample method was applied to the MSHC dataset to develop and internally validate a predictive model. Individual test records were randomly allocated equally to either a development or validation sample.

#### Model development

Generalised linear modelling (GLM) between anorectal test collection and self-report MSM status was undertaken on the development sample to quantify the diagnostic accuracy of anorectal collection in determining MSM status.

Using the development sample, we calculated the following statistics to estimate the accuracy of anorectal swab to predict MSM status at test collection episode: diagnostic odds ratio (of MSM among anorectal *vs.* non-anorectal test groups), sensitivity, specificity, positive and negative predictive values, and positive and negative likelihood ratios.

### Validation

Predicted probabilities from GLM analyses on the development sample were then used to classify test episodes and assess the fit of the univariate model on the internal validation sample. Hosmer–Lemeshow goodness-of-fit statistic [12], area under the receiver operator curve statistic (AUC) and proportional reduction in error of classification ( $\lambda$ p) were estimated to provide a range of fit measures. Given the 2 × 2 nature of the data, the AUC represents the average classification accuracy of anorectal swabbing across sensitivity and specificity; and  $\lambda$ p indicates the % reduction in error in correct classification of MSM given knowledge of anorectal swabbing.

We then assessed whether identifying MSM by anorectal swab site could be used to extrapolate results to the total MSM testing population, by comparing the demographic and behavioural characteristics and urogenital chlamydia positivity of MSM who have had anorectal swabs with those who have not. Quantile regression and  $\chi^2$ -tests of independence were used to determine whether differences were statistically significant.

In all diagnostic accuracy and sample comparative analyses, cluster robust variance estimation was used to account for the lack of independence of observations (multiple test collection episodes per individual). To account for lack of independence in model fit analyses, bootstrap standard errors were used to derive 95% confidence intervals (CI) for AUC point estimation.

All analyses were performed with Stata version  $13 \cdot 1$  [13]. Ethics approval to undertake this study was obtained from the Alfred Health Human Ethics Committee (VPCNSS agreement, project number 213/05 and project number 62/15).

## RESULTS

The overall dataset included records of 30 358 individual men, who underwent 48 554 episodes of chlamydia testing between 2007 and 2012. Male-to-male sex in the previous 12 months was reported in 45% of test collection episodes, and an anorectal swab was performed in 40%. MSM had on average 2.2 episodes of testing, compared with 1.3 among non-MSM (P < 0.001).

#### Model development

Of the 11 002 test collections among MSM allocated to the development sample, 9597 (87%) included anorectal swabs (Table 1). Only 141 (1%) of the test collections among non-MSM included an anorectal swab (specificity 99%; Table 2). Of the 9738 anorectal test collection episodes, 9597 (98.6%) were in men reporting sex with men.

The odds of reporting MSM was 636 (95% CI 534–758) times higher for those who had had an anorectal swab than those who did not (Table 2).

#### Internal validation

Table 3 presents fit statistics for the internal validation dataset. As indicated by the AUC, goodness-of-fit statistic and estimated proportional reduction in error in classification ( $\lambda p$ ), the model fitted the internal validation sample data well.

## CHARACTERISTICS OF MSM WHO HAD ANORECTAL SWABS

Compared with MSM without anorectal collection, MSM with anorectal swabs were younger, a higher proportion reported more than 10 male partners in

| Table 1. Contingency table showing anorectal swab      |
|--|
| collection by MSM status (male-to-male sex reported in |
| last 12 months); development sample                    |

| Anorectal swab<br>done at test<br>collection episode | MSM         | Not MSM      | Total        |
|--|-------------|--------------|--------------|
| Anorectal swab                                       | 9 597 (87%) | 141 (1%)     | 9 738 (40%)  |
| Non-anorectal testing                                | 1 405 (13%) | 13 134 (99%) | 14 539 (60%) |
| Total  | 11 002      | 13 275       | 24 277       |

Cell counts and column per cent.

Table 2. Diagnostic statistics and 95% CI for assessing the diagnostic accuracy of anorectal swabbing: development sample

| Parameter                 | Estimate (95% CI)*  |  |
|---------------------------|---------------------|--|
| Diagnostic odds ratio     | 636 (534–758)       |  |
| Sensitivity               | 87.2 (86.6-87.8)    |  |
| Specificity               | 98.9 (98.7–99.1)    |  |
| Positive predictive value | 98.6 (98.3–98.8)    |  |
| Negative predictive value | 90.3 (89.8–90.8)    |  |
| Positive likelihood ratio | 82.1 (69.7–96.8)    |  |
| Negative likelihood ratio | 0.129 (0.123-0.136) |  |

\* Cluster robust standard errors specified.

Table 3. Model fit statistics for internal validation sample: AUC, Hosmer–Lemeshow goodness of fit (GOF) and proportional reduction in error of classification ( $\lambda p$ )

| Parameter                     | Development sample estimate (95% CI)*  |
|-------------------------------|--|
| AUC <sup>†</sup><br>GOF<br>λp | $\begin{array}{l} 0.93 & (0.93 - 0.93) \\ \chi^2(2) = 1.06, \ P = 0.590 \\ 0.86 \end{array}$ |

\* The 95% CI based on bootstrapped standard errors.

† AUC represents the average of anorectal swabbing sensitivity and specificity.

the previous 12 months, a higher proportion reported always using a condom during anal and vaginal sex, and a lower proportion reported male-to-female sex (Table 4). Chlamydia positivity based on urogenital test results were similar between both groups (P = 0.678).

### DISCUSSION

Anorectal swab site is a highly predictive and valid marker of male-to-male sex within our sample from a metropolitan sexual health service. Its use as a proxy for MSM in surveillance datasets where reliable behavioural data are not available is highly acceptable, particularly for monitoring STI trends among MSM with high-risk behaviour who are more vulnerable to STI.

Despite some demographic and behavioural differences between MSM with and without an anorectal swab at test collection, there was no difference in urogenital chlamydia positivity. This suggests that using anorectal swab site for surveillance would not substantially bias estimates of urogenital chlamydia prevalence among MSM. However, differences in anorectal risk cannot be inferred due to lack of comparative biological data among MSM without anorectal testing. Although MSM with anorectal swabs reported more male sex partners than MSM without anorectal collection, they also reported more consistent condom use, perhaps indicating that men who had anorectal swabs were more aware of their risk and more likely to take actions to mitigate it.

The impact of these differences in behaviour between MSM with and without anorectal swabs on prevalence of other diseases warrants further investigation, as using anorectal swabs as a marker of male-to-male sex may capture men at higher risk of STI acquisition. Men who did not have anorectal swabs were much more likely to be behaviourally bisexual; perhaps these men were less likely to engage in receptive anal intercourse, or were less open about their homosexual activity within the clinical consultation [14, 15]. Interestingly, men who had anorectal testing were statistically younger than those who did not. This may reflect true differences in sexual practices by age (for example, less receptive anal sex among older men) [16] or greater discomfort among older MSM to reveal their sexual practices. The latter is a potential concern, particularly in light of a US study which found that older men were more likely than younger men to have rectal chlamydia [17].

A small number of anorectal swabs (1%) were conducted among non-self-identified MSM. This may reflect other sexual practices, for example anal-digital activity and sharing of sex toys between partners of either sex [9, 18], or misreporting of male-to-male sex. These cases account for a low and acceptable false-positive rate and are unlikely to impact surveillance and trend monitoring where anorectal swabbing is used to identify MSM.

This study had a number of limitations. First, the variables that could be included in the model were

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| Test collection episodes in MSM ( $n = 11002$ )                 | Anorectal collection $(n = 9597)$ | Non-rectal collection $(n = 1405)$ | <i>P</i> -value |
|---|-----------------------------------|------------------------------------|-----------------|
| Median age, years   | 29.8 (29.6–30.0)                  | 33.8 (32.8–34.7)                   | <0.001          |
| Australian born ( $n = 10\ 101$ )                               | 64.5(63.0-66.1)                   | 62.7 (59.5–65.7)                   | 0.252           |
| >10 male partners in last 12 m                                  | 30.1 (28.9-31.3)                  | 20.6 (18.4-23.1)                   | <0.001          |
| Reported female partners in last $12 \text{ m}$ ( $n = 10852$ ) | 9.57 (8.84-10.4)                  | 30.9 (28.3–33.7)                   | <0.001          |
| Always use condom† $(n = 9994)$                                 | 46.7 (45.4-47.9)                  | 40.8 (37.8-43.8)                   | <0.001          |
| Urogenital chlamydia positive at test collection: $(n = 10949)$ | 3.07 (2.74–3.44)                  | 3.28 (2.46-4.35)                   | 0.678           |

Table 4. Characteristics of men reporting MSM at time of test collection, by an orectal non-rectal collection; development sample; per cent  $(95\% CI)^*$ 

Total number of observations (n) = 11002 for all comparisons, unless indicated.

\* Cluster robust standard errors specified.

† Always used a condom during anal and vaginal sex in the previous 12 months.

‡ Refers to a chlamydia-positive urine or urethral test result at test collection episode.

primarily limited to variables regularly collected in surveillance systems in order to optimise applicability: i.e., age, sex, anatomical site of test. An exception was made to enable exclusions to reduce bias in our model (sex workers and HIV-positive men receiving chronic disease care), and these may not be replicable in other surveillance systems. For example, history of ever having an anorectal swab may be a better predictor of MSM status, but is unlikely to be available in surveillance systems without capability to track individuals over time. A subanalysis (data not shown) demonstrated limited additional predictive value of lifetime anorectal swab in this sample. Second, our models were developed using data from one site only. Low rates of anorectal testing outside of sexual health services may limit the utility of anorectal swabs as a surrogate for MSM in some surveillance systems [19, 20]. The external validation of using anorectal swabbing as a marker of MSM in unrelated enhanced surveillance datasets, including those that represent the general community, to assess the performance of the model in diverse contexts is warranted. However, the lack of consistently collected behavioural data in most surveillance systems limits the options available for validation purposes. Finally, MSM status was based on a question asking about sexual contact with males but not specifying anal sex. MSM not engaging in receptive anal sex may decline an anorectal swab.

In conclusion, anorectal swabs are a valid marker of MSM behaviour in surveillance data from specialised sexual health centres. This marker provides a practical and sensitive means of identifying and following trends of STIs among MSM in surveillance systems where behavioural data are not routinely collected, thus extending the utility of laboratory and clinical surveillance data to monitor disease in a key population.

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