Estimating local chlamydia incidence and prevalence using surveillance data: eAppendix 1

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February 23, 2017

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1 A model for chlamydia surveillance data

We propose a three-compartment model of chlamydia infection, testing and screening in a closed population, as illustrated below. Uninfected individuals (U) become infected with a constant incidence, and move to either the asymptomatic-infected (A) or symptomatic-infected (S) pool. Asymptomatic-infected individuals may leave A and return to U by spontaneous clearance of their infection or by detection and treatment under a screening programme. Symptomatic individuals may similarly be screened, but will also seek treatment at a rate which is typically much higher than the rates of spontaneous clearance or screening.

Out[1]:Out[1]:

This dynamic model has a steady-state solution which depends on the transition rates α_{UA} , α_{AU} , α_{US} and α_{SU} :

```
In [2]: import sympy as sym
        from sympy import *
        A, U, S = symbols("A U S")
        alpha_UA, alpha_AU, alpha_US, alpha_SU = symbols("alpha_UA alpha_AU alpha_US alpha_SU")
        model_dyn = [
            alpha_UA*U - alpha_AU*A,
            alpha_AU*A + alpha_SU*S - (alpha_UA + alpha_US)*U,
            alpha_US*U - alpha_SU*S,
            A + U + S - 1 # this equation sets the total population size to 1
        # steady-state solution
        sol_dyn = solve(model_dyn, A, U, S)
        # functions for calculating the proportion of the population in each compartment at
        # steady state, given transition rates between compartments
        dyn_fun = lambdify((alpha_UA, alpha_AU, alpha_US, alpha_SU), sol_dyn[A] + sol_dyn[S])
       U_fun = lambdify((alpha_UA, alpha_AU, alpha_US, alpha_SU), sol_dyn[U])
        A_fun = lambdify((alpha_UA, alpha_AU, alpha_US, alpha_SU), sol_dyn[A])
       S_fun = lambdify((alpha_UA, alpha_AU, alpha_US, alpha_SU), sol_dyn[S])
Out[2]: {S: alpha_AU*alpha_US/(alpha_AU*alpha_US + alpha_SU*(alpha_AU + alpha_UA)),
         U: alpha_AU*alpha_SU/(alpha_AU*alpha_US + alpha_SU*(alpha_AU + alpha_UA)),
         A: alpha_SU*alpha_UA/(alpha_AU*alpha_US + alpha_SU*(alpha_AU + alpha_UA))}
```

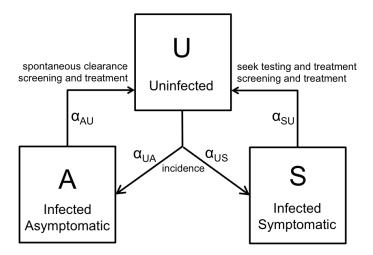


Figure 1: A model of chlamydia infection, clearance, testing and treatment.

The transition rates are functions of parameters describing behaviour and the natural history of infection:

```
\alpha_{UA} = \text{incidence} \times (1 - p_{symptomatic})
\alpha_{AU} = \text{rate of spontaneous clearance} + \text{rate of screening} \times p_{truepositive}
\alpha_{US} = \text{incidence} \times p_{symptomatic}
\alpha_{SU} = (\text{rate of screening} + \text{rate of symptomatic testing}) \times p_{truepositive}
```

Assuming all tests conducted are included in the surveillance data, the number of tests reported per unit time will be:

```
rate of testing = rate of screening + S \times rate of symptomatic testing
```

And the number of diagnoses per unit time will be:

```
rate of new diagnoses = (A + S) \times (\text{rate of screening} \times p_{truepositive})
+ (U \times \text{rate of screening} \times p_{falsepositive})
+ (S \times \text{rate of symptomatic testing} \times p_{truepositive})
```

Let's assume (based on mean sampled values for men; see Table 2 of main text) that:

- 51.0% of incident infections are asymptomatic.
- Infections (whether symptomatic or not) clear spontaneously or through background antibiotic use at a rate 0.47 per year.
- Symptomatic cases seek and obtain testing and treatment at a rate 14.4 per year.
- 97.1% of tests in infected individuals return a positive result.
- $\bullet~0.314\%$ of tests in uninfected individuals return a positive result.

```
In [3]: p_asymp = 0.510
    sc = 0.47
    att_symp = 14.4
    p_true_pos = 0.971
    p_false_pos = 0.00314
```

It is then possible to calculate the steady-state proportion of the population in each compartment given the rate of screening and incidence, and from these proportions to calculate the total prevalence, and the rates of new tests and diagnoses.

```
In [4]: %matplotlib inline
        from numpy import *
        import matplotlib.pyplot as plt
        inc = linspace(0, 0.5, 101) # incidence
        scr = linspace(0, 0.5, 101) # screening
        inc,scr = meshgrid(inc, scr)
        # proportion of population in each compartment
        ZU = U_fun(inc*p_asymp, sc + scr*p_true_pos, inc*(1-p_asymp), scr*p_true_pos + att_symp*p_true_pos)
        ZA = A_fun(inc*p_asymp, sc + scr*p_true_pos, inc*(1-p_asymp), scr*p_true_pos + att_symp*p_true_pos)
       ZS = S_fun(inc*p_asymp, sc + scr*p_true_pos, inc*(1-p_asymp), scr*p_true_pos + att_symp*p_true_pos)
        Zprev = 1 - ZU
        Ztest = scr + ZS*att_symp
        Zdiag = (ZA+ZS)*scr*p_true_pos + ZU*scr*p_false_pos + ZS*att_symp*p_true_pos
In [5]: #plt.rc("savefig", dpi=600) # for high-resolution version
       fig = plt.figure(figsize = (12, 7))
       ax1 = fig.add_subplot(231)
       p = ax1.pcolor(inc,scr, ZU)
        c = ax1.contour(inc,scr, ZU, [0.6,0.7,0.8,0.9], colors=['k','k','k','k'])
       plt.clabel(c, manual = [(0.1,0.05), (0.2,0.05), (0.4,0.05)], fmt='%1.1f')
        cb = fig.colorbar(p, ax=ax1)
       #ax1.set_xlabel('Incidence')
        ax1.set_ylabel('Screening Rate (years $^{-1}$)')
        t = ax1.text(0.25, 0.45, 'Uninfected', ha='center', size='large')
        t.set_bbox(dict(facecolor='white', alpha=0.7, edgecolor='None'))
        ax1.set_ylim(0, 0.5)
       ax1.set_xlim(0, 0.5)
       ax2 = fig.add_subplot(232)
       p = ax2.pcolor(inc,scr, ZS)
       c = ax2.contour(inc,scr, ZS, (0.003,0.006,0.009,0.012), colors='k', manual=True)
       plt.clabel(c, manual = [(0.1, 0.35), (0.2, 0.35), (0.35, 0.35), (0.45, 0.35)])
       cb = fig.colorbar(p, ax=ax2)
       t = ax2.text(0.25, 0.45, 'Infected, Symptomatic', ha='center', size='large')
        t.set_bbox(dict(facecolor='white', alpha=0.7, edgecolor='None'))
        ax2.set_ylim(0, 0.5)
       ax2.set_xlim(0, 0.5)
       ax3 = fig.add_subplot(233)
       p = ax3.pcolor(inc,scr, ZA)
       c = ax3.contour(inc,scr, ZA, (0.1,0.2,0.3), colors='k')
       plt.clabel(c, manual = [(0.1,0.1), (0.2,0.1), (0.4,0.1)], fmt='%1.1f')
        cb = fig.colorbar(p, ax=ax3)
        t = ax3.text(0.25, 0.45, 'Infected, Asymptomatic', ha='center', size='large')
        t.set_bbox(dict(facecolor='white', alpha=0.7, edgecolor='None'))
       ax3.set_ylim(0, 0.5)
        ax3.set_xlim(0, 0.5)
       ax4 = fig.add_subplot(234)
       p = ax4.pcolor(inc,scr, Zprev)
       c = ax4.contour(inc,scr, Zprev, (0.1,0.2,0.3), colors='k')
       plt.clabel(c, manual = [(0.1,0.1), (0.2,0.1), (0.4,0.1)], fmt=\frac{1}{1.1}
        cb = fig.colorbar(p, ax=ax4)
        ax4.set_xlabel('Incidence (years $^{-1}$)')
        ax4.set_ylabel('Screening Rate (years $^{-1}$)')
        t = ax4.text(0.25, 0.45, 'Prevalence', ha='center', size='large')
        t.set_bbox(dict(facecolor='white', alpha=0.7, edgecolor='None'))
        ax4.set_ylim(0, 0.5)
        ax4.set_xlim(0, 0.5)
       ax5 = fig.add_subplot(235)
        p = ax5.pcolor(inc,scr, Ztest)
        c = ax5.contour(inc,scr, Ztest, (0.2,0.4,0.6), colors='k')
        plt.clabel(c, manual = [(0.45,0.05), (0.45,0.25), (0.45,0.45)], fmt='\%1.1f')
```

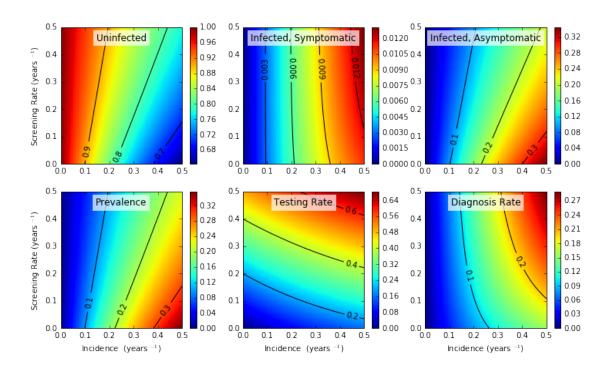


Figure 2: Upper row: the effects of incidence and screening rate on the proportion of individuals who are uninfected, infected-symptomatic and infected-asymptomatic in the model at steady state. Lower row: prevalence, testing and diagnosis rates corresponding to each combination of incidence and screening rate.

```
cb = fig.colorbar(p, ax=ax5)
ax5.set_xlabel('Incidence (years $^{-1}$)')
t = ax5.text(0.25, 0.45, 'Testing Rate', ha='center', size='large')
t.set_bbox(dict(facecolor='white', alpha=0.7, edgecolor='None'))
ax5.set_ylim(0, 0.5)
ax5.set_xlim(0, 0.5)
ax6 = fig.add_subplot(236)
p = ax6.pcolor(inc,scr, Zdiag)
c = ax6.contour(inc,scr, Zdiag, (0.1,0.2), colors='k')
plt.clabel(c, manual = [(0.2,0.2), (0.4,0.25)], fmt=\frac{1.1f}{}
cb = fig.colorbar(p, ax=ax6)
ax6.set_xlabel('Incidence (years $^{-1}$)')
t = ax6.text(0.25, 0.45, 'Diagnosis Rate', ha='center', size='large')
t.set_bbox(dict(facecolor='white', alpha=0.7, edgecolor='None'))
ax6.set_ylim(0, 0.5)
ax6.set_xlim(0, 0.5)
plt.show()
#plt.rc("savefig", dpi=80) # reset resolution
```

From the figures, it is clear that a particular pair of observed testing and diagnosis rates corresponds to a single point in the (incidence, screening rate) plane, which in turn corresponds to a particular prevalence. Note, however, that this mapping depends on the parameter values which have been assumed.

We also produce the same plot, focusing on the lower part of the incidence range:

```
In [6]: inc = linspace(0, 0.1, 101) # incidence - different range
    scr = linspace(0, 0.5, 101) # screening
    inc,scr = meshgrid(inc, scr)

# proportion of population in each compartment
ZU = U_fun(inc*p_asymp, sc + scr*p_true_pos, inc*(1-p_asymp), scr*p_true_pos + att_symp*p_true_pos)
ZA = A_fun(inc*p_asymp, sc + scr*p_true_pos, inc*(1-p_asymp), scr*p_true_pos + att_symp*p_true_pos)
```

```
ZS = S_fun(inc*p_asymp, sc + scr*p_true_pos, inc*(1-p_asymp), scr*p_true_pos + att_symp*p_true_pos)
       Zprev = 1 - ZU
        Ztest = scr + ZS*att_symp
        Zdiag = (ZA+ZS)*scr*p_true_pos + ZU*scr*p_false_pos + ZS*att_symp*p_true_pos
In [7]: fig = plt.figure(figsize = (12, 7))
       ax1 = fig.add_subplot(231)
       p = ax1.pcolor(inc,scr, ZU)
        c = ax1.contour(inc,scr, ZU, [0.92,0.94,0.96,0.98], colors=['k','k','k','k'])
       plt.clabel(c, manual = [(0.02,0.25), (0.05,0.25), (0.07,0.15), (0.09,0.05)], fmt='%1.2f')
       cb = fig.colorbar(p, ax=ax1)
       #ax1.set_xlabel('Incidence')
       ax1.set_ylabel('Screening Rate (years $^{-1}$)')
       t = ax1.text(0.05, 0.45, 'Uninfected', ha='center', size='large')
       t.set_bbox(dict(facecolor='white', alpha=0.7, edgecolor='None'))
       ax1.set_ylim(0, 0.5)
       ax1.set_xlim(0, 0.1)
       ax2 = fig.add_subplot(232)
       p = ax2.pcolor(inc,scr, ZS)
       c = ax2.contour(inc,scr, ZS, (0.001,0.002,0.003), colors='k', manual=True)
       plt.clabel(c, manual = [(0.03, 0.15), (0.06, 0.15), (0.09, 0.15)], fmt='%1.3f')
       cb = fig.colorbar(p, ax=ax2)
       t = ax2.text(0.05, 0.45, 'Infected, Symptomatic', ha='center', size='large')
       t.set_bbox(dict(facecolor='white', alpha=0.7, edgecolor='None'))
        ax2.set_ylim(0, 0.5)
       ax2.set_xlim(0, 0.1)
       ax3 = fig.add_subplot(233)
       p = ax3.pcolor(inc,scr, ZA)
       c = ax3.contour(inc,scr, ZA, (0.02,0.04,0.06,0.08), colors='k')
       plt.clabel(c, manual = [(0.02,0.25), (0.05,0.25), (0.07,0.15), (0.09,0.05)], fmt='%1.2f')
        cb = fig.colorbar(p, ax=ax3)
       t = ax3.text(0.05, 0.45, 'Infected, Asymptomatic', ha='center', size='large')
       t.set_bbox(dict(facecolor='white', alpha=0.7, edgecolor='None'))
       ax3.set_ylim(0, 0.5)
       ax3.set_xlim(0, 0.1)
       ax4 = fig.add_subplot(234)
       p = ax4.pcolor(inc,scr, Zprev)
        c = ax4.contour(inc,scr, Zprev, (0.02,0.04,0.06,0.08), colors='k')
       plt.clabel(c, manual = [(0.02, 0.25), (0.05, 0.25), (0.07, 0.15), (0.09, 0.05)], fmt='%1.2f')
        cb = fig.colorbar(p, ax=ax4)
       ax4.set_xlabel('Incidence (years $^{-1}$)')
       ax4.set_ylabel('Screening Rate (years $^{-1}$)')
       t = ax4.text(0.05, 0.45, 'Prevalence', ha='center', size='large')
        t.set_bbox(dict(facecolor='white', alpha=0.7, edgecolor='None'))
        ax4.set_ylim(0, 0.5)
       ax4.set_xlim(0, 0.1)
       ax5 = fig.add_subplot(235)
       p = ax5.pcolor(inc,scr, Ztest)
        c = ax5.contour(inc,scr, Ztest, (0.2,0.4), colors='k')
       plt.clabel(c, manual = [(0.045,0.2), (0.045,0.5)], fmt=\frac{115}{115}
        cb = fig.colorbar(p, ax=ax5)
       ax5.set_xlabel('Incidence (years $^{-1}$)')
        t = ax5.text(0.05, 0.45, 'Testing Rate', ha='center', size='large')
        t.set_bbox(dict(facecolor='white', alpha=0.7, edgecolor='None'))
       ax5.set_ylim(0, 0.5)
       ax5.set_xlim(0, 0.1)
       ax6 = fig.add_subplot(236)
       p = ax6.pcolor(inc,scr, Zdiag)
       c = ax6.contour(inc,scr, Zdiag, (0.02,0.04,0.06), colors='k')
       plt.clabel(c, manual = [(0.04,0.2), (0.06,0.4), (0.09,0.35)], fmt='\%1.2f')
        cb = fig.colorbar(p, ax=ax6)
        ax6.set_xlabel('Incidence (years $^{-1}$)')
```

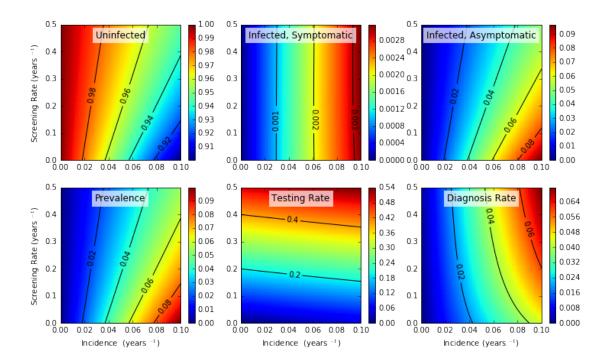


Figure 3: Upper row: the effects of incidence and screening rate on the proportion of individuals who are uninfected, infected-symptomatic and infected-asymptomatic in the model at steady state. Lower row: prevalence, testing and diagnosis rates corresponding to each combination of incidence and screening rate.

```
t = ax6.text(0.05, 0.45, 'Diagnosis Rate', ha='center', size='large')
t.set_bbox(dict(facecolor='white', alpha=0.7, edgecolor='None'))
ax6.set_ylim(0, 0.5)
ax6.set_xlim(0, 0.1)
plt.show()
```

1.1 Steady-state assumption

In using the model to interpret testing and diagnosis data, we assume the system is at steady state. We investigate to what extent this assumption is valid by perturbing the system and observing the return to equilibrium.

First, we use the national coverage and diagnoses per capita in men in the years 2012 - 2015. The analysis proceeds as follows:

- 1. Begin by estimating the steady state in 2012, using 2012 data.
- 2. Assuming a (potentially different) steady state in 2013, estimate incidence and screening rate in 2013.
- 3. Starting at the 2012 steady state, simulate the evolution of the system for one year with 2013 incidence and screening figures. Compare the prevalence after one year with the steady-state prevalence estimated for 2013 from that year's surveillance data, to see how closely they agree.
- 4. Repeat steps 2 and 3 for 2014 and 2015, each time starting the system in the state it had reached at the end of the previous one-year period.

The results of the simulations are then plotted.

```
model_test_diag = [
            tsym - ( ssym + (1 - A - U)*test_sym ),
            dsym - ( A*ssym*p_true_pos + U*ssym*p_false_pos + (1 - A - U)*test_sym*p_true_pos )
        sol_test_diag = solve(model_test_diag, tsym, dsym)
        test_fun = lambdify((A, U, ssym, test_sym), sol_test_diag[tsym])
        diag_fun = lambdify((A, U, ssym, test_sym), sol_test_diag[dsym])
        def test_diag_fun(parms):
            # parms = (incidence, screening rate)
            inc = parms[0]
            scr = parms[1]
             \texttt{A} = \texttt{A\_fun(inc*p\_asymp, sc} + \texttt{scr*p\_true\_pos, inc*(1 - p\_asymp), scr*p\_true\_pos} + \texttt{att\_symp*p\_true\_pos)} 
            U = U_fun(inc*p_asymp, sc + scr*p_true_pos, inc*(1 - p_asymp), scr*p_true_pos + att_symp*p_true_pos)
            return [test_fun(A, U, scr, att_symp), diag_fun(A, U, scr, att_symp)]
        # set up a function to simulate system dynamics when perturbed from steady state
        from scipy.integrate import odeint
        def dydt(y, t, parms):
            return([
            parms[1]*y[1] + parms[3]*y[2] - (parms[0] + parms[2])*y[0],
            parms[0]*y[0] - parms[1]*y[1],
parms[2]*y[0] - parms[3]*y[2]
In [9]: # find steady state based on 2012 data
        cov_2012 = 566908. / 3519015.
        adpc_2012 = 48387. / 3519015.
        [incsol, scrsol] = fsolve(
            lambda x: [test_diag_fun(x)[0] - cov_2012, test_diag_fun(x)[1] - adpc_2012],
            [0.09, 0.25]
        U_2012 = U_fun(
            incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos
        A 2012 = A fun(
            incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos
        S_2012 = S_fun(
            incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos
            )
        # find incidence and screening based on 2013 data
        cov_2013 = 531428. / 3519015.
        adpc_2013 = 48825. / 3519015.
        [incsol, scrsol] = fsolve(
            lambda x: [test_diag_fun(x)[0] - cov_2013, test_diag_fun(x)[1] - adpc_2013],
            [0.09, 0.25]
            )
        # solve, 2012-2013
        inc = incsol
        scr = scrsol
        parms = \
            [incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos]
        sol_12_13 = odeint(dydt,
                [U_2012,A_2012,S_2012],
               linspace(0,10,1000),
               args = (parms,)
```

```
In [10]: # incidence and screening based on 2014 data
         cov_2014 = 493327. / 3500026.
         adpc_2014 = 47437. / 3500026.
         [incsol, scrsol] = fsolve(
             lambda x: [test_diag_fun(x)[0] - cov_2014, test_diag_fun(x)[1] - adpc_2014],
             [0.09, 0.25]
         inc = incsol
         scr = scrsol
         parms = \
             [incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos]
         # solve, 2013-2014
         sol_13_14 = odeint(dydt,
                sol_12_13[999,:],
                linspace(0,10,1000),
                args = (parms,)
In [11]: # incidence and screening based on 2015 data
         cov_2015 = 446279. / 3496125.
         adpc_2015 = 44609. / 3496125.
         [incsol, scrsol] = fsolve(
             lambda x: [test_diag_fun(x)[0] - cov_2015, test_diag_fun(x)[1] - adpc_2015],
         inc = incsol
         scr = scrsol
         parms = \
             [incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos]
         # solve, 2013-2014
         sol_14_15 = odeint(dydt,
                sol_13_14[999,:],
                linspace(0,10,1000),
                args = (parms,)
In [12]: # plot solutions
         plt.plot(linspace(2012,2013,1000), sol_12_13[:,1]+sol_12_13[:,2], label='2012-2013')
         plt.plot(linspace(2013,2014,1000), sol_13_14[:,1]+sol_13_14[:,2], label='2013-2014')
         plt.plot(linspace(2014,2015,1000), sol_14_15[:,1]+sol_14_15[:,2], label='2014-2015')
        plt.ylim(0,0.025)
         plt.ylabel('Prevalence')
         plt.xticks([2012,2013,2014,2015], ['2012','2013','2014','2015'])
         plt.legend(loc=4)
Out[12]: <matplotlib.legend.Legend at 0x113a81710>
```

The plot shows that prevalence was very close to the steady state, with differences being very small compared to the uncertainty in prevalence estimates illustrated in the Figures in the main text.

To investigate the validity of the steady state assumption at a local level, we identified the local authorities with the largest changes in prevalence between 2012 and 2013:

	Test rate	$(year^{-1})$	Diagnosis rate	$(year^{-1})$	Prevalence	
Year	2012	2013	2012	2013	2012	2013
North Lincolnshire	0.101	0.173	0.022	0.011	2.1%	3.9%
Haringey	0.267	0.191	0.035	0.018	4.2%	2.7%
Dudley	0.075	0.239	0.020	0.006	1.1%	2.4%

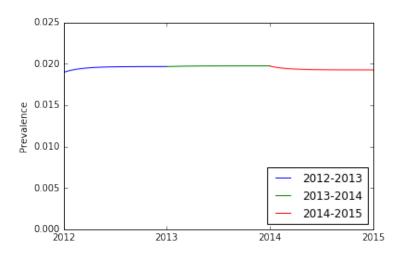


Figure 4: Simulated dynamics of national prevalence in 15-24-year-old men, 2012-2015.

```
lambda x: [test_diag_fun(x)[0] - cov_2012, test_diag_fun(x)[1] - adpc_2012],
             [0.09, 0.25]
             )
         U_2012 = U_fun(
             incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos
         A_2012 = A_fun(
             incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos
         S_2012 = S_fun(
             incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos
             )
         # find incidence and screening based on 2013 data
         cov_2013 = 0.173269822929
         adpc_2013 = 0.0216211803756
         [incsol, scrsol] = fsolve(
             lambda x: [test_diag_fun(x)[0] - cov_2013, test_diag_fun(x)[1] - adpc_2013],
             [0.09, 0.25]
             )
         # solve, 2012-2013
         inc = incsol
         scr = scrsol
         parms = \
             [incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos]
         sol_n_lincs = odeint(dydt,
                [U_2012,A_2012,S_2012],
                linspace(0,10,1000),
                args = (parms,)
In [14]: # Haringey
         # find steady state based on 2012 data
         cov_2012 = 0.267007002375
         adpc_2012 = 0.0346976493046
         [incsol, scrsol] = fsolve(
             lambda x: [test_diag_fun(x)[0] - cov_2012, test_diag_fun(x)[1] - adpc_2012],
             [0.09, 0.25]
```

```
U_2012 = U_fun(
             incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos
         A 2012 = A fun(
             incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos
         S_2012 = S_fun(
             incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos
         # find incidence and screening based on 2013 data
         cov_2013 = 0.190544970144
         adpc_2013 = 0.0184872060681
         [incsol, scrsol] = fsolve(
             lambda \ x: \ [test\_diag\_fun(x)[0] \ - \ cov\_2013, \ test\_diag\_fun(x)[1] \ - \ adpc\_2013],
         # solve, 2012-2013
         inc = incsol
         scr = scrsol
         parms = \
             [incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos]
         sol_haringey = odeint(dydt,
                [U_2012,A_2012,S_2012],
                linspace(0,10,1000),
                args = (parms,)
In [15]: # Dudley
         # find steady state based on 2012 data
         cov_2012 = 0.0750667240187
         adpc_2012 = 0.0057129570304
         [incsol, scrsol] = fsolve(
             lambda x: [test_diag_fun(x)[0] - cov_2012, test_diag_fun(x)[1] - adpc_2012],
             [0.09, 0.25]
             )
         U_2012 = U_fun(
             incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos
         A_2012 = A_fun(
             incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos
         S_2012 = S_fun(
             incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos
             )
         # find incidence and screening based on 2013 data
         cov_2013 = 0.238873910562
         adpc_2013 = 0.0199612670162
         [incsol, scrsol] = fsolve(
             lambda x: [test_diag_fun(x)[0] - cov_2013, test_diag_fun(x)[1] - adpc_2013],
             [0.09, 0.25]
             )
         # solve, 2012-2013
         inc = incsol
         scr = scrsol
         parms = \
             [incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos]
         sol_dudley = odeint(dydt,
                [U_2012,A_2012,S_2012],
```

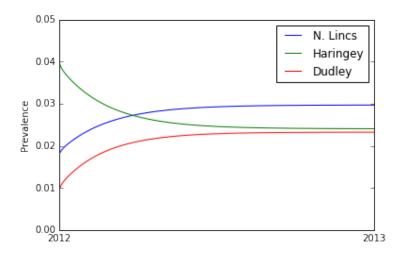


Figure 5: Simulated dynamics of local authority prevalence in 15-24-year-old men, 2012-2013.

At local level changes in prevalence can be more pronounced than at national level, but even with the largest changes in prevalence the new steady state is reached after much less than a year.

1.2 Different testing rates in different populations

We also investigate the sensitivity of the model to different testing rates in subpoulations with different prevalences. This analysis makes use of results reported in Woodhall Sex. Transm. Infect. 92:21-227 (2016) for the proportion of 16-24-year-old men in Natsal-3 reporting different risk behaviours and chlamydia testing and diagnosis in the last year.

Taking each risk factor in turn, we estimate prevalence for each risk level and take the weighted average as an estimate of population prevalence. We also estimate prevalence from the proportion tested and and diagnosed in the whole population, for comparison.

```
[0.351, 0.261, 0.388],
                    [0.573, 0.187, 0.136, 0.104],
                    [0.421, 0.326, 0.253],
                    [0.331, 0.475, 0.194],
                    [0.527, 0.224, 0.249],
                    [0.517, 0.483],
                    [0.712, 0.143, 0.145],
                    [0.526, 0.202, 0.273],
                    [0.08, 0.92]]
         # proportion reporting testing in the last year, by risk factor level
         n3_{test} = [[0.404, 0.311],
                    [0.345, 0.333, 0.352],
                    [0.336, 0.378],
                    [0.256, 0.334, 0.453],
                    [0.26, 0.403, 0.43, 0.609],
                    [0.26, 0.367, 0.463],
                    [0.270, 0.343, 0.488],
                    [0.253,0.396,0.492],
                    [0.335, 0.38],
                    [0.329, 0.497, 0.385],
                    [0.281, 0.399, 0.432],
                    [0.339, 0.424]]
         # proportion reporting diagnosis in the last year, by risk factor level
         n3_{diag} = [[0.404*0.047, 0.311*0.067],
                    [0.345*0.052, 0.333*0.05, 0.352*0.065],
                    [0.336*0.053, 0.378*0.072],
                    [0.256*0.028, 0.334*0.047, 0.453*0.078],
                    [0.26*0.034, 0.403*0.009, 0.43*0.015, 0.609*0.212],
                    [0.26*0.055, 0.367*0.04, 0.463*0.08],
                    [0.270*0.018, 0.343*0.049, 0.488*0.109],
                    [0.253*0.01, 0.396*0.038, 0.492*0.123],
                    [0.335*0.043, 0.38*0.074],
                    [0.329*0.067, 0.497*0.065, 0.385*0.013],
                    [0.281*0.035, 0.399*0.04, 0.432*0.098],
                    [0.339*0.058, 0.424*0.051]]
In [18]: plt.figure(figsize=(8,6))
         for j in xrange(len(n3_test)):
             wav = 0
             wav_pos = 0
             plt.plot([0,0.09],[2*(j+1), 2*(j+1)], '0.8')
             for i in xrange(len(n3_test[j])):
                 cov = -log(1 - n3_test[j][i])
                 adpc = -log(1 - n3_diag[j][i])
                 [incsol, scrsol] = fsolve(
                     lambda x: [test_diag_fun(x)[0] - cov, test_diag_fun(x)[1] - adpc], [0.09, 0.25]
                 prev = 1 - U_fun(incsol*p_asymp,
                                  sc + scrsol*p_true_pos,
                                  incsol*(1-p_asymp),
                                  scrsol*p_true_pos + att_symp*p_true_pos
                 plt.plot(prev, 2*(j+1), 'ob', markerfacecolor='None', markersize=20*sqrt(n3_props[j][i]))
                 wav = wav + n3_props[j][i]*prev
                 wav_pos = wav_pos + n3_props[j][i]*adpc/cov
             plt.plot(wav, 2*(j+1), 'ob')
             plt.text(0.082, 2*(j+1), factors[j], verticalalignment='center')
         # overall
         cov = -log(1 - 0.346)
         adpc = -log(1 - 0.02)
         [incsol, scrsol] = fsolve(
             lambda x: [test_diag_fun(x)[0] - cov, test_diag_fun(x)[1] - adpc], [0.09, 0.25]
         prev = 1 - U_fun(
```

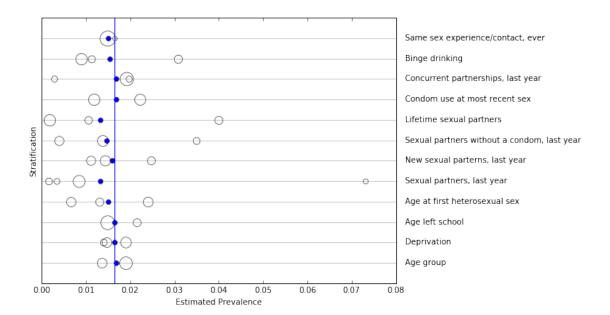


Figure 6: Sensitivity of prevalence estimates in 16-24-year-old men to risk-dependent differences in testing. Hollow markers: risk-level-specific estimates. Marker area is proportional to the proportion of the population in each risk category. Solid markers: weighted mean of level-specific estimates. Vertical line: estimate using aggregated proportions tested and diagnosed.

```
incsol*p_asymp, sc + scrsol*p_true_pos, incsol*(1-p_asymp), scrsol*p_true_pos + att_symp*p_true_pos
)

plt.plot([prev,prev],[0,100])

cur_axes = plt.gca()
 cur_axes.axes.get_yaxis().set_ticks([])
 plt.xlim([0,0.08])
 plt.ylim([0,26])
 plt.xlabel('Estimated Prevalence')
 plt.ylabel('Stratification')
Out [18]: <matplotlib.text.Text at 0x114ad1250>
```

In this figure, hollow markers show risk-level-specific prevalence estimates and their area represents the proportion of the population in each risk group. Large markers show the weighted average of these level-specific estimates. The solid line shows prevalence estimated from aggregated testing and diagnosis (ie. not stratified by risk). It should be emphasised that due to limitations of the data, the analysis is intended as an illustration of the model's theoretical properties rather than an accurate estimate of prevalence in the different risk categories. The data from Natsal-3 is some of the best available, but nonetheless relies on participants' recall and accurate self-reporting. It was collected at a national level, and equivalent information is not available at a local level for incorporation into local-level prevalence estimates.

Although aggregating across the population does affect prevalence estimates, the differences are small compared with the 1-2% uncertainty which we found in our analyses of the surveillance data.

In []: