

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

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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.

```
In [1]: from IPython.display import Image
        Image(filename="figures/3_comp.png", width=500)
```

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])

        sol_dyn

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))}
```

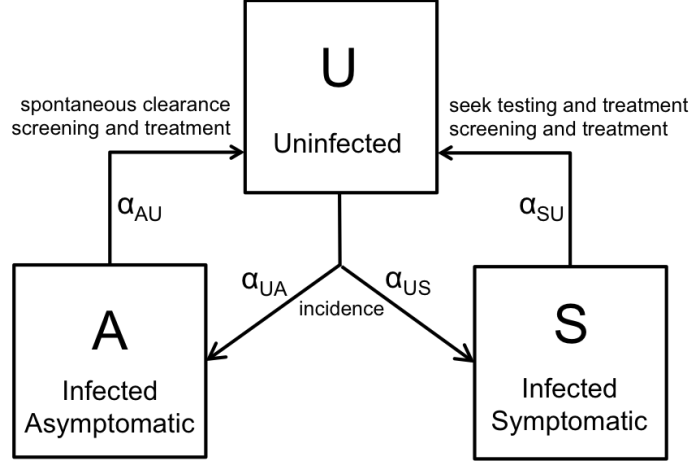


Figure 1: A model of chlamydia infection, clearance, testing and treatment.

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

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

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

$$\text{rate of testing} = \text{rate of screening} + S \times \text{rate of symptomatic testing}$$

And the number of diagnoses per unit time will be:

$$\begin{aligned}
 \text{rate of new diagnoses} &= (A + S) \times (\text{rate of screening} \times p_{\text{truepositive}}) \\
 &\quad + (U \times \text{rate of screening} \times p_{\text{falsepositive}}) \\
 &\quad + (S \times \text{rate of symptomatic testing} \times p_{\text{truepositive}})
 \end{aligned}$$

Let's assume 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.
- 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]: 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='%1.1f')
        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')
        cb = fig.colorbar(p, ax=ax5)
        ax5.set_xlabel('Incidence (years  $^{-1}$ )')

```

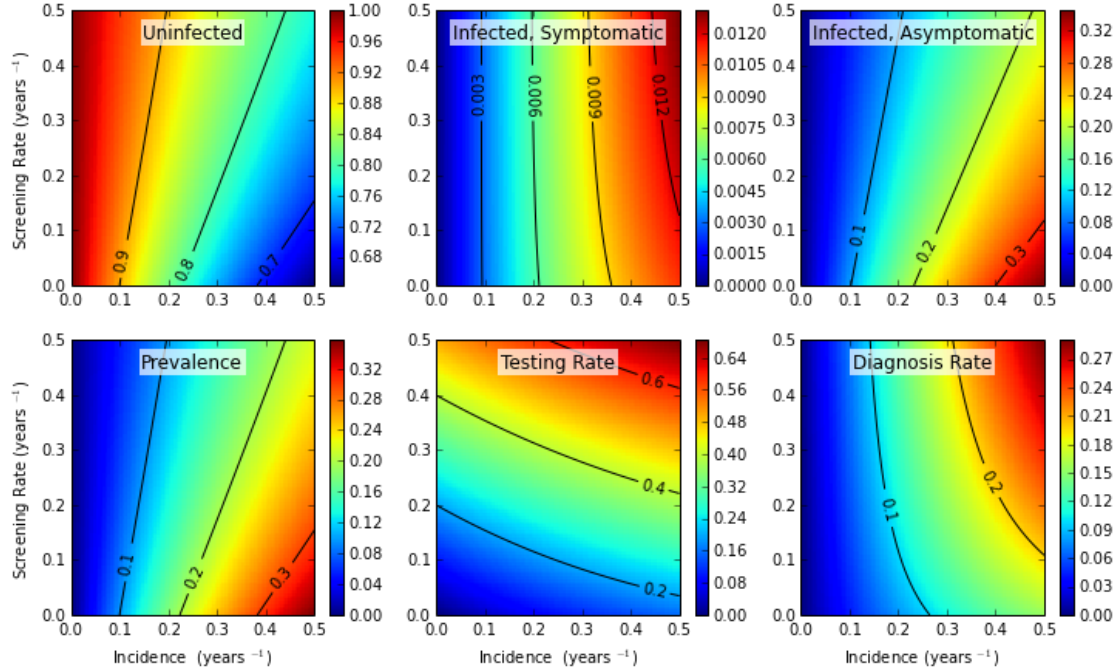


Figure 2: Upper row: the effects of incidence and screening rate on the proportion of individuals who are uninfected, infected-symptomatic and infected-asymptomatic in the model at steady state. Lower row: prevalence, testing and diagnosis rates corresponding to each combination of incidence and screening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

```

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='%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()

```

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, focussing 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, scr + scr*p_true_pos, inc*(1-p_asymp), scr*p_true_pos + att_symp*p_true_pos)
ZA = A_fun(inc*p_asymp, scr + scr*p_true_pos, inc*(1-p_asymp), scr*p_true_pos + att_symp*p_true_pos)
ZS = S_fun(inc*p_asymp, scr + 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  $\sim$ 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  $\sim$ 1$)')
ax4.set_ylabel('Screening Rate (years  $\sim$ 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='%1.1f')
cb = fig.colorbar(p, ax=ax5)
ax5.set_xlabel('Incidence (years  $\sim$ 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  $\sim$ 1$)')
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)

```

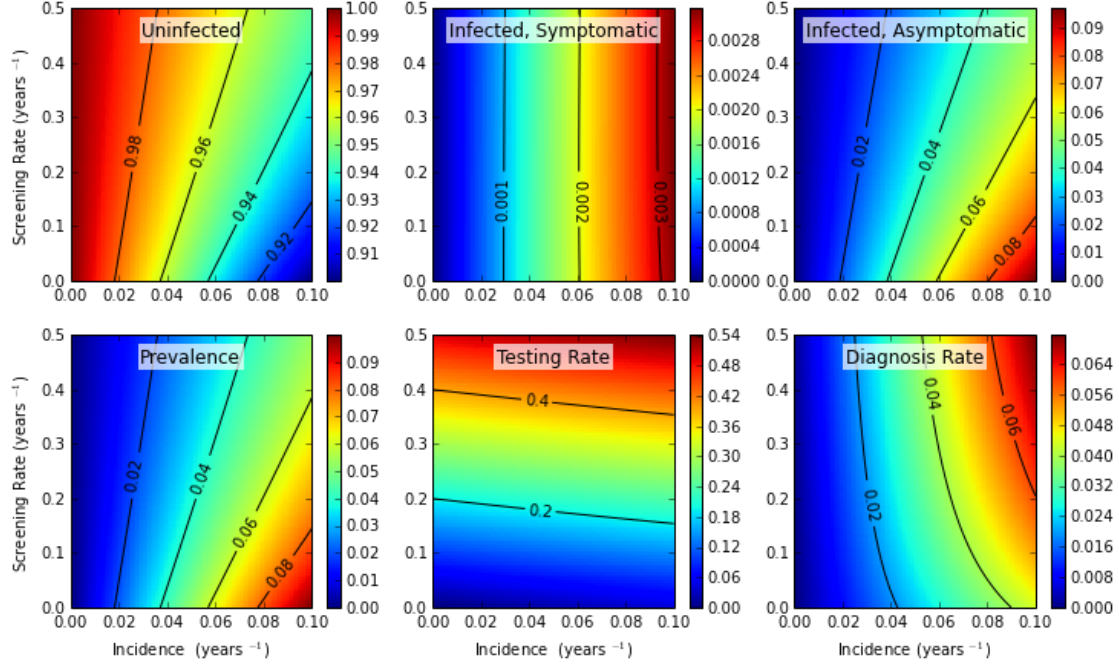


Figure 3: Upper row: the effects of incidence and screening rate on the proportion of individuals who are uninfected, infected-symptomatic and infected-asymptomatic in the model at steady state. Lower row: prevalence, testing and diagnosis rates corresponding to each combination of incidence and screening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

```
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.
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.

```
In [8]: from scipy.optimize import fsolve

tsym, dsym, ssym, test_sym = symbols('tsym dsym ssym test_sym')

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]

    A = A_fun(inc*p_asymp, sc + scr*p_true_pos, inc*(1 - p_asymp), scr*p_true_pos + 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(

```

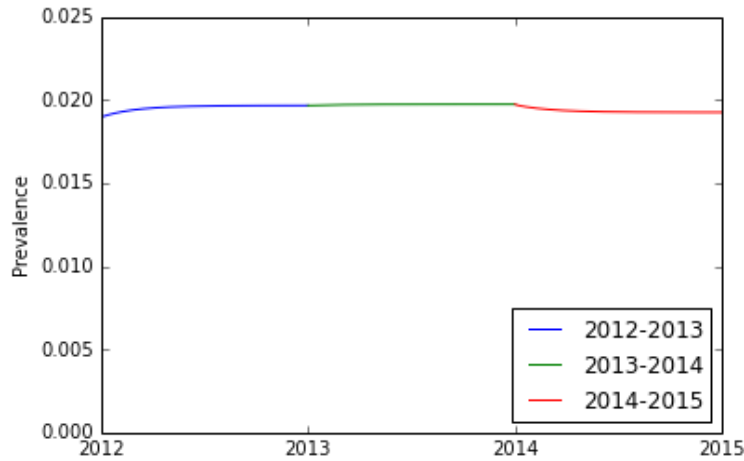


Figure 4: Simulated dynamics of national prevalence in 15-24-year-old men, 2012-2015.

```

    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],
    [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_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 0x107567a50>

The plot shows that although there were inevitably some perturbations to the steady state, they were

small compared to the uncertainty in prevalence estimates illustrated in 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 (analysis not shown):

	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%

```
In [13]: # North Lincolnshire
# find steady state based on 2012 data

cov_2012 = 0.100807801953
adpc_2012 = 0.0111652211547
[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 = 0.173269822929
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],
    [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_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],
    linspace(0,10,1000),
    args = (parms,)
)

In [16]: # plot solutions

```

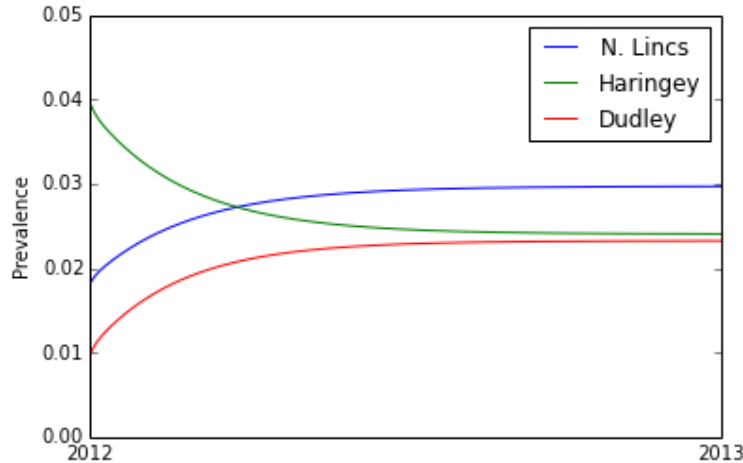


Figure 5: Simulated dynamics of national prevalence in 15-24-year-old men, 2012-2015.

```
plt.plot(linspace(2012,2013,1000), sol_n_lincs[:,1]+sol_n_lincs[:,2], label='N. Lincs')
plt.plot(linspace(2012,2013,1000), sol_haringey[:,1]+sol_haringey[:,2], label = 'Haringey')
plt.plot(linspace(2012,2013,1000), sol_dudley[:,1]+sol_dudley[:,2], label = 'Dudley')
plt.ylim(0,0.05)
plt.xlim(2012,2013)
plt.ylabel('Prevalence')
plt.xticks([2012,2013], ['2012','2013'])
plt.legend()
```

Out [16]: <matplotlib.legend.Legend at 0x1075f7c90>

At local level there can be more pronounced temporal changes, but even the largest changes in prevalence have time to reach steady state over the course of a year.

1.2 Different testing rates in different populations

We also investigate the sensitivity of the model to different testing rates in subpoulation with different prevalences. This analysis makes use of results reported in Woodhall *Sex. Transm. Infect.* **92**:21-227 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.

```
In [17]: # analysis by identified risk factors for prevalent infection:
# age group, deprivation index and lifetime number of sexual partners
factors = ['Age group', 'Deprivation', 'Lifetime sexual partners']

# proportion of those surveyed reporting each risk factor level
n3_props = [[0.373, 0.627],[0.369,0.182,0.449],[0.527, 0.224, 0.249]]

# proportion reporting testing in the last year, by risk factor level
n3_test = [[0.404, 0.311],[0.345, 0.333, 0.352],[0.253,0.396,0.492]]

# 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],
            [0.253*0.01,0.396*0.038,0.492*0.123]]
```

```
In [18]: plt.figure(figsize=(6,4))

for j in xrange(len(n3_test)):
```

```

wav = 0
wav_pos = 0
for j 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), '.b')
    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(0.05, 2*(j+1), factors[j])

# 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(
    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([0,0.09])
plt.ylim([0,8])
plt.xlabel('Estimated Prevalence')

```

Out[18]: <matplotlib.text.Text at 0x1067bef10>

In the figure above, small markers show risk-level-specific prevalence estimates and large markers show the weighted average of the level-specific estimates. The solid line shows prevalence estimated from aggregated testing and diagnosis (ie. not stratified by risk).

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.

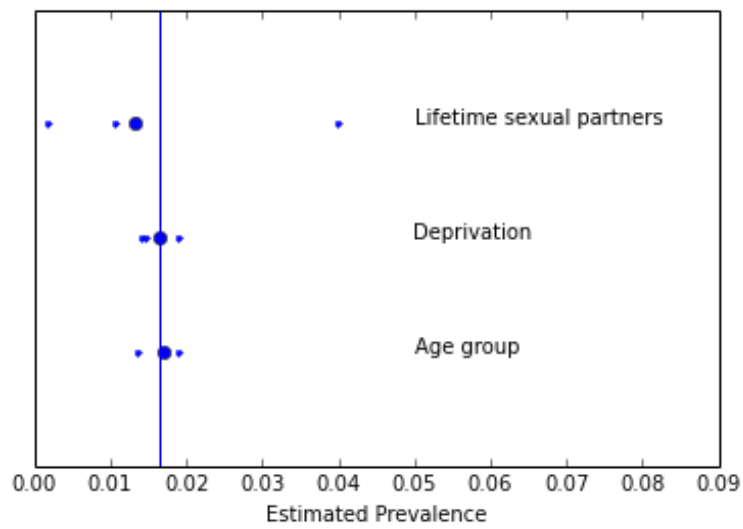


Figure 6: Sensitivity of prevalence estimates in 16-24-year-old men to risk-dependent differences in testing. Small markers: risk-level-specific estimates. Large markers: weighted mean of level-specific estimates. Vertical line: estimate using aggregated proportions tested and diagnosed.