

Appendix (unedited, as supplied by the authors)

Details of the mathematical model

The model used for the simulations in this manuscript was adapted from a previously described model of HCV transmission among PWID¹, which is described below. Note that in the Victoria simulations we assume no immunity ($\sigma=1$ and $\xi=0$ in the model equations below, such that all those who attain SVR after treatment, or who spontaneously clear the acute stage, become susceptible).

In this system, X denotes susceptible PWID (including those who have cleared the infection and are Ab^+), C_1 denotes both chronically infected and acutely infected PWID which will proceed to chronic infections, C_2 denotes chronically infected PWID who did not achieve sustained viral response (SVR) after treatment, T denotes PWID in treatment, Z denotes immune PWID, τ is time in years, and where $N=X+Z+C_1+C_2+T$. The model equations are as follows:

$$\frac{dX}{d\tau} = \theta - \pi(1 - \delta + \delta\xi) \frac{C_1 + C_2}{N} X + \omega\alpha\sigma T - \mu X$$

$$\frac{dC_1}{d\tau} = \pi(1 - \delta) \frac{C_1 + C_2}{N} X - f(C_1) - \mu C_1$$

$$\frac{dT}{d\tau} = f(C_1) - \omega T - \mu T$$

$$\frac{dZ}{d\tau} = \pi\delta\xi \frac{C_1 + C_2}{N} X + \omega\alpha(1 - \sigma)T - \mu Z$$

$$\frac{dC_2}{d\tau} = \omega(1 - \alpha)T - \mu C_2$$

where

$$f(C_1) = \begin{cases} \Phi, & \Phi < C_1 \\ C_1, & 0 \leq C_1 < \Phi. \end{cases}$$

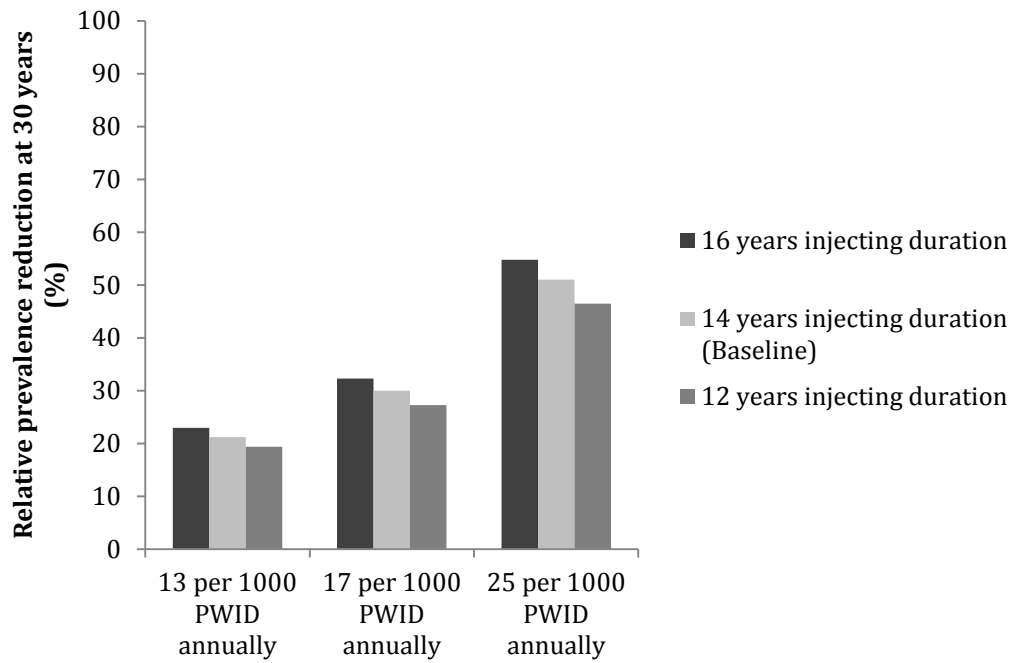
New injectors enter the PWID population at a fixed rate θ , and leave each compartment (due to death or ceasing injection) proportional to the rate μ . Susceptible PWID can become infected at a rate which is proportional to the number of susceptibles, the fraction of the population chronically infected, and the infection rate, π . The acute infection spontaneously clears in the proportion δ , a fraction of which become immune at a proportion ξ . The remaining infected fraction which do not spontaneously clear, $1-\delta$, progress to chronic infection.

Chronically infected PWID can move into treatment at a rate $f(C)$. In the treatment function, infected PWID are recruited onto treatment at a fixed rate, Φ people per 1000 PWID annually. If the infected population is driven below Φ people, all the infected PWID are treated. PWID exit treatment at a rate ω . A proportion of those treated do not respond (proportion $'1-\alpha'$) and move from the treatment compartment to the non-responder compartment. The non-responders cannot be retreated, but can leave the PWID population proportional to the same exit rate as the other PWID, μ . In this model, both compartments of chronic infections contribute to the spread of the infection to susceptibles. The remaining proportion respond to treatment and are cured, a fraction of which (σ) become susceptible again with the remainder $(1-\sigma)$ becoming immune.

Table 1. **Model parameters and values.** ^aWeighted average of genotype distribution and SVR rates. $\alpha = g_1 * \alpha_1 + g_{2/3} * \alpha_{2/3}$. ^bWeighted average of genotype distribution and treatment durations (assuming 24 weeks genotype 2/3, 48 weeks genotype 1). $1/\omega = (48 * g_1 + 24 * g_{2/3}) / 52$. ^cSum of the cessation and death rates. $\mu = \mu_1 + \mu_2$. ^dModel solved until steady state with this baseline treatment rate, and equilibrium values used as initial conditions for model projections. ^eInitial conditions calculated from model equilibrium values with current baseline treatment rates in Victoria.

Symbol	Definition	Value	Units
g_1	Proportion population genotype 1	0.56	-
$g_{2/3}$	Proportion population genotype 2/3	0.44	-
δ	Proportion who spontaneously clear the acute stage	0.26	-
α	Sustained viral response (SVR) Baseline ^a		-
α_1	Genotype 1	0.45	-
$\alpha_{2/3}$	Genotype 2/3	0.80	-
$1/\omega$	Average duration of treatment	0.72 ^b	years
μ	Exit rate from injecting	0.08 ^c	Per year
$1/\mu_1$	Average duration of injecting career	14	years
μ_2	Average PWID death rate	0.0083%	Per year
θ	New PWID entry rate	Fit to total population of 1000 injectors	Per year
π	Infection rate	Fit to give 50% chronic prevalence	Per year
Φ	Antiviral treatment rate		
	Baseline in Victoria ^d	1 per 1000 PWID	Per year
	Model projections ^e	5-40 per 1000 PWID	Per year
ξ	Proportion who become immune after spontaneous clearance	0	-
$1-\sigma$	Proportion who become immune after SVR	0	-

Figure 1. Relative reduction in chronic HCV prevalence at 30 years depending on the length of injecting and treatment rate.



* The baseline prevalence is 50%, and baseline treatment rate 1/1000 PWID annually.

1. Martin NK, Vickerman P, Foster GR, Hutchinson SJ, Goldberg DJ, Hickman M. Can antiviral therapy for hepatitis C reduce the prevalence of HCV among injecting drug user populations? A modelling analysis of its prevention utility. *Journal of Hepatology* 2011;54:1137-1144.