The Models used

The reproductive power model.

The early stage of an outbreak can be described by the infectiousness of the outbreak process, but in later stages of the outbreak, this is complicated by factors such as changing contact patterns and the impact of control measures. It is important to take these factors into account in order to get a good, if approximate, model for an outbreak process. For the covid-19 outbreak in the Netherlands, a non-homogeneous birth process is used for the (quasi) reproductive power function, similar to models in discrete survival analysis. A baseline reproductive power function gives a description of the outbreak.

The non-homogeneous birth model (negative binomial) for outbreak data depends on the reproductive power (probability) ("Modelling the reproductive power function", van den Broek, J., 2020, Journal of Applied Statistics). This reproductive power is taken to be non-homogeneous in order to deal with developments later on in the outbreak. This non-homogeneous is important because, it allows the (quasi) reproductive power to adapt to other dynamics besides the infectious disease outbreak; dynamics such as changing contact patterns, the changing population of susceptible individuals, and control measures taken. Furthermore, in the case of a non-homogeneous reproductive power, there is no need to include a homogeneous mixing assumption. This is because such non-homogeneous mixing – for example if there are periods in which the infected individuals mix well with other individuals and periods in which this is less the case – can influence the number of infected individuals at certain time points and the non-homogeneous reproductive power can adapt to it. The model has the following features:

- The model is a non-homogeneous birth model so it can describe the early phase of the outbreak well and, due to its non-homogeneous nature, can deal with other aspects of the outbreak such as changing behavior or control measures taken besides.
- It does not need the size of an at-risk population; such a population is often hard to determine and can change during the outbreak.

- An infection is usually observed as having taken place in the past. This model deals with this in a way similar to current status data.
- Because the model is a Markov model, it deals with dependence in the data.
- Modelling the log-odds of the reproductive power probabilities (proportion new cases of all the infections at any one time) is the same as modelling the log of the reproductive power (proportion of new cases per existing case).

 $R_p(t_j)$ is the discrete quasi-reproductive power probability function hereafter referred to as the reproductive power probability function or shortly as the reproductive power probability. This is the probability that an infection will occur at t_j given that it did not occur before that point in time. Or the probability that an previous infected individual will produce an infected at t_j .

The conditional probability of observing z_{t_j} new infected individuals at time t_j given $y_{t_{j-1}}$ infected individuals at the previous time point t_{j-1} is:

$$P(Z(t_j) = z_{t_j}|Y(t_{j-1}) = y_{t_{j-1}}) = \begin{pmatrix} z_{t_j} + y_{t_{j-1}} - 1 \\ y_{t_{j-1}} - 1 \end{pmatrix} [1 - R_p(t_j)]^{y_{t_{j-1}}} [R_p(t_j)]^{z_{t_j}},$$

$$z_{t_i} = 0, 1, \dots$$

The expected value is:

$$E[Z(t_j)] = \mu_{t_j} = y_{t_{j-1}} \frac{R_p(t_j)}{1 - R_p(t_j)},$$
(1)

Taking a log link for the expected value of the number of new cases $(Z(t_j))$, at time interval t_j , given the total number of cases in the previous interval, gives:

$$\log[\mu_{t_j}] = \log[y_{t_{j-1}}] + \log\left[\frac{R_p(t_j)}{1 - R_p(t_j)}\right]$$

One can assume a parametric model for the survival function and use this to calculate the reproductive power probability function, as was done in (Van Den Broek, J. and Heesterbeek, J.A.P. (2007) "Non-Homogeneous Birth and Death Models for Epidemic Outbreak Data". Biostatistics8, 453-467) with

members from the Burr-family. To avoid this parametric assumption, one can model the time effects with a piece wise constant function in discrete time. $\$ The log-odds of the reproductive probability can be modelled linearly in the covariates. If the covariates all have baseline values (usually zero), the model for the log-odds of the base line reproductive probability, $R_p(t_i)$, is

$$\log\left[\frac{R_p(t_j)}{1 - R_p(t_j)}\right] = \alpha_{t_j},$$

so α_{t_j} is the log-odds of the reproductive power probability at time t_j , and

$$R_p(t_j) = \frac{e^{\alpha_{t_j}}}{1 - e^{\alpha_{t_j}}}.$$

An extension: A Markov-switching model

A hidden Markov process consists of two parts. The first is an unobserved parameter process. So the reproductive power function might change over time because the underlying probability goes through a number of states. These states form a Markov model. So the state on time-point t depends only on the previous time point. The transition matrix determines the probability to go from one state to another.

The second part is the state-dependent part. Given the state it gives the probability of observing a number of new cases. For more details see (Zucchini w., Macdonald I.L. and Langrock R.(2016), "Hidden Markov Models for TimeSeries. An Introduction Using R". Second edition Boca Raton: CRC Press).

The state dependent distribution is taken to be the above negative binomial distribution which depends on the reproductive power probability. Since this distribution also depends on the history in a Markov way (it depends on the total number of infected in a previous period), in this model there are 2 of these dependencies, one in the state dependent distribution and one in the parameter process. Such a model is called a Markov switching model (see chapter 10. of Zucchine at.all.).

One is often interested in the most likely sequence of hidden states. This global decoding is achieved by maximizing the conditional probability of all the states given the data. This can be calculated using the Viterbi algorithm. (see chapter 5.4 about decoding in Zucchine at.all.)