About Statistics of Disease Dynamics

LiJin Joo lijin.joo at gmail.com

2020-09-14

Reproduction number (R) and epidemic growth rate (r) are two statistics to describe the disease dynamics, or transmission of COVID-19 completely and neither of the two is replaceable to the other. Reproduction number R tells a direction of growth, whether an epidemic is growing or shrinking, yet does not tell how fast a series of cases is growing at each time point. On the other hand, growth rate r provides the information on both size and speed of change but does not predict how likely a number of cases will grow or decline at the next time point. The goal of measuring the transmission is to assess the effectiveness of public health intervention, such as lockdown, and to predict the impact of relaxing an intervention (Anderson et al. 2020) (Science and Scientific Advisory Group for Emergencies 2020).

Reproduction Number (R)

The reproduction number (R) is the average number of secondary infections produced by a single infected person and shows the potential of transmission. If R is greater than 1, every infected patient would produce more than one new case on average, thus the disease continues to spread. If R is smaller than 1, a new case from an existing patient is less than one on average, thus the disease would shrink and eventually stop.

R is also called "time-varying" reproduction number, Rt, or "effective" reduction number Re, to emphasize its ability to reflect time varying parameters, thus, to evaluate the impact of a public health intervention, or evolution of viruses. In contrast to R, the basic reproduction number, R0, is the number of secondary infections produced by a single individual at the beginning of an epidemic and shows the potential of disease transmission in a population never exposed to the pathogen (Delamater et al. 2019). As more individuals are infected over time, Rt include both types of new cases from a population susceptible and from an exposed/immune population. Rt is always less than R0 (Anderson et al. 2020).

Various models for transmission of infection and methods to estimate parameters exist, which uses different approaches to infer cases from susceptible, incubation, infectious and recovered (SEIR) populations (Anderson et al. 2020). We used the method developed by (Cori et al. 2013), which takes a series of incidences only and learns the values of necessary parameters by its empirical density and Bayesian model assumptions. The formulas are provided by R package {EpiEstim}.

- We used estimate_R() function with a "parametric_si" option assuming a gamma prior with a mean of 4 days and a standard deviation of 4.75 days and 7 days of smoothing window are used suggested in (Abbott et al. 2020).
- In the previous studies, a gamma prior with a mean of 7 days and a standard deviation of 4.5 days with 4 days of smoothing window is suggested in (Basu et al. 2020). However, this old prior gives a unreasonably small variation (sd < 0.01) for recent data where Rt is lingering around 1 and new cases remain large.
- The formula for the time varying reproduction number Rt from {EpiEstim} is

$$R_t = \frac{I_t}{\sum_{s=1}^t I_{t-s} \omega_s}$$

where I_t is the incidence at time t and ω is a density function of time since infection at s and can be estimated from serial intervals, the average time between symptoms of infection in the transmitter and

newly infected. Often, Gamma is assumed.

 I_t is determined by past cases I_0, I_1, \dots, I_{t-1} , which follows a Poisson model:

$$P(I_t|I_0, I_1, \dots, I_{t-1}, \omega, R_t) = \frac{(R_t \Lambda_t)^{I_t}}{I_t!}$$

with $\Lambda_t = \sum_{s=1}^t I_{t-s} \omega_s$.

• As I_t is determined by a stochastic process, so R_t is. At time t, R_t is right censored as the unreported cases due to 1) the delay in reporting, 2) incubation period of a new case, 3) generation time, a time between infection events. The three additional parameters are additionally considered in {EpiNow2} by (Abbott et al. 2020). One major improvement by additional parameters is more sensible uncertainty estimates.

Growth Rate (r)

The growth rate r shows how quickly the number of infections is changing and is calculated by the second order derivative of the incidence. The growth rate represents who many new cases are occurring for the next time point under the exponential growth assumption and can be an early or instantaneous indicator for the impact of a new intervention, such as a release in lock down. If the growth rate is greater than zero (or positive), the epidemic is growing. If the growth rate is less than zero (or negative) then the epidemic is declining (Anderson et al. 2020).

- Calculation of r is simpler than that of Rt. However, its interpretation on an epidemic growth curve from raw incidence is not straightforward due to the non-linear and non-parametric characteristics of the curve. Instead, we provide two more statistics based on r to illustrate the implications on the change in growth rate.
- Doubling Time (d_t) : We used the formula from (Basu et al. 2020) as follows.

$$d_t = \tau \frac{log(2)}{log(1+r)}$$

where the growth rate r is combined by

$$r = \frac{I_t - I_{t-1}}{I_{t-1}}$$

for t = 0, 1, ..., T and τ is a window size of smoothing. We used $\tau = 7$ days.

This indicates that the doubling time for Sep 7 represents how long would take to observe cases doubled, if it increases at a constant growth between Sep 1 to Sep 7.

• Levitt Statistics: We compute Levitt statistics (Levitt, Scaiewicz, and Zonta 2020) for deaths only.

$$H(t) = log(\frac{X_t}{X_{t-1}})$$

where X_t is a cumulative count until day t. The statistics shows a rate of incidence changes. If H(t) < 0, a time of a new infection is slow down. H(t) enjoys a great compatibility being free of model assumptions, yet its application to incidence is warranted in (Raman 2020) due to the influence of testing capacities.

Reference

Abbott, Sam, Joel Hellewell, Robin N Thompson, Katharine Sherratt, Hamish P Gibbs, Nikos I Bosse, James D Munday, et al. 2020. "Estimating the Time-Varying Reproduction Number of Sars-Cov-2 Using National and Subnational Case Counts." Wellcome Open Research 5 (112). F1000 Research Limited: 112.

Anderson, Roy, Christl Donnelly, Deirdre Hollingsworth, Matt Keeling, Vegvari Carolin, Rebecca Baggaley, and Rosie Maddren. 2020. "Reproduction Number (R) and Growth Rate (R) of the Covid-19 Epidemic

in the Uk: Methods of Estimation, Data Sources, Causes of Heterogeneity, and Use as a Guide in Policy Formulation." Royal Society SET-C (Science in Emergencies Tasking – COVID). https://royalsociety.org/-/media/policy/projects/set-c/set-covid-19-R-estimates.pdf; The Royal Society.

Basu, Deepankar, Maxwell Salvatore, Debashree Ray, Michael Kleinsasser, Soumik Purkayastha, Rupam Bhattacharyya, and Bhramar Mukherjee. 2020. "A Comprehensive Public Health Evaluation of Lockdown as a Non-Pharmaceutical Intervention on Covid-19 Spread in India: National Trends Masking State Level Variations." medRxiv. Cold Spring Harbor Laboratory Press. https://doi.org/10.1101/2020.05.25.20113043.

Cori, Anne, Neil M Ferguson, Christophe Fraser, and Simon Cauchemez. 2013. "A New Framework and Software to Estimate Time-Varying Reproduction Numbers During Epidemics." *American Journal of Epidemiology* 178 (9). Oxford University Press: 1505–12.

Delamater, Paul L, Erica J Street, Timothy F Leslie, Y Tony Yang, and Kathryn H Jacobsen. 2019. "Complexity of the Basic Reproduction Number (R0)." *Emerging Infectious Diseases* 25 (1). https://wwwnc.cdc.gov/eid/article/25/1/17-1901_article; Centers for Disease Control; Prevention: 1.

Gostic, Katelyn M, Lauren McGough, Edward Baskerville, Sam Abbott, Keya Joshi, Christine Tedijanto, Rebecca Kahn, et al. 2020. "Practical Considerations for Measuring the Effective Reproductive Number, Rt." medRxiv. Cold Spring Harbor Laboratory Press. https://doi.org/10.1101/2020.06.18.20134858.

Levitt, Michael, Andrea Scaiewicz, and Francesco Zonta. 2020. "Predicting the Trajectory of Any Covid19 Epidemic from the Best Straight Line." medRxiv. Cold Spring Harbor Laboratory Press.

Obadia, Thomas, Romana Haneef, and Pierre-Yves Boëlle. 2012. "The R0 Package: A Toolbox to Estimate Reproduction Numbers for Epidemic Outbreaks." *BMC Medical Informatics and Decision Making* 12 (1). BioMed Central: 1–9.

Raman, Bhaskaran. 2020. "When Will the Covid-19 Pandemic End? Levitt's Metric on Indian Data."

Science, Government Office for, and UK Scientific Advisory Group for Emergencies. 2020. "The R Number and Growth Rate in the Uk." https://www.gov.uk/guidance/the-r-number-in-the-uk. 2020.

Wallinga, Jacco, and Peter Teunis. 2004. "Different Epidemic Curves for Severe Acute Respiratory Syndrome Reveal Similar Impacts of Control Measures." *American Journal of Epidemiology* 160 (6). Oxford University Press: 509–16.