

# Generation intervals in space

Understanding the effects of spatial and network structure on links between generation interval and growth rate

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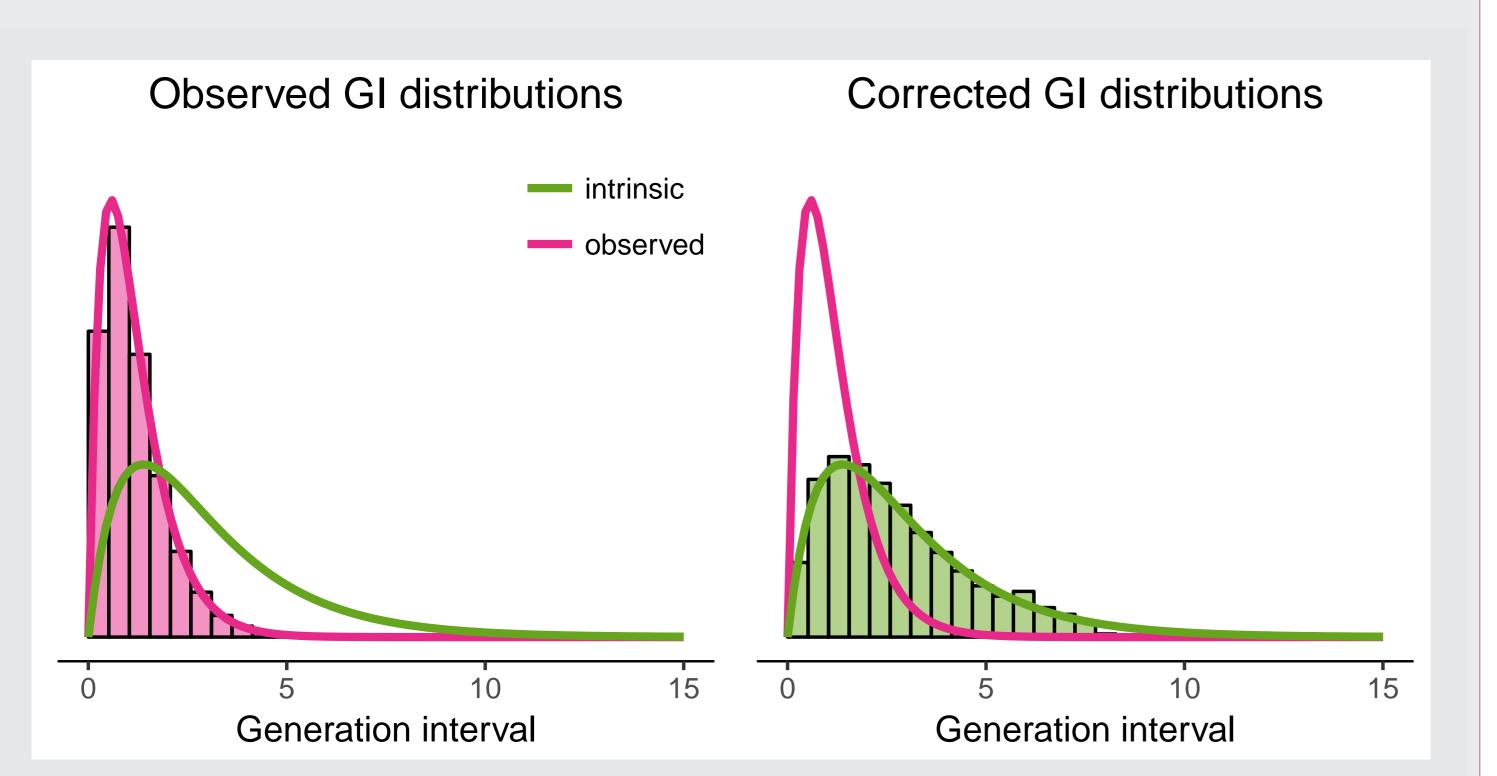
### Background

- ► Generation interval (GI) measures time between when a person is infected and when that person infects another person
- ► GI distribution,  $g(\tau)$ , links exponential growth rate, r, and reproductive number,  $\mathcal{R}$ , of an epidemic [1]:  $1/\mathcal{R} = \int g(\tau) \exp(-r\tau) d\tau$
- ▶ If r is known, longer generation times imply higher estimates of  $\mathcal{R}$ .
- ► Measuring GI through contact tracing data can introduce bias [2]
- ► Trapman *et al.* [3] investigated the effects of network structure on the relationship between r and  $\mathcal{R}$ .

#### Overview Homogeneous Intrinsic GI Observed GI in early epidemic assumption • patient-based • contact-tracing based Temporal • infectiousness profile of an censored at observation time infected individual correction Local correc-Temporal correction | weight observed periods by $\exp(r\tau)$ Locally corrected GI Effective GI • based on degree distribution + spatial structure • reflects network structure, but and contact rate [3] corrects for time censoring • depends on between-individual • gives the correct link between r and $\mathcal{R}$ variation

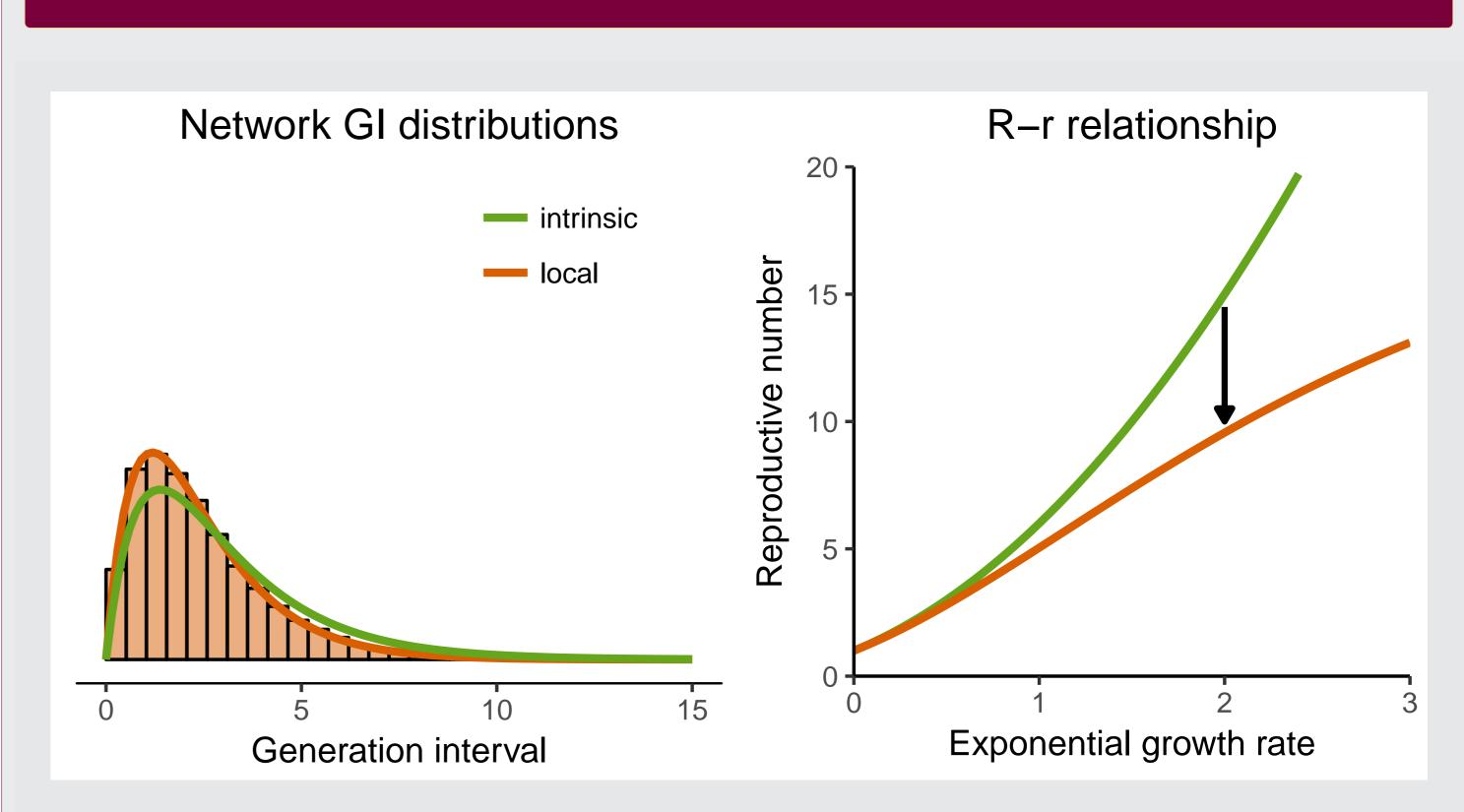
Patient-based GI observations do not account for spatial effects, while contact tracing introduces temporal bias (especially early in outbreaks).

## Temporal correction on a homogeneous network



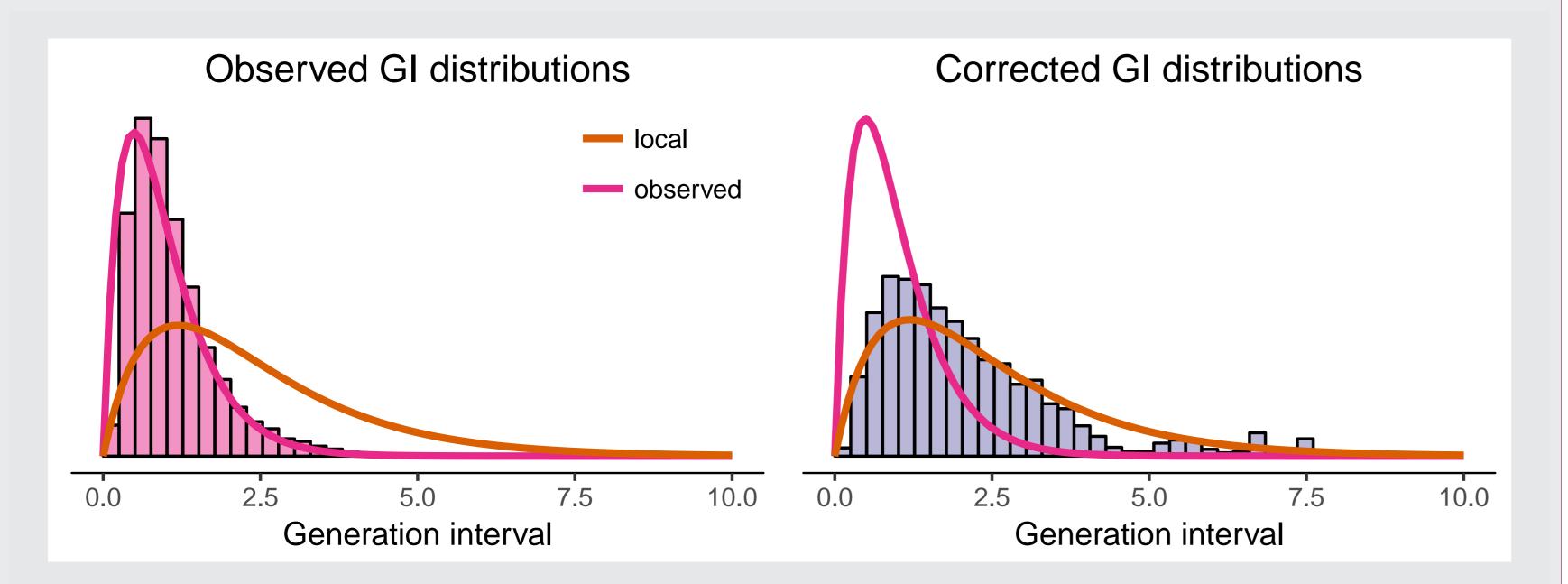
The intrinsic GI distribution can be estimated from observations during an outbreak. During an outbreak, shorter intervals are more likely to be observed (since longer ones may not have completed yet). (Left) Intervals observed during the exponential phase. (Right) The same intervals, but reweighted to remove the sampling bias. Stochastic SEIR simulation on a homogeneous network (histograms) match analytically derived distributions (solid curves). Model parameters:  $\mathcal{R}=6$ , mean latent, infectious periods of 1, 2 time units.

#### Local correction on an idealized network



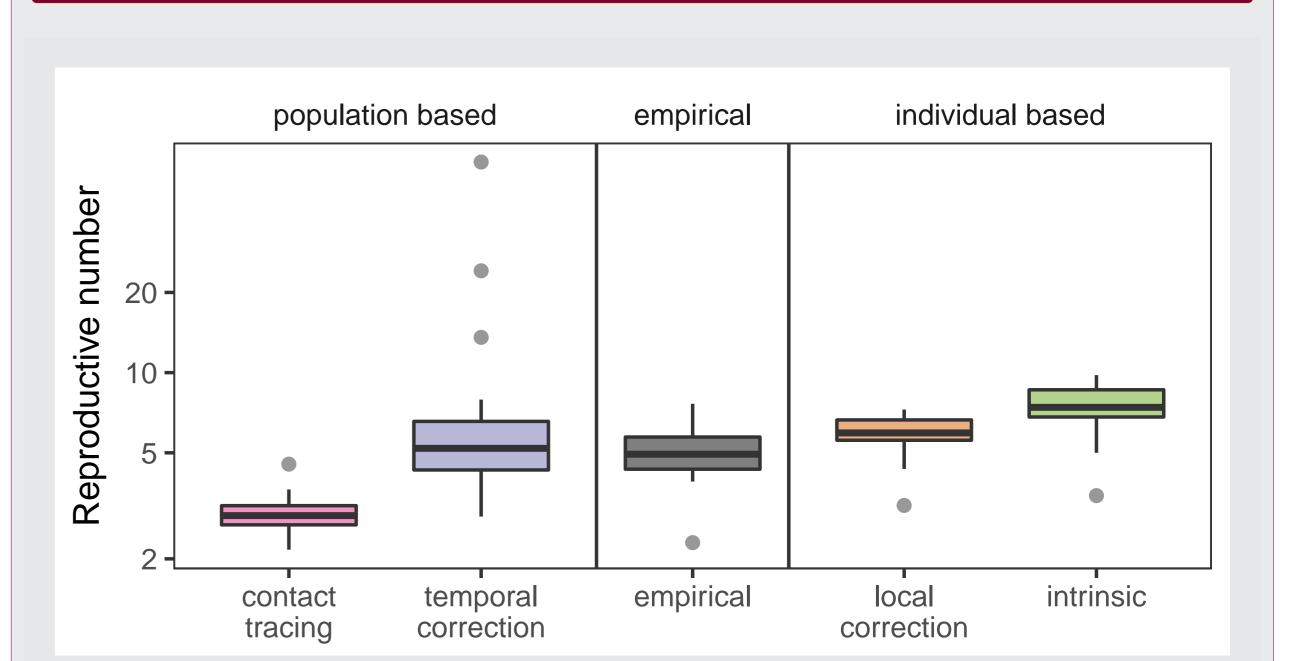
Effects of locality on GI can be calculated in simple cases. Local spatial effects make GIs shorter, because repeated contacts only infect the first time. (Left) Observed GIs from simulations on idealized, strictly local networks. These do not match the known intrinsic GI, but do match the calculated local correction. (Right) If a disease is spreading on a network, calculations based on the (longer) intrinsic GI will lead to over-estimates of  $\mathcal{R}$ . The local correction accounts for local effects, but not other network effects [3].

#### Estimating GIs on an empirical network



**Local correction partially accounts for spatial effects on an empirical network** [4]. This correction requires an estimate of degree distribution, and either an explicit model structure (like SEIR) or information about individual-level variation in infectiousness.

#### Estimating R on an empirical network



Appropriate corrections improve estimates. Estimates of R using directly observed Gls either result in under-estimates (contact tracing) or over-estimates (intrinsic). Temporal correction of censored contact-tracing Gls can produce unstable estimates. Local correction of intrinsic Gls [3] is difficult and approximate.

#### Conclusion and future direction

- ► Different approaches measure different Gls
- Naive use of intrinsic GI may over-estimate  $\mathcal{R}$ , while naive use of contact-tracing GI may underestimate  $\mathcal{R}$ . We can understand how spatial and temporal effects change

#### Bibliography

- [1] Jacco Wallinga and Marc Lipsitch. How generation intervals shape the relationship between growth rates and reproductive numbers. Proceedings of the Royal Society of London B: Biological Sciences, 274(1609):599–604, 2007.
- [2] David Champredon and Jonathan Dushoff. Intrinsic and realized generation intervals in infectious-disease transmission. 282(1821), 2015.
- [3] Pieter Trapman, Frank Ball, Jean-Stéphane Dhersin, Viet Chi Tran, Jacco Wallinga, and Tom Britton. Inferring r0 in emerging epidemicsthe