

# RtEstim: Effective reproduction number estimation with trend filtering

Jiaping Liu<sup>1\*</sup>, Zhenglun Cai<sup>2</sup>, Paul Gustafson<sup>1</sup>, Daniel J. McDonald<sup>1</sup>

**1** Department of Statistics, The University of British Columbia, Vancouver, British Columbia, Canada

**2** Centre for Health Evaluation and Outcome Sciences, The University of British Columbia, Vancouver, British Columbia, Canada

\* [jiaping.liu@stat.ubc.ca](mailto:jiaping.liu@stat.ubc.ca)

## Abstract

To understand the transmissibility and spread of infectious diseases, epidemiologists turn to estimates of the effective reproduction number. While many estimation approaches exist, their utility may be limited. Challenges of surveillance data collection, model assumptions that are unverifiable with data alone, and computationally inefficient frameworks are critical limitations for many existing approaches. We propose a discrete spline-based approach **RtEstim** that solves a convex optimization problem—Poisson trend filtering—using the proximal Newton method. It produces a locally adaptive estimator for effective reproduction number estimation with heterogeneous smoothness. **RtEstim** remains accurate even under some process misspecifications and is computationally efficient, even for large-scale data. The implementation is easily accessible in a lightweight R package [rtestim](#).

## Author summary

Effective reproduction number estimation presents many challenges due to data collection, modelling assumptions, and computational burden. Such limitations hinder the accurate estimation of the effective reproduction number. Our motivation is to

develop a model that produces accurate estimates, is robust to model misspecification, and is straightforward to use and computationally efficient, even for large counts and long time periods. We propose a convex optimization model with an  $\ell_1$  trend filtering penalty. It couples accurate estimation of the effective reproduction number with desired smoothness. We solve the optimization using the proximal Newton method, which converges rapidly and is numerically stable. Our software, conveniently available in the R package `RtEstim`, can produce estimates in seconds for incidence sequences with hundreds of observations. These estimates are produced for a sequence of tuning parameters and can be selected using a built-in cross validation procedure.

## 1 Introduction

The effective reproduction number is defined to be the average number of secondary infections caused by a primary infection that occurred sometime in the past. Also called the instantaneous reproduction number, it is a key quantity for understanding infectious disease dynamics including the potential size of an outbreak and the required stringency of control measures. Tracking the time series of this quantity is useful for understanding whether or not future infections are likely to increase or decrease from the current state. Let  $\mathcal{R}(t)$  denote the effective reproduction number at time  $t$ . Practically, as long as  $\mathcal{R}(t) < 1$ , infections will decline gradually, eventually resulting in a disease-free equilibrium, whereas when  $\mathcal{R}(t) > 1$ , infections will continue to increase, resulting in endemic equilibrium. While  $\mathcal{R}(t)$  is fundamentally a continuous time quantity, it can be related to data only at discrete points in time  $t = 1, \dots, n$ . This sequence of effective reproduction numbers over time is not observable, but, nonetheless, is easily interpretable and retrospectively describes the course of an epidemic. Therefore, a number of procedures exist to estimate  $\mathcal{R}_t$  from different types of observed incidence data such as cases, deaths, or hospitalizations, while relying on various domain-specific assumptions. Importantly, accurate estimation of effective reproduction numbers relies heavily on the quality of the available data, and, due to the limitations of data collection, such as underreporting and lack of standardization, estimation methodologies rely on various assumptions to compensate. Because model assumptions may not be easily verifiable from data alone, it is also critical for any estimation procedure to be

robust to model misspecification.

Many existing approaches for effective reproduction number estimation are Bayesian: they estimate the posterior distribution of  $\mathcal{R}_t$  conditional on the observations. One of the first such approaches is the software **EpiEstim** [1], described in [2]. This method is prospective, in that it uses only observations available up to time  $t$  in order to estimate  $\mathcal{R}_t$  for each  $i = 1, \dots, t$ . An advantage of **EpiEstim** is its straightforward statistical model: new incidence data follows the Poisson distribution conditional on past incidence combined with the conjugate gamma prior distribution for  $\mathcal{R}_t$  with fixed hyperparameters. Additionally, the serial interval distribution, the distribution of the period between onsets of primary and secondary infections in a population, is fixed and known. For this reason, **EpiEstim** requires little domain expertise for use, and it is computationally fast. [3] modified this method to distinguish imported cases from local transmission and simultaneously estimate the serial interval distribution. [4] further extended **EpiEstim** by using “reconstructed” daily incidence data to handle irregularly spaced observations. Recently, [5] proposed a Bayesian latent variable framework, **EpiNow2** [6], which leverages incident cases, deaths or other available streams simultaneously along with allowing additional delay distributions (incubation period and onset to reporting delays) in modelling. [7] proposed an extension that handles missing data by imputation followed by a truncation adjustment. These modifications are intended to increase accuracy at the most recent (but most uncertain) timepoints, to aid policymakers. [8] also proposed a Bayesian approach, **EpiFilter** based on the (discretized) Kalman filter and smoother. **EpiFilter** also estimates the posterior of  $\mathcal{R}_t$  given a Gamma prior and Poisson distributed incident cases. Compared to **EpiEstim**, however, **EpiFilter** estimates  $\mathcal{R}_t$  retrospectively using all available incidence data both before and after time  $t$ , with the goal of being more robust in low-incidence periods. [9] proposed a Bayesian P-splines approach, **EpiLPS**, that assumes negative Binomial distributed observations. [10] also proposed a Bayesian model estimated with particle filtering to incorporate spatial structures. Bayesian approaches estimate the posterior distribution of the effective reproduction numbers and possess the advantage that credible intervals may be easily computed. A limitation of many Bayesian approaches, however, is that they usually require more intensive computational routines, especially when observed data sequences are long or hierarchical structures are complex. Below,

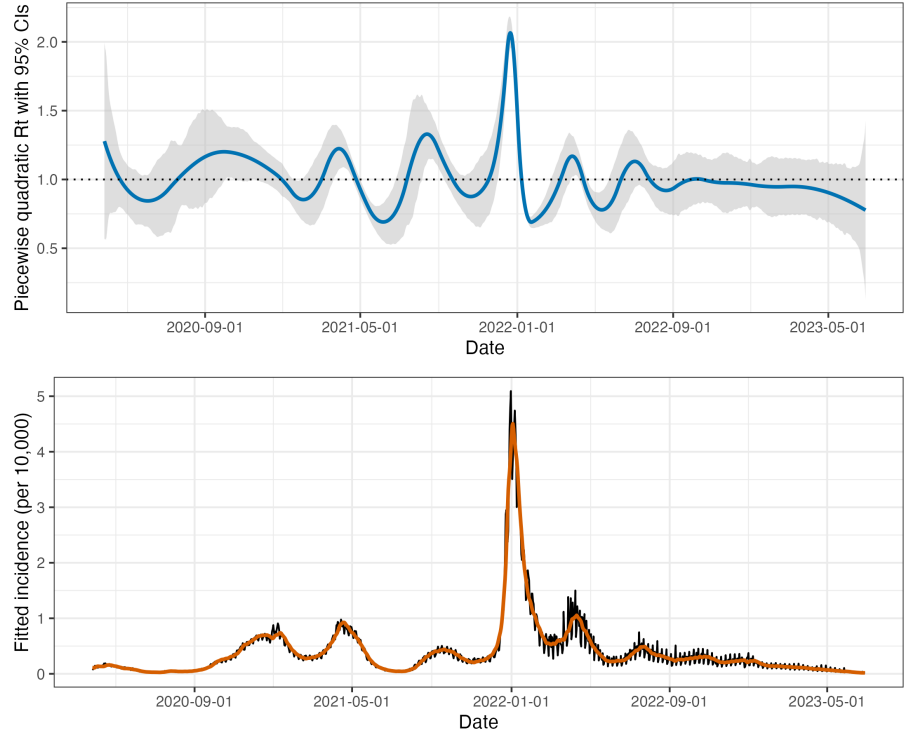
we compare our method to two of the more computationally efficient Bayesian models, EpiEstim and EpiLPS.

There are also frequentist approaches for  $\mathcal{R}_t$  estimation. [11] proposed regularizing the smoothness of  $\mathcal{R}_t$  through penalized regression with second-order temporal regularization, additional spatial penalties, and with Poisson loss. [12] extended this procedure by introducing another penalty on outliers. [13] proposed a spline-based model relying on the assumption of exponential-family distributed incidence. [14] estimates  $\mathcal{R}_t$  while monitoring the time-varying level of overdispersion. There are other spline-based approaches such as [15, 16], autoregressive models with random effects [17] that are robust to low incidence, and generalized autoregressive moving average (GARMA) models [18] that are robust to measurement errors in incidence data.

We propose a retrospective effective reproduction number estimator called **RtEstim** that requires only incidence data. Our model makes the conditional Poisson assumption, similar to much of the prior work described above, but is empirically more robust to misspecification. This estimator is defined by a convex optimization problem with Poisson loss and  $\ell_1$  penalty on the temporal evolution of  $\log(\mathcal{R}_t)$  to impose smoothness over time. As a result, **RtEstim** generates discrete splines, and the estimated curves (in logarithmic space) appear to be piecewise polynomials of an order selected by the user. Importantly, the estimates are locally adaptive, meaning that different time ranges may possess heterogeneous smoothness. Because we penalize the logarithm of  $\mathcal{R}_t$ , we naturally accommodate the positivity requirement, in contrast to related methods, can handle large or small incidence measurements, and are automatically (reasonably) robust to outliers without additional constraints. A small illustration using three years of Covid-19 case data in Canada is shown in Fig 1 [19].

While our approach is straightforward and requires little domain knowledge for implementation, we also implement a number of refinements. We use a proximal Newton method to solve the convex optimization problem along with warm starts to produce estimates efficiently, typically in a matter of seconds, even for long sequences of data. In a number of simulation experiments, we show empirically that our approach is more accurate than existing methods at estimating the true effective reproduction numbers.

The manuscript proceeds as follows. We first introduce the methodology of **RtEstim** including the renewal equation and the development of Poisson trend filtering estimator.



**Fig 1.** A demonstration of effective reproduction number estimation by `RtEstim` and the corresponding predicted incident cases for the Covid-19 epidemic in Canada during the period from March 28, 2020 to June 28, 2023. In the top panel, the blue curve is the estimated piecewise quadratic  $\mathcal{R}_t$  and the gray ribbon is the corresponding 95% confidence band. The black curve in the bottom panel is the observed Covid-19 daily confirmed cases, and the orange curve is the predicted incident cases corresponding to the estimated  $\mathcal{R}_t$ .

We explain how this method could be interpreted from the Bayesian perspective, 86  
connecting it to previous work in this context. We provide illustrative experiments 87  
comparing our estimator to `EpiEstim` and `EpiLPS`. We then apply our `RtEstim` on the 88  
Covid-19 pandemic incidence in British Columbia and the 1918 influenza pandemic 89  
incidence in the United States. Finally, we conclude with a discussion of the advantages 90  
and limitations of our approach and describe practical considerations for effective 91  
reproduction number estimation. 92

## 2 Methods

### 2.1 Renewal model for incidence data

The effective reproduction number  $\mathcal{R}(t)$  is defined to be the expected number of secondary infections at time  $t$  produced by a primary infection sometime in the past. To make this precise, denote the number of new infections at time  $t$  as  $y(t)$ . Then the total primary infectiousness can be written as  $\eta(t) := \int_0^\infty p(i)y(t-i)di$ , where  $p(i)$  is the probability that a new secondary infection is the result of a primary infection that occurred  $i$  time units in the past. The effective reproduction number is then given as the value that equates

$$\mathbb{E}[y(t) \mid y(j), j < t] = \mathcal{R}(t)\eta(t) = \mathcal{R}(t) \int_0^\infty p(i)y(t-i)di, \quad (1)$$

otherwise known as the renewal equation. The period between primary and secondary infections is exactly the generation time of the disease, but given real data, observed at discrete times (say, daily) this delay distribution must be discretized into contiguous time intervals, say,  $(0, 1], (1, 2], \dots$ , which results in the sequence  $\{p_i\}_0^\infty$  corresponding to observations  $y_t$  and resulting in the discretized version of Eq (1),

$$\mathbb{E}[y_t \mid y_1, \dots, y_{t-1}] = \mathcal{R}_t \eta_t = \mathcal{R}_t \sum_{i=1}^\infty p_i y_{t-i}. \quad (2)$$

Many approaches to estimating  $\mathcal{R}_t$  rely on Eq (2) as motivation for their procedures, among them, **EpiEstim** [2] and **EpiFilter** [8].

In most cases, it is safe to assume that infectiousness disappears beyond  $\tau$  timepoints ( $p(i) = 0$  for  $i > \tau$ ), resulting in the truncated integral of the generation interval distribution  $\int_0^\tau p(i)di = 1$ . Generation time, however, is usually unobservable and tricky to estimate, so common practice is to approximate it by the serial interval: the period between the symptom onsets of primary and secondary infections. If the infectiousness profile after symptom onset is independent of the incubation period (the period from the time of infection to the time of symptom onset), then this approximation is justifiable: the serial interval distribution and the generation interval distribution share the same mean. However, other properties may not be similarly

shared, and, in general, the generation interval distribution is a convolution of the serial  
interval distribution with the distribution of the difference between independent draws  
from the delay distribution from infection to symptom onset. See, for example, [20] for  
a fuller discussion of the dangers of this approximation. Nonetheless, treating these as  
interchangeable is common [2] and doing otherwise is beyond the scope of this work.  
Additionally, we assume that the generation interval (and, therefore, the serial interval),  
is constant over time  $t$ . That is, the probability  $p(i)$  depends only on the gap between  
primary and secondary infections and not on the time  $t$  when the secondary infection  
occurs. For our methods, we will assume that the serial interval can be accurately  
estimated from auxiliary data (say by contact tracing, or previous epidemics) and we  
will take it as fixed, as is common in existing studies, e.g., [2, 11, 12].

The renewal equation in Eq (2) relates observable data streams (incident cases)  
occurring at different timepoints to the effective reproduction number given the serial  
interval. The fact that it depends only on the observed incident counts makes it  
reasonable to estimate  $\mathcal{R}_t$ . However, data collection idiosyncrasies can obscure this  
relationship. Diagnostic testing targets symptomatic individuals, omitting  
asymptomatic primary infections which can lead to future secondary infections. Testing  
practices, availability, and uptake can vary across space and time [21, 22]. Finally,  
incident cases as reported to public health are subject to delays due to laboratory  
confirmation, test turnaround times, and eventual submission to public health [23]. For  
these reasons, reported cases are lagging indicators of the course of the pandemic.  
Furthermore, they do not represent the actual number of new infections that occur on a  
given day, as indicated by exposure to the pathogen. The assumptions described above  
(constant serial interval distribution, homogenous mixing, similar susceptibility and  
social behaviours, etc.) are therefore consequential. That said, Eq (2) also provides some  
comfort about deviations from these assumptions. If  $y_t$  is scaled by a constant (in time)  
describing the reporting ratio, then it will cancel from both sides. Similar arguments  
mean that even if such a scaling varies in time, as long as it varies slowly relative to the  
set of  $p_i$  that are larger than 0, Eq (2) will be a reasonably accurate approximation, so  
that  $\mathcal{R}_t$  can still be estimated well from reported incidence data. Finally, even a sudden  
change in reporting ratio, say from  $c_1$  for  $i = 1, \dots, t_1$  to  $c_2$  for  $i > t_1$  would only result  
in large errors for  $t$  in the neighbourhood of  $t_1$  (where the size of this neighbourhood is

again determined by the effective support of  $\{p_i\}$ ). This robustness to certain types of data reporting issues partially justifies using Eq (2) to calculate  $\mathcal{R}_t$ .

## 2.2 Poisson trend filtering estimator

We use the daily confirmed incident cases  $y_t$  on day  $t$  to estimate the observed infectious cases under the model that  $y_t$ , given previous incident cases  $y_{t-1}, \dots, y_1$  and a constant serial interval distribution, follows a Poisson distribution with mean  $\Lambda_t$ . That is,

$$y_t \mid y_1, \dots, y_{t-1} \sim \text{Poisson}(\Lambda_t), \text{ where } \Lambda_t = \mathcal{R}_t \sum_{i=1}^{t-1} p_i y_{t-i} = \mathcal{R}_t \eta_t. \quad (3)$$

Given a history of  $n$  confirmed incident counts  $\mathbf{y} = (y_1, \dots, y_n)^\top$ , our goal is to estimate  $\mathcal{R}_t$ . A natural approach is to maximize the likelihood, producing the maximum likelihood estimator (MLE):

$$\begin{aligned} \hat{\mathcal{R}} &= \underset{\mathcal{R} \in \mathbb{R}_+^n}{\operatorname{argmax}} \mathbb{P}(\mathcal{R} \mid \mathbf{y}, \mathbf{p}) = \underset{\mathcal{R} \in \mathbb{R}_+^n}{\operatorname{argmax}} \prod_{t=1, \dots, n} \frac{(\mathcal{R}_t \eta_t)^{y_t} \exp\{-\mathcal{R}_t \eta_t\}}{y_t!} \\ &= \underset{\mathcal{R} \in \mathbb{R}_+^n}{\operatorname{argmin}} \sum_{t=1}^n \mathcal{R}_t \eta_t - y_t \log(\mathcal{R}_t \eta_t). \end{aligned} \quad (4)$$

This optimization problem, however, is easily seen to yield a one-to-one correspondence between the observations and the estimated effective reproduction number, i.e.,  $\hat{\mathcal{R}}_t = y_t / \eta_t$ , so that the estimated sequence  $\hat{\mathcal{R}}$  will have no significant smoothness.

The MLE is an unbiased estimator of the true parameter  $\mathcal{R}_t$ , but unfortunately has high variance: changes in  $y_t$  result in proportional changes in  $\hat{\mathcal{R}}_t$ . To avoid this behaviour, and to match the intuition that  $\mathcal{R}_t \approx \mathcal{R}_{t-1}$ , we advocate enforcing smoothness of the effective reproduction numbers. This constraint will decrease the estimation variance, and hopefully lead to more accurate estimation of  $\mathcal{R}$ , as long as the smoothness assumption is reasonable. Smoothness assumptions are common (see e.g., [8] or [20]), but the type of smoothness assumed is critical. [1] imposes smoothness indirectly by estimating  $\mathcal{R}_t$  with moving windows of past observations. The Kalman filter procedure of [8] would enforce in  $\ell_2$ -smoothness  $(\int_0^n (\hat{\mathcal{R}}''(t))^2 dt < C$  for some  $C$ ), although the computational implementation results in  $\hat{\mathcal{R}}$  taking values over a discrete grid. [12] produces piecewise linear  $\hat{\mathcal{R}}_t$ , which turns out to be closely related to a special



case of our methodology. Smoother estimated curves will provide high-level information about the entire epidemic, obscuring small local changes in  $\mathcal{R}(t)$ , but may also remove the ability to detect large sudden changes, such as those resulting from lockdowns or other major containment policies.

To enforce smoothness of  $\hat{\mathcal{R}}_t$ , we add a trend filtering penalty to Eq (5) [24–27]. Because  $\mathcal{R}_t > 0$ , we explicitly penalize the divided differences (discrete derivatives) of neighbouring values of  $\log(\mathcal{R}_t)$ . Let  $\theta := \log(\mathcal{R}) \in \mathbb{R}^n$ , so that  $\Lambda_t = \eta_t \exp(\theta_t)$ , and  $\log(\eta_t \mathcal{R}_t) = \log(\eta_t) + \theta_t$ . For evenly spaced incident case, we write our estimator as the solution to the optimization problem

$$\hat{\mathcal{R}} = \exp(\hat{\theta}) \quad \text{where} \quad \hat{\theta} = \underset{\theta \in \mathbb{R}^n}{\operatorname{argmin}} \eta^\top \exp(\theta) - \mathbf{y}^\top \theta + \lambda \|D^{(k+1)} \theta\|_1, \quad (5)$$

where  $\exp(\cdot)$  applies elementwise. Here,  $D^{(k+1)} \in \mathbb{Z}^{(n-k-1) \times n}$  is the  $(k+1)^{\text{th}}$  order divided difference matrix for any  $k \in \{0, \dots, n-1\}$ .  $D^{(k+1)}$  is defined recursively as  $D^{(k+1)} := D^{(1)} D^{(k)}$ , where  $D^{(1)} \in \{-1, 0, 1\}^{(n-k-1) \times (n-k)}$  is a sparse matrix with diagonal band:

$$D^{(1)} = \begin{pmatrix} -1 & 1 & & & \\ & -1 & 1 & & \\ & & \ddots & \ddots & \\ & & & -1 & 1 \end{pmatrix}. \quad (6)$$

The tuning parameter (hyperparameter)  $\lambda$  balances data fidelity with desired smoothness. When  $\lambda = 0$ , the problem in Eq (5) reduces to the MLE in Eq (4). Larger tuning parameters privilege the regularization term and yield smoother estimates. Finally, there exists  $\lambda_{\max}$  such that any  $\lambda \geq \lambda_{\max}$  will result in  $D^{(k+1)} \hat{\theta} = 0$  and  $\hat{\theta}$  will be the Kullback-Leibler projection of  $\mathbf{y}$  onto the null space of  $D^{(k+1)}$  (see subsection 2.3).

The solution to Eq (5) will result in piecewise polynomials, specifically called discrete splines. For example,  $0^{\text{th}}$ -degree discrete splines are piecewise constant,  $1^{\text{st}}$ -degree curves are piecewise linear, and  $2^{\text{nd}}$ -degree curves are piecewise quadratic. For  $k \geq 1$ ,  $k^{\text{th}}$ -degree discrete splines are continuous and have continuous discrete differences up to degree  $k-1$  at the knots. This penalty results in more flexibility compared to the homogeneous smoothness that is created by the squared  $\ell_2$  norm.

Using different orders of divided differences result in estimated effective reproduction numbers with different smoothness properties.

For unevenly-spaced data, the spacing between neighbouring parameters varies with the time between observations, and thus, the divided differences must be adjusted by the times that the observations occur. Given observation times  $\mathbf{x} = (x_1, \dots, x_n)^\top$ , for  $k \geq 1$ , define a  $k^{\text{th}}$ -order diagonal matrix

$$X^{(k)} = \text{diag} \left( \frac{k}{x_{k+1} - x_1}, \frac{k}{x_{k+2} - x_2}, \dots, \frac{k}{x_n - x_{n-k}} \right). \quad (7)$$

Letting  $D^{(\mathbf{x},1)} := D^{(1)}$ , then for  $k \geq 1$ , the  $(k+1)^{\text{th}}$ -order divided difference matrix for unevenly spaced data can be created recursively by  $D^{(\mathbf{x},k+1)} := D^{(1)} X^{(k)} D^{(\mathbf{x},k)}$ . No adjustment is required for  $k = 0$ .

Due to the penalty structure, this estimator is locally adaptive, meaning that it can potentially capture local changes such as the initiation of control measures. [11, 12] considered only the 2<sup>nd</sup>-order divided difference of  $\mathcal{R}_t$  rather than its logarithm. In comparison to their work, our estimator (i) allows for arbitrary degrees of temporal smoothness and (ii) avoids the potential numerical issues of penalizing/estimating positive real values. Furthermore, as we will describe below, our procedure is computationally efficient for estimation over an entire sequence of penalty strengths  $\lambda$  and provides methods for choosing how smooth the final estimate should be.

## 2.3 Solving over a sequence of tuning parameters

We can solve the Poisson trend filtering estimator over an arbitrary sequence of  $\lambda$  that produces different levels of smoothness in the estimated curves. We consider a candidate set  $\boldsymbol{\lambda} = \{\lambda_m\}_{m=1}^M$  that is strictly decreasing.

Let  $D := D^{(k+1)}$  for simplicity in the remainder of this section. As  $\lambda \rightarrow \infty$ , the penalty term  $\lambda \|D\theta\|_1$  dominates the Poisson objective, so that minimizing the objective is asymptotically equivalent to minimizing the penalty term, which results in  $\|D\theta\|_1 = 0$ . In this case, the divided differences of  $\theta$  with order  $k+1$  is always 0, and thus,  $\theta$  must lie in the null space of  $D$ , that is,  $\theta \in \mathcal{N}(D)$ . The same happens for any  $\lambda$  beyond this threshold, so define  $\lambda_{\max}$  to be the smallest  $\lambda$  that produces  $\theta \in \mathcal{N}(D)$ . It turns out that this value can be written explicitly as  $\lambda_{\max} = \|(D^\dagger)^\top (\eta - y)\|_\infty$ , where  $D^\dagger$  is the

(left) generalized inverse of  $D$  satisfying  $D^\dagger D = I$ . Therefore, we use  $\lambda_1 = \lambda_{\max}$  and  
then choose the minimum  $\lambda_M$  to be  $r\lambda_{\max}$  for some  $r \in (0, 1)$  (typically  $r = 10^{-5}$ ).  
Given any  $M \geq 3$ , we generate a sequence of  $\lambda$  values to be equally spaced on the  
log-scale between  $\lambda_1$  and  $\lambda_M$ .

To compute the sequence efficiently, the model is estimated sequentially by visiting  
each component of  $\lambda$  in order. The estimates produced for a larger  $\lambda$  are used as the  
initial values (warm starts) for the next smaller  $\lambda$ . By solving through the entire  
sequence of tuning parameters, we have a better chance to achieve a better trade-off  
between bias and variance, and accordingly, improved accuracy relative to procedures  
examining one fixed value of  $\lambda$ .

## 2.4 Choosing a final $\lambda$

We estimate model accuracy over the candidate set through  $K$ -fold cross validation  
(CV) to choose the best tuning parameter. Specifically, we divide  $\mathbf{y}$  (except the first and  
last observations) roughly evenly and randomly into  $K$  folds, estimate  $\mathcal{R}_t$  for all  $\lambda$   
leaving one fold out, and then predict the held-out observations. Model accuracy can be  
measured by multiple metrics such as mean squared error  $\text{MSE}(\hat{y}, y) = n^{-1} \|\hat{y} - y\|_2^2$  or  
mean absolute error  $\text{MAE}(\hat{y}, y) = n^{-1} \|\hat{y} - y\|_1$ , but we prefer to use the (average)  
deviance, to mimic the likelihood in Eq (4):

$D(y, \hat{y}) = n^{-1} \sum_{i=1}^n 2(y_i \log(y_i) - y_i \log(\hat{y}_i) - y_i + \hat{y}_i)$ , with the convention that  
 $0 \log(0) = 0$ . Note that for any  $K$  and any  $M$ , we will end up estimating the model  
 $(K + 1)M$  times rather than once.

## 2.5 Approximate confidence bands

We also provide empirical confidence bands of the estimators with approximate  
coverage. Consider the related estimator  $\tilde{\mathcal{R}}_t$  defined as

$$\tilde{\mathcal{R}} = \exp(\tilde{\theta}) \quad \text{where} \quad \tilde{\theta} = \underset{\theta \in \mathbb{R}^n}{\operatorname{argmin}} \eta^\top \exp(\theta) - \mathbf{y}^\top \theta + \lambda \|D\theta\|_2^2. \quad (8)$$

Let  $\tilde{\mathbf{y}} = \eta \circ \tilde{\mathcal{R}}$ , and then it can be shown (for example, Theorem 2 in [28]) that an  
estimator for  $\text{Var}(\tilde{\mathbf{y}})$  is given by  $(\text{diag}(\tilde{\mathbf{y}}^{-2}) + \lambda D^\top D)^\dagger$ . Finally, an application of the  
delta method shows that  $\text{Var}(\tilde{\mathbf{y}}_t)/\eta_t^2$  is an estimator for  $\text{Var}(\tilde{\mathcal{R}}_t)$  for each  $t = 1, \dots, n$ .

We therefore use  $(\text{diag}(\hat{\mathbf{y}}^{-2}) + \lambda D^T D)_t^\dagger / \eta_t^2$  as an estimator for  $\text{Var}(\hat{\mathcal{R}}_t)$ . An  
approximate  $(1 - \alpha)\%$  confidence interval then can be written as  $\hat{\mathcal{R}}_t \pm s_t \times T_{\alpha/2, n-\text{df}}$ ,  
where  $s_t$  is the square-root of  $\text{Var}(\hat{\mathcal{R}}_t)$  for each  $t = 1, \dots, n$  and df is the number of  
change-points in  $\hat{\theta}$  plus  $k + 1$ .

## 2.6 Bayesian perspective

Unlike many other methods for  $\mathcal{R}_t$  estimation, our approach is frequentist rather than  
Bayesian. Nonetheless, it has a corresponding Bayesian interpretation: as a state-space  
model with Poisson observational noise, autoregressive transition equation of degree  
 $k \geq 0$ , e.g.,  $\theta_{t+1} = 2\theta_t - \theta_{t-1} + \varepsilon_{t+1}$  for  $k = 1$ , and Laplace transition noise  
 $\varepsilon_{t+1} \sim \text{Laplace}(0, 1/\lambda)$ . Compared to **EpiFilter** [8], we share the same observational  
assumptions, but our approach has a different transition noise. **EpiFilter** estimates  
the posterior distribution of  $\mathcal{R}_t$ , and thus it can provide credible interval estimates as  
well. Our approach produces the maximum *a posteriori* estimate via an efficient convex  
optimization, obviating the need for MCMC sampling. But the associated confidence  
bands are created differently.

## 3 Results

Implementation of our approach is provided in the R package **rtestim**. All experiments  
are run in R version 4.3.1 on a MacBook with an Apple M1 Pro chip and 32GB RAM  
running under macOS Sonoma 14.0. The R packages used for simulation and real-data  
application are **EpiEstim** 2.2-4, **EpiLPS** 1.2.0, and **rtestim** 0.0.4.

### 3.1 Synthetic experiments

We simulate four scenarios of the time-varying effective reproduction number, intended  
to mimic different epidemics. The first two scenarios are rapidly controlled by  
intervention, where the  $\mathcal{R}(t)$  consists of one discontinuity and two segments. Scenario 1  
has constant  $\mathcal{R}(t)$  before and after an intervention, while Scenario 2 grows exponentially,  
then decays. The other two scenarios are more complicated, where more waves are  
involved. Scenario 3 has four linear segments with three discontinuities, which reflect  
the effect of an intervention, resurgence to rapid transmission, and finally suppression of

the epidemic. Scenario 4 involves sinusoidal waves throughout the epidemic. The first three scenarios and the last scenario are motivated by [8] and [9] respectively. We name the four scenarios as (1) *piecewise constant*, (2) *piecewise exponential*, (3) *piecewise linear*, and (4) *periodic* lines or curves respectively.

In all cases, the times of observation are regular, and epidemics are of length  $n = 300$ . Specifically, in Scenario 1,  $\mathcal{R}_t = 2, 0.8$  before and after  $t = 70$ . In Scenario 2,  $\mathcal{R}_t$  increases and decreases exponentially with rates 0.015, 0.005 pre and post  $t = 50$ . In Scenario 3,  $\mathcal{R}_t$  is piecewise linear with four discontinuous segments following

$$\begin{aligned} \mathcal{R}(t) = & \left(2.5 - \frac{0.5}{59}(t-1)\right) \mathbf{1}_{[1,61)}(t) + \left(0.8 - \frac{0.2}{49}(t-61)\right) \mathbf{1}_{[61,111)}(t) \\ & + \left(1.7 + \frac{0.3}{39}(t-111)\right) \mathbf{1}_{[111,151)}(t) + \left(0.9 - \frac{0.4}{149}(t-151)\right) \mathbf{1}_{[151,300]}(t), \end{aligned} \quad (9)$$

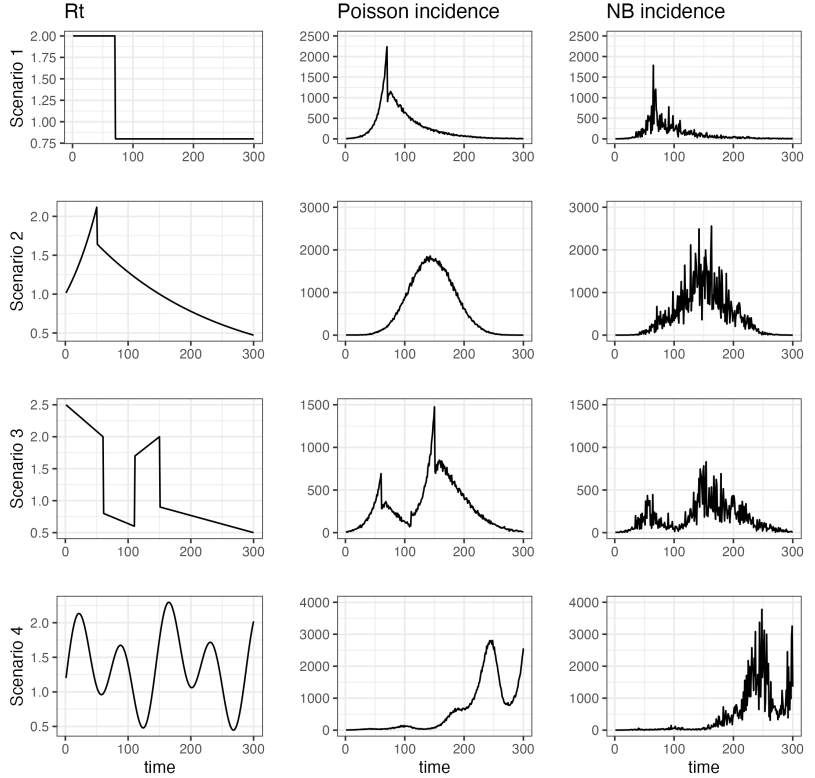
where  $\mathbf{1}_A(t) = 1$ , if  $t \in A$ , and  $\mathbf{1}_A(t) = 0$  otherwise. In Scenario 4,  $\mathcal{R}_t$  is realization of the continuous, periodic curve generated by the function

$$\mathcal{R}(t) = 0.2 \left( (\sin(\pi t/12) + 1) + (2 \sin(\pi t/6) + 2) + (3 \sin(\pi t/1.2) + 3) \right), \quad (10)$$

evaluated at equally spaced points  $t \in [0, 10]$ . These settings are illustrated in the left column of Fig 2.

We assume that the serial interval follows the Gamma distribution with fixed shapes and scales (3, 3), (2.5, 2.5), (3.5, 3.5) and (3.5, 3.5) for Scenarios 1–4 respectively. We initialize all epidemics with  $y_1 = 2$  cases and generating for  $t = 2, \dots, 300$ . We compute the expected incidence  $\Lambda_t$  using the renewal equation, and generate the incident infections from the Poisson distribution  $y_t \sim \text{Pois}(\Lambda_t)$ . To verify the performance of our model under the violation of this distributional assumption, we also generate incident infections using the negative Binomial distribution with dispersion size 5, i.e.,  $y_t \sim \text{NB}(\Lambda_t, \text{size} = 5)$ . For each setting, we generate 50 random samples, resulting in 400 total experiments. An example of each effective reproduction number scenario with a single corresponding Poisson and negative Binomial sample incidence sequences is displayed in Fig 2.

We compare RtEstim to EpiEstim and EpiLPS. Unfortunately, EpiFilter frequently fails to converge due to the large incident counts in these settings, so we are



**Fig 2.** The effective reproduction numbers (left column) and corresponding sample incident cases drawn from a Poisson (middle column) or negative Binomial (right column) distribution. The rows correspond to the four  $\mathcal{R}_t$  settings.

unable to include its results. **EpiEstim** estimates the posterior distribution of the effective reproduction number given a Gamma prior and Poisson distributed observations over a trailing window, under the assumption that the effective reproduction number is constant during that window. A larger window averages out more fluctuations, leading to smoother estimates, whereas, a shorter sliding window is more responsive to sudden spikes or declines. We tried the default, a weekly sliding window, as well as a monthly window. However, since neither considerably outperforms the other across all scenarios, we defer the monthly results to the supplementary document. **EpiLPS** is another Bayesian approach that estimates P-splines based on the Laplace approximation to the conditional posterior with negative Binomial likelihood. For **RtEstim** on the four scenarios respectively, we estimate (1) piecewise constant  $k = 0$ , (2) piecewise linear & cubic  $k = 1, 3$ , (3) piecewise linear  $k = 1$  and (4) piecewise cubic polynomials  $k = 3$ . In each case, we examine a grid of 50  $\lambda$  values, selecting the best using 10-fold cross validation. For all models and problems, we use the same serial

interval distribution for estimation that was used to create the data.

To measure estimation accuracy, we compare  $\widehat{\mathcal{R}}$  to  $\mathcal{R}$  using the Kullback-Leibler (KL) divergence. We use the KL divergence for the Poisson distribution (summed over across all  $t$ ) to measure the accuracy of the  $\mathcal{R}_t$  estimates

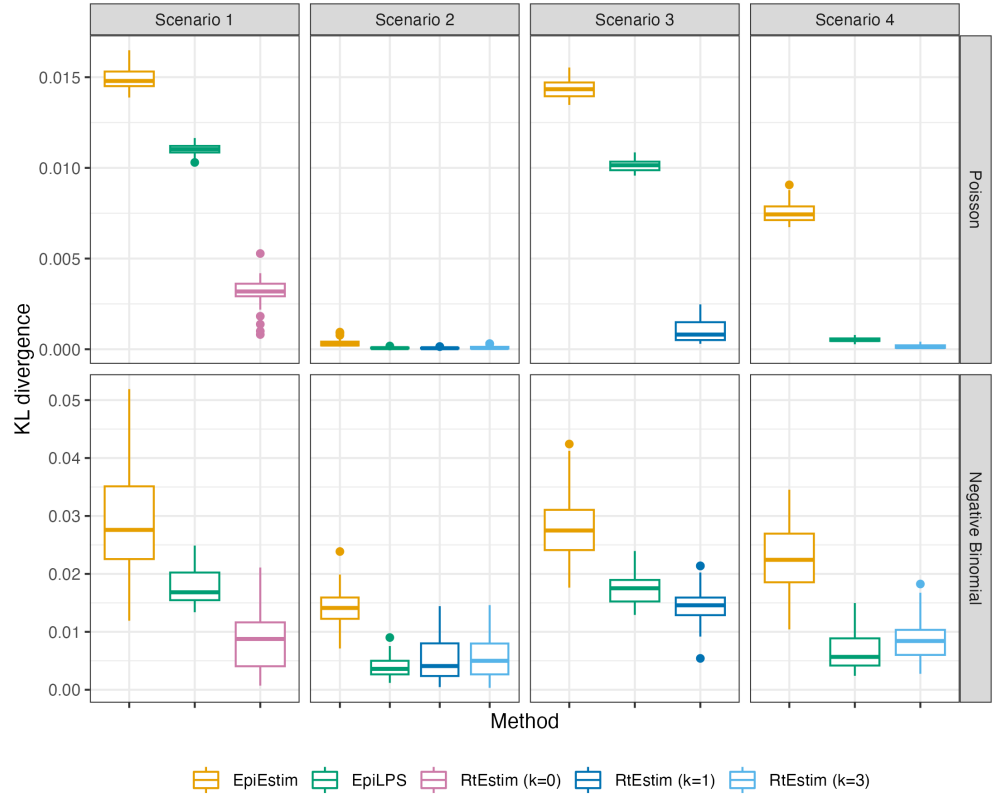
$$D_{KL}(\mathcal{R} \parallel \widehat{\mathcal{R}}) = \sum_{t=1}^n w_t \left( \mathcal{R}_t \log \left( \frac{\mathcal{R}_t}{\widehat{\mathcal{R}}_t} \right) + \widehat{\mathcal{R}}_t - \mathcal{R}_t \right), \quad (11)$$

where  $\mathcal{R} = \{\mathcal{R}_t\}_{t=1}^n$  and  $w_t = \eta_t / \sum_t \eta_t$  is the rescaled total infectiousness. To fairly compare across methods, we drop the estimates during the first week because estimates from `EpiEstim` do not begin until  $t = 8$  (using a weekly window). KL divergence is more appropriate for measuring accuracy because it connects directly to the Poisson likelihood used to generate the data, whereas standard measures like the mean-squared error correspond to Gaussian likelihood. Using Poisson likelihood has the effect of increasing the relative cost of mistakes when  $\Lambda_t$  is small. Other details of the experimental settings are deferred to the supplementary document.

### 3.2 Results for synthetic data

`RtEstim` overall outperforms `EpiEstim` and `EpiLPS` in the experimental study. [Fig 3](#) visualizes the KL divergence across the three models. Under both Poisson and negative Binomial distributions, `RtEstim` is easily the most accurate for Scenarios 1 and 3: the median of KL divergence is much lower and the boxes frequently fail to overlap indicating better performance than the other two methods across all 50 simulations. The advantage is less pronounced for the negative Binomial configuration, but still obvious. `RtEstim` and `EpiLPS` have similar performance in Scenarios 2 and 4. For the Poisson case, `RtEstim` and `EpiLPS` both have very small KL scores, which are very close to zero. In Scenario 4, `RtEstim` is slightly better for Poisson and `EpiLPS` is better for negative Binomial, but the boxes largely overlap each other. `EpiLPS` has a slightly lower median and a smaller IQR in Scenario 2 for the negative Binomial case. Both smoothness choices for `RtEstim` in Scenario 2 perform similarly across noise distributions, implying good performance under model misspecification. We will examine a single realization of each experiment to investigate these global conclusions in more detail.

[Fig 4](#) shows one realization for the estimated effective reproduction number under



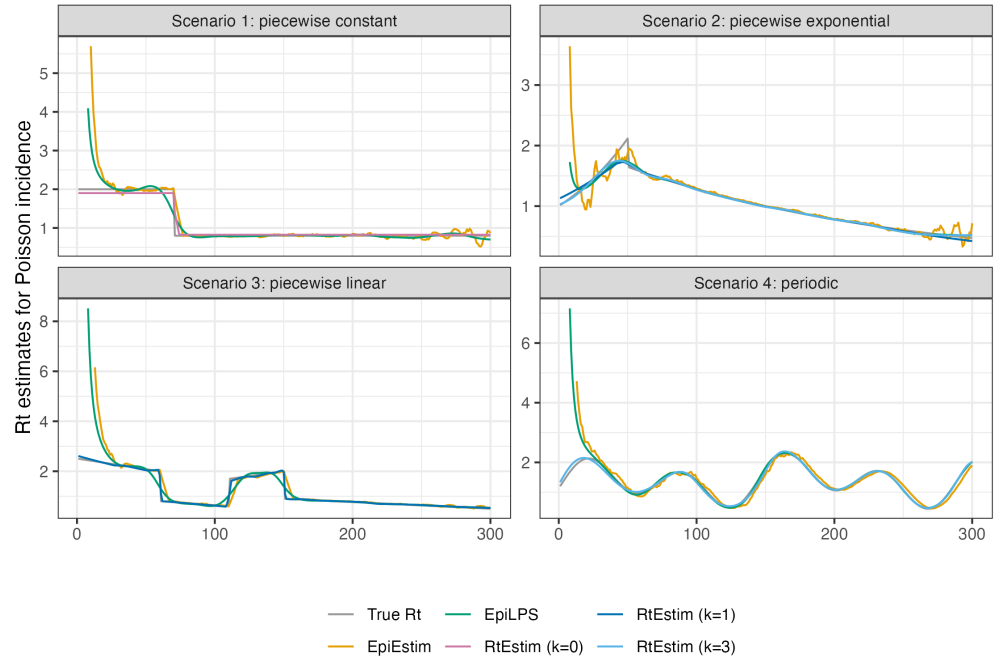
**Fig 3.** Boxplot of KL divergence between the estimated  $\hat{\mathcal{R}}_t$  and the true  $\mathcal{R}_t$  across 50 random samples for each approach given Poisson incidence (*in top panels*) and negative Binomial incidence (*in bottom panels*) respectively. Outliers are excluded.

the Poisson generative model for all four scenarios. Compared to EpiEstim and EpiLPS, which have rather severe difficulties at the beginning of the time series, RtEstim estimates are more accurate—they nearly overlap with the true values—without suffering from the initialization problem. Scenario 1 is the simplest case with only one knot and two constant segments. Besides the edge problem, EpiEstim and EpiLPS produce “smooth” estimated curves that are continuous at the changepoint, which results in large mistakes in that neighbourhood. Since the piecewise constant RtEstim estimator does not force any smoothness in  $\mathcal{R}_t$ , it easily captures the sharp change. Scenario 2 is relatively easy for all methods, except at the changepoint occurring at the end of the exponential growth. Although the truth is likely best represented with a discontinuous piecewise cubic curve, the actual curvature is so gentle that linear estimation ( $k = 1$ ) appears potentially reasonable. However, RtEstim has difficulty recovering the acute rise in the growth phase because it enforces continuity at the



changepoint.

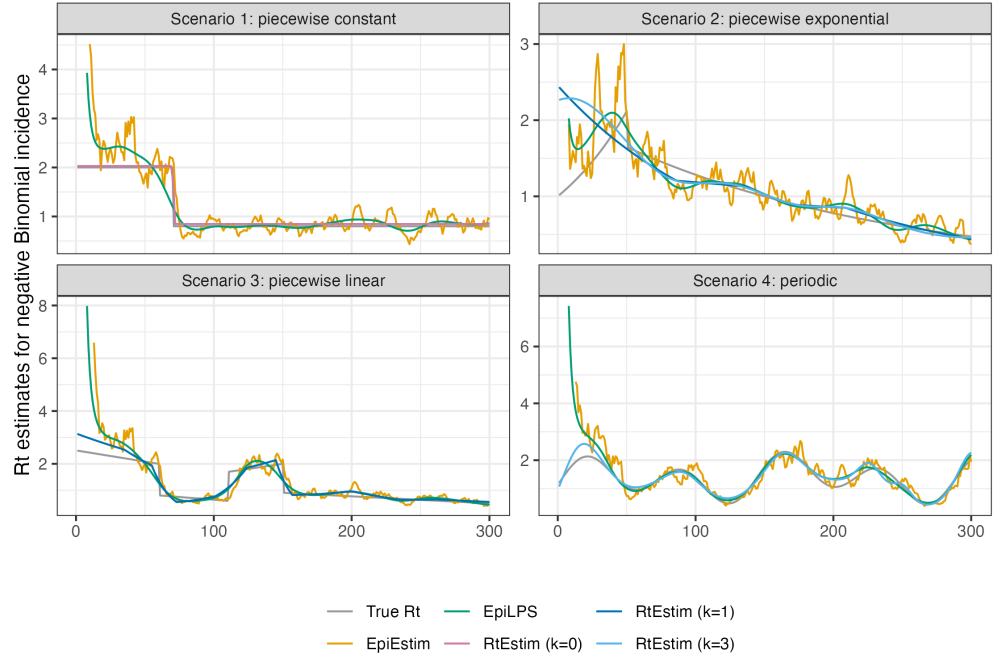
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**Fig 4.** Example of effective reproduction number estimation for Poisson observations.

To investigate the performance when the Poisson assumption (imposed by both **RtEstim** and **EpiEstim**) is violated, we also examine estimation accuracy with negative Binomial data. **Fig 5** displays a realization, analogous to the previous case, for all methods and scenarios. **RtEstim** has more difficulty relative to the Poisson setting, especially at the beginning of the outbreak. This is most pronounced in Scenario 4, where **RtEstim** is overly smooth, except in the last wave. In Scenario 2, **RtEstim** successfully captures the changepoint, but suffers from the same discontinuity problem as in the Poisson setting. In Scenario 3, the piecewise linear version of **RtEstim** recovers the curvature of  $\mathcal{R}_t$  well, but is less accurate than in the Poisson case.

Finally, it is important to provide a brief comparison of the running times of all three models across the 8 experimental settings. We find that almost all models across all experiments complete within 10 seconds. **RtEstim** generally takes the longest, due to a relatively large number of estimates—50 values of  $\lambda$  and 10 folds of cross validation require 550 estimates—while other models run only a single time for a fixed setting of hyperparameters per experiment. Additional results on timing comparisons are deferred



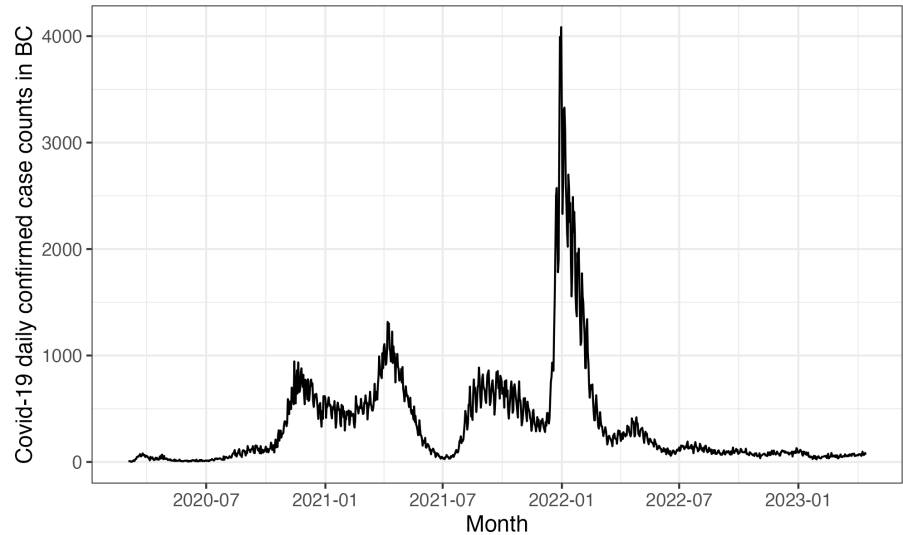
**Fig 5.** Example of effective reproduction number estimation for negative Binomial observations.

to the supplementary document.

### 3.3 Real-data results: Covid-19 incident cases in British Columbia

We implement `RtEstim` on Covid-19 confirmed incident cases in British Columbia (B.C.) as reported on May 18, 2023 (visualized in Fig 6) by the B.C. Centre for Disease Control [29]. We use the gamma distribution with shape 2.5 and scale 2.5 to approximate the serial interval function, which is similar to empirical estimates [30].

Considering the first, second, and third polynomial degrees,  $\hat{\mathcal{R}}_t$  for Covid-19 in British Columbia (illustrated in Fig 7) is always less than 3 except at the very early stage, which means that one distinct infected individuals on average infects less than three other individuals in the population. Examining three different settings for  $k$ , the temporal evolution of  $\hat{\mathcal{R}}$  (across all regularization levels  $\lambda$ ) are similar near the highest peak around the end of 2021 before dropping shortly thereafter. Throughout the estimated curves, the peaks and troughs of the effective reproduction numbers precede



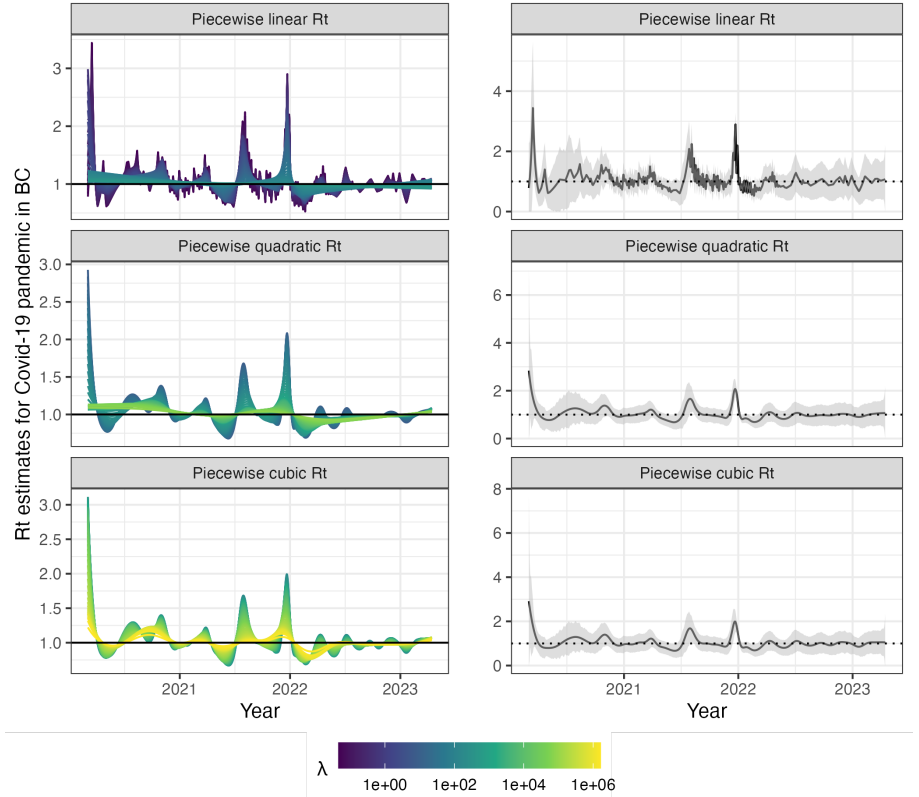
**Fig 6.** Covid-19 daily confirmed incident cases between March 1st, 2020 and April 15th, 2023 in British Columbia, Canada.

the growth and decay cycles of confirmed cases, as expected. We also visualize 95% confidence bands for the point estimates with  $\lambda$  chosen by minimizing cross-validated KL divergence in Fig 7.

The estimated effective reproduction numbers are relatively unstable before April, 2022. The highest peak coincides with the emergence and global spread of the Omicron variant. The estimated effective reproduction numbers fall below 1 during two time periods—roughly from April, 2021 to July, 2021 and from January, 2022 to April, 2022. The first trough coincides with the introduction of Covid-19 vaccines in British Columbia. The second trough, shortly after the largest peak may be due to variety of factors resulting in the depletion of the susceptible population such as increased self-isolation in response to media coverage of the peak or immunity incurred via recent infection. Since April, 2022, the estimated effective reproduction number has remained relatively stable (fluctuating around one) corresponding to low reported cases, though reporting behaviours also changed significantly since the Omicron wave.

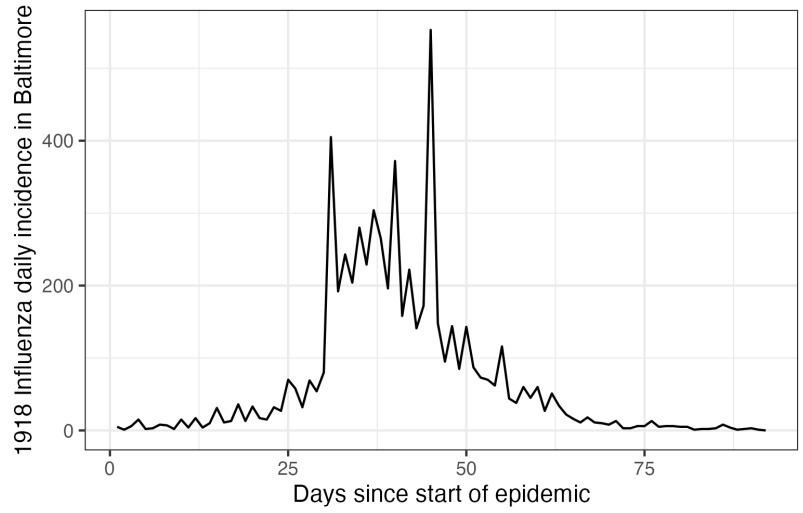
### 3.4 Real-data results: influenza in Baltimore, Maryland, 1918

We also apply `RtEstim` to daily reported influenza cases in Baltimore, Maryland occurring during the world-wide pandemic of 1918 from September to November [31]. The data, shown in Fig 8, is included in the `EpiEstim` R package. The 1918 influenza

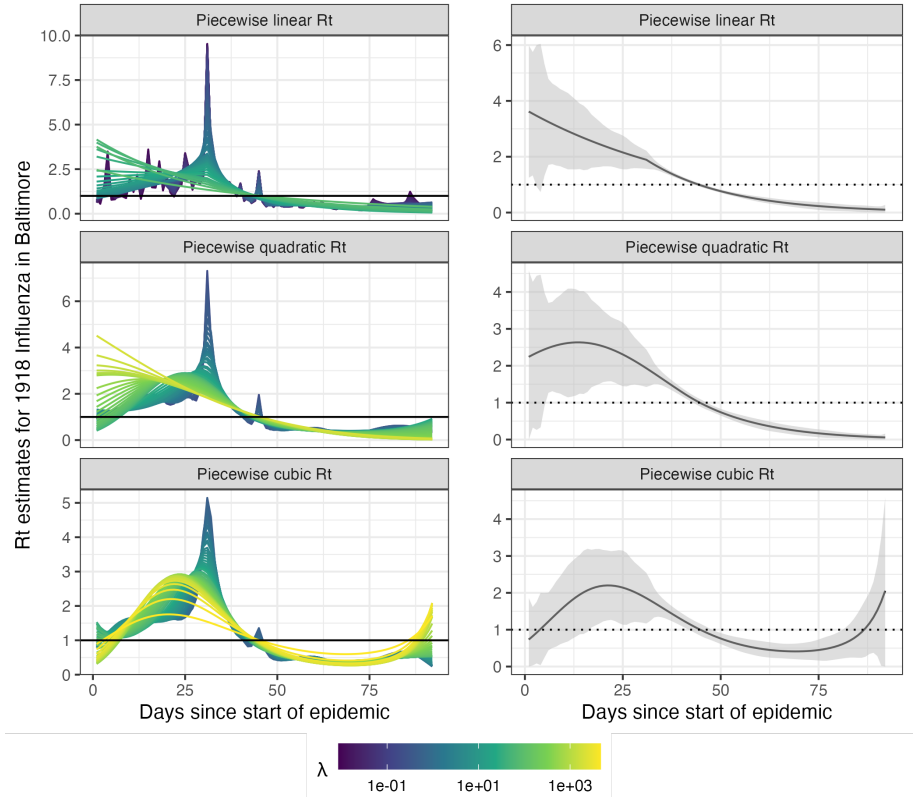


**Fig 7.** Estimated effective reproduction number based on Covid-19 daily confirmed incident cases between March 1st, 2020 and April 15th, 2023 in British Columbia, Canada. The left panels show estimates corresponding to 50 tuning parameters. The right panels show the CV-tuned estimate along with approximate 95% confidence bands. The top, middle and bottom panels show the estimated  $\mathcal{R}_t$  using the Poisson trend filtering in Eq (5) with degrees  $k = 1, 2, 3$  respectively.

outbreak, caused by the H1N1 influenza A virus, was unprecedentedly deadly with case 409  
fatality rate over 2.5%, infecting almost one-third of the population across the 410  
world [32]. The CV-tuned piecewise cubic estimates in Fig 9 better capture the growth 411  
at the beginning of the pandemic in Fig 8. The estimated  $\mathcal{R}_t$  curve suggests that the 412  
transmissibility of the pandemic grew rapidly over the first 30 days before declining 413  
below one after 50 days. However, it also suggests an increase in infectiousness toward 414  
the end of the period. With this data, it is difficult to determine if there is a second 415  
wave or a steady decline ahead. The CV-tuned piecewise constant and linear estimates 416  
in Fig 9 both suggest a steady decline. This conclusion is supported by the fact that 417  
incident cases decline to zero at the end of the period and matches  $\mathcal{R}$  estimates in [2], 418  
which are all lower than one. 419



**Fig 8.** Daily incident influenza cases in Baltimore, Maryland between September and November in 1918.



**Fig 9.** Estimated effective reproduction numbers for influenza in Baltimore, Maryland in 1918. The left panels show estimates for a set of 50 tuning parameters. The right column displays the CV-tuned estimates with approximate 95% confidence bands. The rows (top to bottom) show estimated effective reproduction numbers ( $\mathcal{R}_t$ ) using the Poisson trend filtering in Eq (5) with  $k = 1, 2, 3$  respectively.

## 4 Discussion

The **RtEstim** methodology provides a locally adaptive estimator using Poisson trend filtering on univariate data. It captures the heterogeneous smoothness of effective reproduction numbers given observed incidence data rather than resulting in global smoothness. This is a nonparametric regression model which can be written as a convex optimization (minimization) problem. Minimizing the distance (KL divergence across all coordinates) between the estimators and (functions of) observations guarantees data fidelity while the penalty on divided differences between pairs of neighbouring parameters imposes smoothness. The  $\ell_1$ -regularization results in sparsity of the divided differences, which leads to heterogeneous smoothness across time.

The property of local adaptivity (heterogeneous smoothness) is useful to automatically distinguish, for example, seasonal outbreaks from outbreaks driven by other factors (behavioural changes, foreign introduction, etc.). Given a well-chosen polynomial degree, the growth rates can be quickly detected, potentially advising public health to implement policy changes. The effective reproduction numbers can be estimated retrospectively to examine the efficacy of such policies, whether they result in  $\mathcal{R}_t$  falling below 1 or the speed of their effects. The smoothness of  $\mathcal{R}_t$  curves (including the polynomial degrees and tuning parameters) should be chosen based on the purpose of the study in practice, e.g., epidemic forecasting may require less smoothness while retrospective studies that solely target understanding of the pandemic may prefer a smoother estimate.

Our method **RtEstim** provides a natural way to deal with missing data, for example, on weekends and holidays or due to changes in reporting frequency. While solving the convex optimization problem, our method can easily handle uneven spacing or irregular reporting. Computing the total primary infectiousness is also easily generalized to irregular reporting by modifying the discretization of the serial interval distribution. Additionally, because the  $\ell_1$  penalty introduces sparsity (operating like a median rather than a mean), this procedure is relatively insensitive to outliers compared to  $\ell_2$  regularization.

There are a number of limitations that may influence the quality of  $\mathcal{R}_t$  estimation. While our model is generic for incidence data rather than tailored to any specific

disease, it does assume that the generation interval is short relative to the period of data collection. More specialized methodologies would be required for diseases with long incubation periods such as HIV or Hepatitis. Our approach, does not explicitly model imported cases, nor distinguish between subpopulations that may have different mixing behaviour. While the Poisson assumption is common, it does not handle overdispersion (observation variance larger than the mean). The negative binomial distribution is a good alternative, but more difficult to estimate in this context. As described in [section 1](#), the expression for  $\mathcal{R}$  assumes that a relatively constant proportion of true infections is reported. However, if this proportion varies with time (say, due to changes in surveillance practices or testing recommendations), the estimates may be biased over this window. A good example is in early January 2022, during the height of the Omicron wave, British Columbia moved from testing all symptomatic individuals to testing only those in at-risk groups. The result was a sudden change that would render  $\mathcal{R}_t$  estimates on either side of this timepoint incommensurable.

As currently implemented, `RtEstim` uses a fixed serial interval throughout the period of study, but as factors such as population immunity vary, the serial interval may vary as well [4]. Another issue relates to equating serial and generation intervals (also mentioned above). The serial interval distribution is generally wider than that of the generation interval, because the serial interval involves the convolution of two distributions, and is unlikely to actually follow a named distribution like Gamma, though it may be reasonably well approximated by one. Our implementation allows for an arbitrary distribution to be used, but requires the user to specify the discretization explicitly, requiring more nuanced knowledge than is typically available. Pushing this analysis further, to accommodate other types of incidence data (hospitalizations or deaths), a modified generation interval distribution would be necessary, and further assumptions would be required as well. Or else, one would first need to deconvolve deaths to infection onset before using our software.

Nonetheless, our methodology is implemented in a lightweight R package `rtestim` and computed efficiently, especially for large-scale data, with a proximal Newton solver coded in C++. Given available incident case data, prespecified serial interval distribution, and a choice of degree  $k$ , `RtEstim` is able to produce accurate estimates of effective reproduction number and provide efficient tuning parameter selection via cross

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# Supplementary details on experiments of effective reproduction number estimation with trend filtering

Jiaping Liu, Zhenglun Cai, Paul Gustafson, and Daniel J. McDonald

## 1 Supplementary details on experimental settings of RtEstim for Section 3.1

We run 10-fold cross validation (CV) to choose the best tuning parameter from the candidate set of size 50, i.e.,  $\lambda = \{\lambda_1, \dots, \lambda_{50}\}$ . Specifically, we divide all samples (except the first and last entries) into ten folds evenly and randomly, and build models on each sample set by leaving a fold out across all hyperparameters. We select the tuning parameter that gives the lowest averaged *deviance* between the estimated incidence and the observed samples averaged over all folds.

## 2 Experimental comparisons between EpiEstim with monthly sliding windows and other methods for Section 3.1

Fig 10 displays the KL divergence values for negative Binomial incidence. Comparing across  $\mathcal{R}_t$  estimates by EpiLPS, RtEstim and EpiEstim with *monthly* sliding windows, KL divergence computation excludes the first month of  $\mathcal{R}_t$  estimates for all approaches, since EpiEstim estimates with the monthly sliding windows are not available until the second month. The  $y$ -axis is displayed on a logarithmic scale for a better visualization, since a few values are much larger than others.

The relative performance of EpiEstim with monthly sliding windows, in general, is not as good as its weekly sliding window based on the relative positions of its boxes and the counterparts of the other methods, except for the Scenario 2 with negative Binomial incidence. It can be explained by EpiEstim with longer sliding windows assume similarity of neighbouring  $\mathcal{R}_t$  across longer periods, and thus, is smoother and less accurate compared to the one with shorter sliding windows.

## 3 Supplementary results of experiments for Fig 3 in Section 3.2

The full KL divergence values for Poisson incidence without excluding the outliers are shown in Fig 11. The  $y$ -axis is displayed on a logarithmic scale for the same reason in Fig 10.

Comparing among  $\mathcal{R}_t$  estimations by EpiLPS, RtEstim and EpiEstim with *weekly* sliding windows, KL divergence computation excludes the first week of  $\mathcal{R}_t$  estimates for all approaches. The outliers are mainly EpiLPS and EpiEstim  $\mathcal{R}_t$  estimations in Scenario 2, which is generally an easy problem except for the

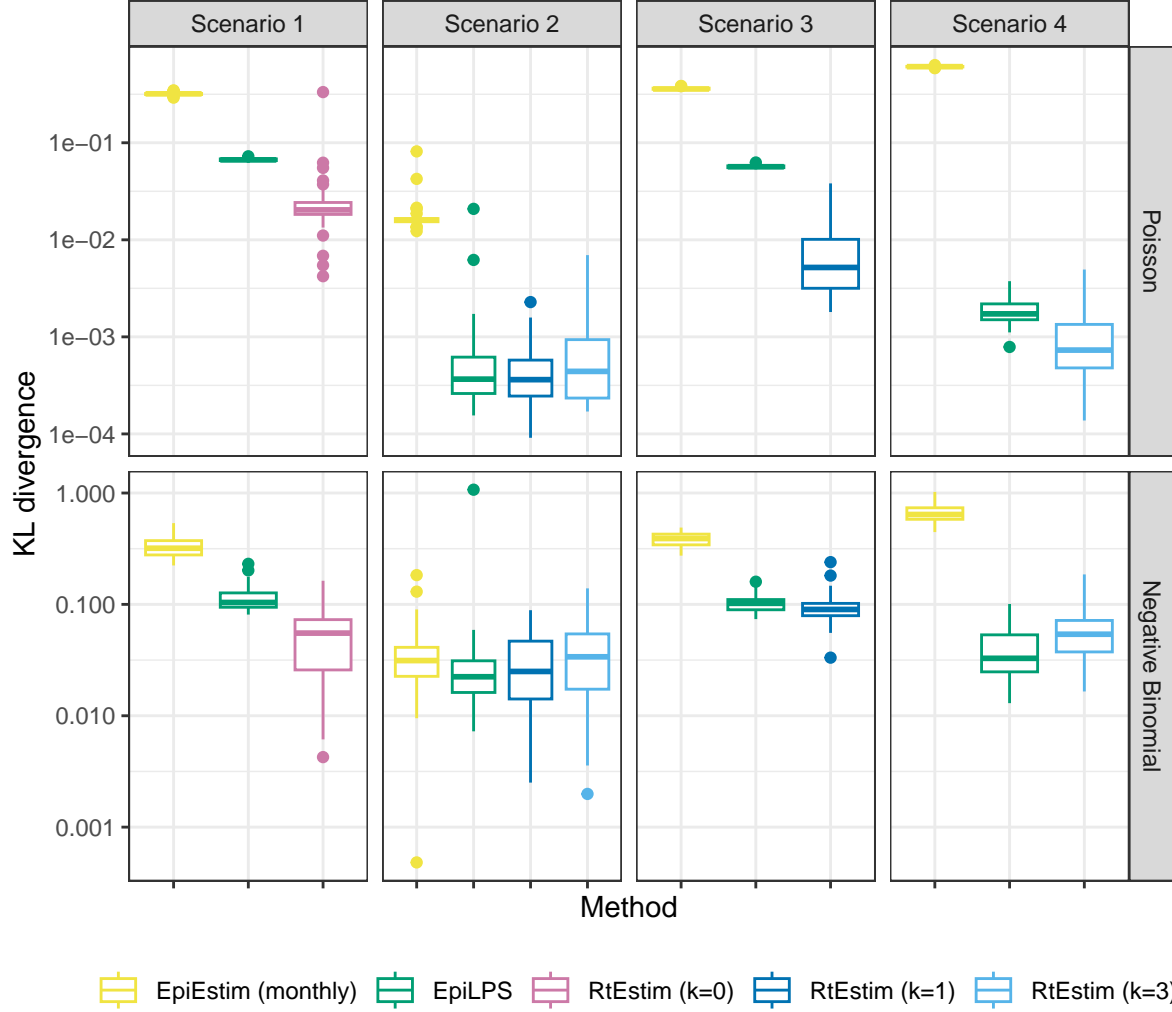


Fig 10. KL divergence for EpiEstim with monthly sliding windows and other methods. Y-axis is on a logarithmic scale.

change point at where the two segments are discontinuous. Since EpiEstim produces a continuous  $\mathcal{R}_t$  curve, the change point can be difficult to recover and make estimation less accurate.

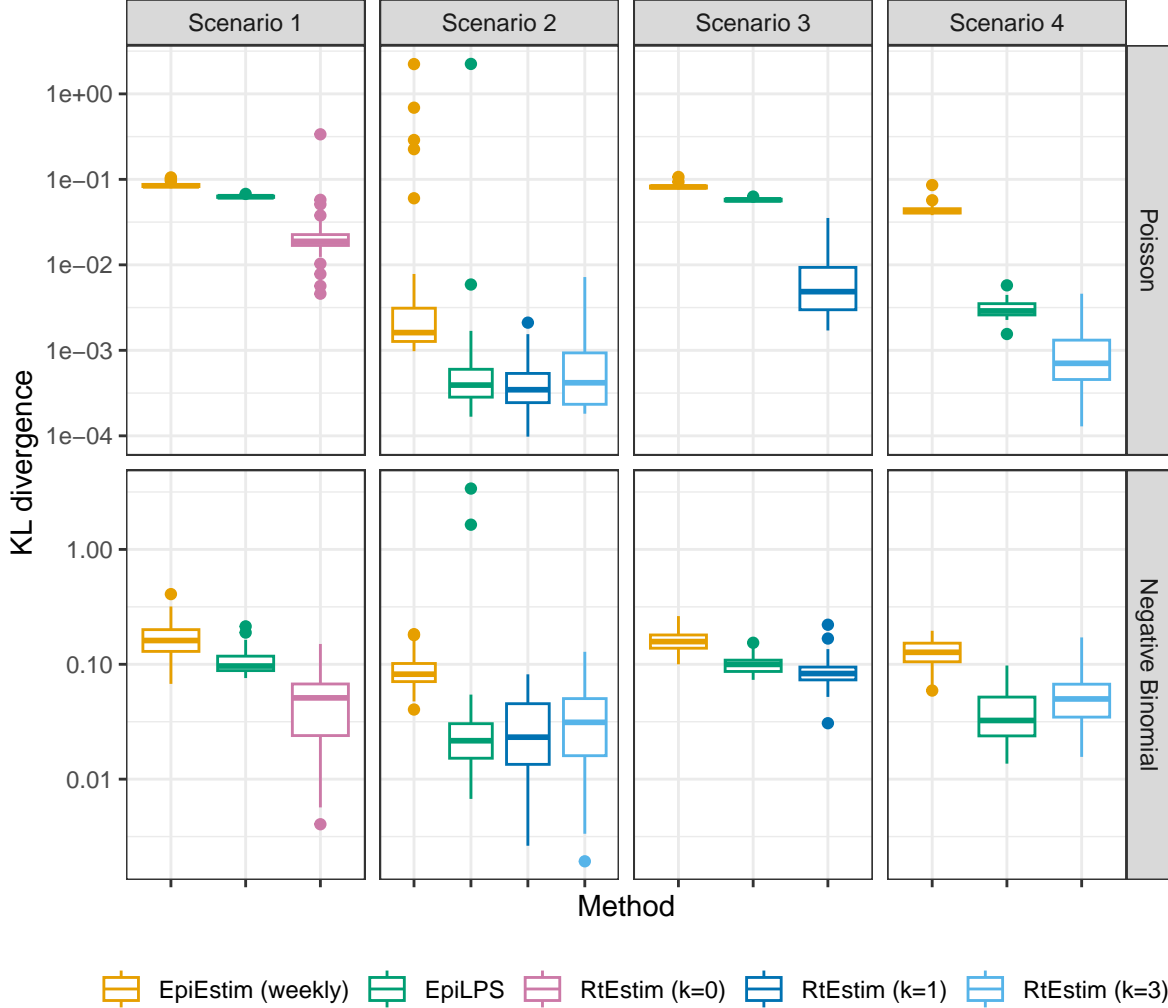


Fig 11. KL divergence for methods with weekly sliding windows. Y-axis is on a logarithmic scale.

## 4 Time comparisons of methods for Section 3.2

Fig 12 shows the time comparisons across all methods. EpiEstim with both weekly and monthly sliding windows are very fast and converge in less than 0.1 seconds. Piecewise constant RtEstim (with  $k=0$ ) estimates can be generated within 0.1 seconds as well. EpiLPS is slightly slower, but still very fast and within 1 second for all experiments. Piecewise linear and cubic RtEstim (with  $k=1$  and  $k=3$  respectively) are slower, but mostly within 10 seconds.

It is remarkable that our RtEstim computes 50 lambda values with 10-fold CV for each experiment, which results in 550 times of modelling per experiment (including modelling for all folds). The running times are no more than 10 seconds for most of the experiments, which means the running time for each time of modelling is very fast, and on average can be less than 0.02 seconds. The other two methods only run once for a fixed set of hyperparameters for each experiment.

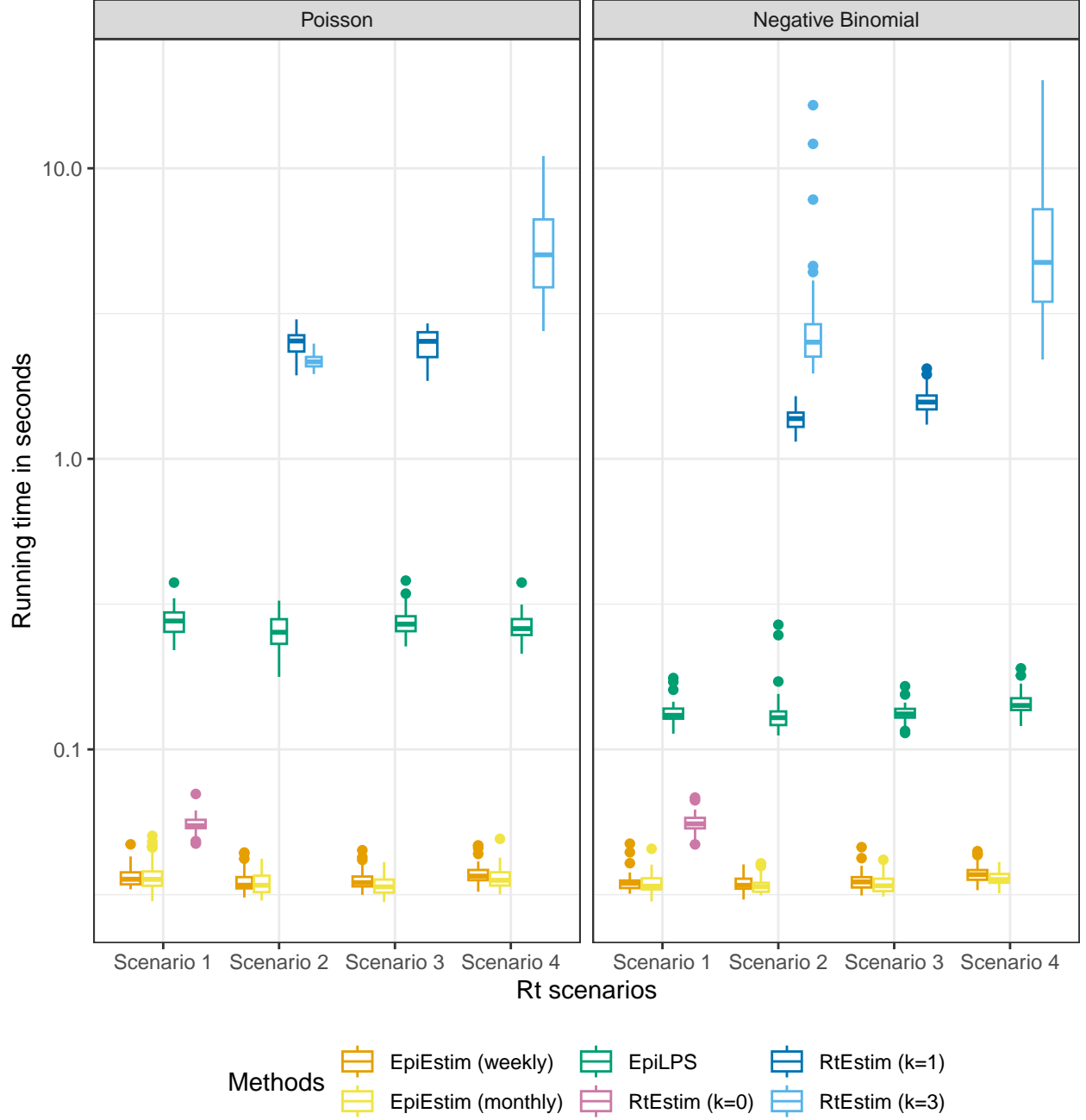


Fig 12. Time comparisons of methods (excluding one outlier of *RtEstim* ( $k=1$ ) in Scenario 2 with negative Binomial incidence). Y-axis is on a logarithmic scale.