Original Article

Bayesian estimation of real-time epidemic growth rates using Gaussian processes: local dynamics of SARS-CoV-2 in England

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Abstract

Quantitative assessments of the recent state of an epidemic and short-term projections for the near future are key public-health tools that have substantial policy impacts, helping to determine if existing control measures are sufficient or need to be strengthened. Key to these quantitative assessments is the ability to rapidly and robustly measure the speed with which an epidemic is growing or decaying. Frequently, epidemiological trends are addressed in terms of the (time-varying) reproductive number *R*. Here, we take a more parsimonious approach and calculate the exponential growth rate, *r*, using a Bayesian hierarchical model to fit a Gaussian process to the epidemiological data. We show how the method can be employed when only case data from positive tests are available, and the improvement gained by including the total number of tests as a measure of the heterogeneous testing effort. Although the methods are generic, we apply them to SARS-CoV-2 cases and testing in England, making use of the available high-resolution spatial heterogeneity.

Keywords: Bayesian hierarchical modelling, epidemiological trends, Gaussian processes, growth rate estimation, public-health tools, spatial heterogeneity

1 Introduction

Statistical analysis of the SARS-CoV-2 pandemic has been instrumental in both assessing the current status of infection at a local or national level (Davies et al., 2021, 2020; Flaxman et al., 2020; Hellewell et al., 2020; The Royal Society, 2020), and extrapolating to generate short-term projections. Arguably good statistical knowledge is a key to the control of epidemics, as it provides a quantitative assessment of control measures and can highlight sectors of the population in which additional targeted controls may be needed. Five elements combine to make the statistical analysis of the SARS-CoV-2 pandemic difficult: many infections are asymptomatic and go undetected; the regular use of lateral flow devices, which would detect asymptomatic infection, is heterogeneous across time, space, and age groups; the use of polymerase chain reaction (PCR) testing (adopted as the gold standard in the UK) also changes across time and space, presumably as individuals react to changes in perceived risk; infection and testing are inherently stochastic processes; and there are distributed lags between infection and detection. These five factors mean that the prompt

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identification of rising infection (especially in relatively small populations) requires sophisticated statistical methods.

The reproductive number, R, has gained substantial media and political interest during the SARS-CoV-2 pandemic as a simple statistical indicator of the current epidemiological trends, with R < 1 corresponding to a declining outbreak and R > 1 corresponding to a growing outbreak (Vegvari et al., 2021). During 2020 in England, the nationwide estimate of R (UK Health Security Agency, 2020a) was one of the key metrics in determining the national alert level with implications for changes in control measures (UK Health Security Agency, 2020b); hence placing great political, economic, and public-health importance on this single value. A robust and rapid estimation of R (or the epidemic growth rate r), together with levels of uncertainty, remains a key public-health tool. The estimation needs to be rapid, such that prompt action can be taken before the burden on health services becomes too great; the estimation also needs to be robust, as the economic and social consequences of action can be costly and so should only be enacted when there is considerable certainty that such measures are needed. As such there is a clear need for continued development of statistical methods that can extract a meaningful signal from complex and noisy epidemiological data.

Obtaining an accurate and timely measure of R generally requires a robust estimate of either the generation time or the infectiousness profile over time (Wallinga & Lipsitch, 2007) (capturing the expected level of transmission at time t after infection). Both of these necessitate detailed individual-level observations (Abbott et al., 2021, 2020; Hart et al., 2022) and may therefore be context dependent, leading to a diversity of R estimates from the same population-level data (Funk et al., 2020). Here, we adopt the more parsimonious approach of working with the growth rate r (such that the number of infections grows like $I(t) \sim exp(rt)$), in which case our threshold for a growing or declining outbreak becomes where r is greater or less than zero, respectively.

Given the importance of real-time estimation of the growth rate, r, or the reproductive number, R, multiple statistical methods have been developed (Gostic et al., 2020; The Royal Society, 2020). All methods have advantages and potential problems, with an inevitable trade-off between robustness and timeliness (Favero et al., 2022; Parag et al., 2021). Most naively, the growth rate can be estimated by simply measuring the rate of change of log (infection), where infections are often approximated as being proportional to reported cases. This naive approach is confounded by the stochastic nature of transmission and reporting, requiring either smoothing of the data or fitting the growth rate over a defined time window—longer windows and more smoothing eliminate stochastic effects, but mean that real-time estimates of the growth rate and R are considerably lagged. The UK government dashboard (UK Health Security Agency, 2020c) expands on these simple ideas to produce estimates of the growth rate at the national scale, calculated as the relative change over seven days in the smoothed number of cases (smoothed by taking a mean over a seven-day window). In recent years, EpiEstim (Cori et al., 2013) has grown in popularity as a method of estimating changing Rvalues, due to its flexibility and accuracy (Funk et al., 2020). EpiEstim uses a Bayesian framework to compare the reported number of cases over a time window with the projection based on the infectiousness profile and historic reporting to generate an estimate of R in a given window.

In this paper, we develop a novel method to generate a real-time estimate of the growth rate of infection in small stochastic populations. Our flexible method uses a Bayesian approach to compute the posterior distribution of the growth rate at any point in time and produces samples of the joint posterior distribution of the growth rate for any given interval. We use Gaussian processes (GPs) to fit the reported data, which gives us flexibility in smoothing the count of new cases according to the GP parameters. We fit two different measures: the raw number of recorded cases in a region, as defined by PCR positives in the community; or the proportion of community PCR tests that are positive. We show that using both measures copes well with potential biases from time-changing patterns in testing, as the latter provides a more stable estimate when testing patterns are changing rapidly.

We first outline the basic methodology and illustrate its use on surrogate data sets where the growth rate is known. We then apply our model to data on SARS-CoV-2 cases in England, initially at a national level by estimating the daily growth rate of SARS-CoV-2 from 1 September 2020 to 6 December 2021 (as available at the time of writing). Finally, we explore the spatial heterogeneity in cases at the lower tier local authority (LTLA) level in April 2021 when the Delta variant was increasing in the North–West of England—a time when the spatial variability of epidemic behaviour was key to understanding the impact of the new variant.

2 Methods

We describe a model framework to estimate the growth rate, r, of an epidemic based on the count of reported infections (cases). If the counts are recorded through a testing programme, to adjust for changes in testing effort the model can also incorporate the total number of tests performed over time. We assume that the underlying process generating the count of cases is given by a onedimensional Gaussian process (Section 2.1), and obtain the growth rate by sampling from the derivative of the process (Section 2.2).

2.1 Model structure

For a given community, let $T = \{1, ..., T\}$ be a set of time indices at which data are collected. For each $t \in T$, y_t denotes the number of positive test results at time t and n_t denotes the total number of tests. In the context of SARS-CoV-2, data are generally collated daily with the potential for missing data, which our proposed models allow for; for other infections data may be collected over larger or irregularly spaced intervals.

2.1.1 Positives model

We first propose the following Bayesian hierarchical model, labelled as the *positives model*, which only requires knowledge of y_t (the number of positive cases at time t) and is therefore applicable in situations where n_t (the number of tests at time t) is not available. The model assumes that y_t follows a negative binomial distribution parameterised by its mean μ_t and a time-homogeneous overdispersion parameter η . The probability mass function of y_t under this parameterisation is

$$\operatorname{Prob}(y_t|\mu_t,\eta) = \frac{\Gamma(y_t+\eta)}{\Gamma(\eta)\Gamma(y_t+1)} \frac{(\mu_t/\eta)^{y_t}}{(1+(\mu_t/\eta))^{\eta+y_t}}$$

The parameter log (η) is assigned a normal prior $\mathcal{N}(m_{\eta}, \tau_{\eta}^{-1})$. The log relative risk, log (μ_t) , is decomposed into the sum of a smooth term x_t and an optional error term ϵ_t whose distribution and dependencies are problem-dependent. The model can therefore be expressed as

$$y_t | \mu_t, \eta \sim \text{Negative binomial}(\mu_t, \eta) \log(\eta) \sim \mathcal{N}(m_\eta, \tau_\eta^{-1})$$
(1)

$$\log(\mu_t) = x_t + \epsilon_t$$

where the hyperparameters underpinning the distribution of the overdispersion (m_{η} and τ_{η}^{-1}) are specific to the problem and quoted in the results. Choices for the distribution of the error term ϵ_t are discussed at the end of the section.

The prior on the smooth terms x_t is given by a Gaussian process f on \mathbb{R} such that $x_t = f(t)$, where f has mean zero, covariance $k_{\theta}(f(s), f(s'))$ between the value of the process f at times s and s', and hyperparameters θ

$$\begin{aligned} x_t &= f(t) \\ f(s)|\theta \sim \mathcal{GP}(0, \, k_\theta(f(s), \, f(s'))) \end{aligned} \tag{2}$$

A comprehensive summary of such regression models using Gaussian processes can be found in (Rasmussen & Williams, 2006, Ch. 2). Here, we use a one-dimensional Matérn covariance family (Stein, 1999), since the resulting process f is stationary and isotropic, and the smoothness can be specified through a single smoothing parameter v. We choose v = 3/2 which results in the covariance function

$$k_{(l,\sigma)}(f(s), f(s')) = \sigma^2 \left(1 + \sqrt{3} \frac{|s-s'|}{l}\right) \exp\left(-\sqrt{3} \frac{|s-s'|}{l}\right)$$

which also depends on the additional hyperparameters $\theta = (l, \sigma)$; where *l* is the length-scale, and σ^2 is the marginal variance of the process (other covariance functions are explored in Appendix E).

We set the joint prior of l and σ as $(\log (l), \log (\sigma)) \sim \mathcal{N}((\log (l_0), \log (\sigma_0)), B^{-1})$ where l_0 and σ_0 are baseline values for the length scale and precision, respectively, and B is the precision matrix of the joint prior. A diagram of the model is shown in Appendix A.

The error term ϵ_t could be omitted in this formulation since variation in the model is included through the use of the negative binomial distribution. However, its inclusion can be used to capture other elements of the dynamics that are not otherwise present in the model. For example, reported SARS-CoV-2 cases in England have a pronounced day-of-the-week effect, with fewer cases reported on weekends (see Figure 2, Panel a). The day-of-the-week effect can be included in the model by allowing $\epsilon_t = w_{d(t)}$, where d(t) is the day of the week associated with time t, thus capturing the weekly pattern of testing and reporting. This is the approach we adopt for the remainder of the paper, assuming that the $w_{d(t)}$ are a priori independent identically distributed with Gaussian hyperprior with zero mean and precision $\tau_w \sim \Gamma(a_w, b_w)$. To avoid identifiability issues with the terms x_t , it is important to impose a sum-to-zero constraint $\sum_d w_d = 0$. For other applications, the error term could be used to capture other temporal factors such as seasonal effects or known changes in reporting or test availability.

2.1.2 Proportions model

If the number of tests is known, then an alternative model formulation is possible that accounts for changes in testing behaviour over time; we label this model the *proportions model* and seek to capture the proportion of tests that are positive. In this case, y_t (the number of positive cases at time *t*) given n_t (the number of tests at time *t*) is assumed to follow a beta-binomial distribution with mean parameter μ_t , overdispersion parameter ρ , and number of trials n_t . We use a beta-binomial distribution to account for both the bounded nature of y_t (which is bounded above by n_t) and the overdispersion.

The probability mass function of y_t under this parameterisation is given by

$$\operatorname{Prob}(y_t|\mu_t, M, n_t) = \binom{n_t}{y_t} \frac{\Gamma(M)}{\Gamma(M\mu_t)\Gamma(M(1-\mu_t))} \frac{\Gamma(y_t + M\mu_t)\Gamma(n_t - y_t + M(1-\mu_t))}{\Gamma(n_t + M)}$$

where $M = (1/\rho) - 1$. Given the bounded nature of the positive tests, such that $\mu_t \in (0, 1)$, we utilise the inverse logit transform $(\log it^{-1})$, and assume that $\log it^{-1}(\mu_t)$ is decomposed into the sum of a smooth term x_t and an optional error term ϵ_t whose distribution and dependencies are problem-dependent. The transformed overdispersion parameter $\log it^{-1}(\rho)$ is assigned a normal prior $\mathcal{N}(m_\rho, \tau_\rho^{-1})$. As in the positives model, the prior on x_t is given by the Gaussian process described in Section 2.1.1

$$y_t | \mu_t, \rho, n_t \sim \text{Beta-binomial}(\mu_t, \rho, n_t)$$

$$\log i t^{-1}(\rho) \sim \mathcal{N}(m_\rho, \tau_\rho^{-1}) \tag{3}$$

$$\log i t^{-1}(\mu_t) = x_t + \epsilon_t$$

2.2 Growth rate sampling

The instantaneous growth rate is defined as the per capita change in the number of new cases per time period. In other words, if w_t is the process generating new cases at time t, the growth rate corresponds to $r_t = \partial_t(w_t)/w_t$, or equivalently, $r_t = \partial_t(\log w_t)$; where ∂_t signifies the time derivative. However, w_t is unknown in practice, so we instead approximate the growth rate using our fitted Gaussian process. For the positives model, we approximate r_t as the growth rate of the process fitting the number of newly reported cases, $\exp(f(t))$. In other words, $r_t \approx r_t^A = \partial_t (\log [\exp \{f(t)\}]) = \partial_t f(t)$ and therefore, r_t can be estimated as the derivative of the Gaussian process f. For the proportions model, we approximate r_t as $r_t^B = [\partial_t \{f(t)\}]/[1 + \exp \{f(t)\}]$, such that r_t^B corresponds to the growth rate of new reported cases minus the growth rate the new tests performed (see Appendix B).

To capture the inherent uncertainty in the process f, we sample from the derivative of the process f to obtain samples of the growth rate. Note that Gaussian processes with the Matérn covariance are mean-square differentiable if v > 1, which is satisfied by our choice of v = 3/2 (Stein, 1999). We obtain samples of the derivative by taking numerical approximations of the derivative (∂_t) of

samples drawn from the process *f*. In other words, for a given sample *g* of *f*, we approximate the derivative as $g'(s) \approx \frac{g(s+h)-g(s-h)}{h}$ with error $\mathcal{O}(h)$, where *h* is the window size of the approximation. Samples of the derivative can be directly sampled from the derivative of the Gaussian process, providing more accurate results but requiring more computational time (see Appendix F).

2.3 Implementation

We implement the model in R using the package INLA (Rue et al., 2009), where the posterior distribution of the parameters of the model is obtained using a Laplace approximation. The Gaussian process with Matérn kernel is computed as the solution of a stochastic partial differential equation (Lindgren et al., 2011), obtained by the finite element method (FEM). To fit the model using the FEM implementation in INLA, we create a one-dimensional mesh with equally-spaced nodes that represent time points. The nodes are located according to the frequency of reported counts; that is, if the data are reported daily, one node per day is located, even if there is missing data. To avoid boundary effects, the mesh domain is extended by at least the length of the studied period (extra nodes are added before the first observation and after the last observation) (Lindgren & Rue, 2015). The code is available in GitHub/juniper-consortium/growth-rate-estim.

The Laplace approximation is suitable if the target distribution is unimodal. To verify the correctness of this approximation for the positives and proportions models, we also implement both models using the Hamiltonian Monte Carlo algorithms in the software STAN and compare them to the results in Section 3.2, as explained in Appendix C. It also allows us to implement different covariance functions not available in INLA.

For the rest of the paper, we use weakly informative priors to the overdispersion parameters of the models, such that $\log(\eta) \sim \mathcal{N}(0, 1)$ and $\operatorname{logit}^{-1}(\rho) \sim \mathcal{N}(0, 0.5^{-1})$. More restrictive priors could reject values of the hyperparameters possibly explained by the data (Rasmussen & Williams, 2006, Ch. 5). Choices of τ_{ϵ} , l_0 , σ_0 , and B are case-specific and are detailed in the results. Sensitivity analyses of the Gaussian process hyperparameters are explored in detail in Appendix D.

2.4 Model validation

To validate the accuracy of the models, we generate synthetic epidemiological data from a single homogeneous population of size N = 1,000,000. We assume for the first 100 days there is an underlying growth rate of r = 0.03 per day. For the second 100 days, we assume that controls are enacted and the epidemic goes into decline with a rate of r = -0.02. More precisely, the number of infections y(t) on day t are sampled from a Poisson distribution with rate $r \sum_{i=3}^{6} y(t-i)$ for t > 6 and $\exp(rt)$ for $t \le 6$ (where r = 0.03 for $t \le 100$, r = -0.02 for t > 100, and y(0) = 100). This approach aims to simulate data with a known growth rate using a model different to the methodology proposed in Section 2.1 to test if the positives and proportions models are capable of recovering the truth growth rate under a quick behaviour change (at t = 100).

We compare two scenarios for the number of daily tests n(t). As our purpose is to test the model accuracy under a known growth rate, rather than discuss the effect of the test sampling, we make highly optimistic assumptions for the frequency of testing. In the first scenario, a random 10% of the population is tested daily, n(t) = 0.1N; in the second scenario, tests increase linearly from n(0) = 0.01N to n(200) = 0.1N.

We run both the positives model and proportions model for each scenario. We set $l_0 = 50$, $\sigma_0 = 1$, and impose B = 1 to have non-informative priors for the parameters of the Gaussian process (where 1 is the identity matrix). For the approximation of the derivative, we set a window of h = 3 days for all times except near the boundary, where we choose h = 1 for t = 1,200, and h = 2 for t = 2,199.

For simulation 1, with a constant daily testing rate, for both models the true growth rate is in the posterior credible interval (CI) for all time steps, except near t = 100 (Figure 1, top row). The lack of abrupt transition at the t = 100 breaking point is due to the smoothness of the Gaussian process (as captured by the assumed length-scale, l_0). Although we could use a less smooth covariance function, such a covariance function choice would overfit the data, responding to small stochastic variations and hence not capturing the true underlying growth rate.

For simulation 2, which has a linearly increasing daily testing rate, the positives model generally overestimates the growth rate. The overestimation in the positives model is more dramatic for the

first 100 time steps, where the exponential growth rate of testing was higher (Figure 1, bottom left panel). In contrast, for the proportions model the true growth rate lies within the posterior credible interval for the majority of time steps, as in simulation 1 (Figure 1, bottom right panel).

2.5 Heterogeneity measure

Although the method is not inherently spatial, treating each set of temporal data as statistically independent, we can nevertheless use the spatial position of each spatially sampled location to address localised effects. In this way, we introduce a heterogeneity measure to assess whether exceptionally high or low growth rates within a given spatial location are a localised pattern or are caused by a larger, more widespread, phenomenon. We define the *heterogeneity* h_i of a spatial element i as

$$b_i = \frac{1}{N_i} \sum_{j \in Nbd(i)} (r_i - r_j)^2 \operatorname{sign}(r_i - r_j)$$

where r_i is the growth rate within location $i, j \in Nbd(i)$ denotes all spatial locations that neighbour element *i* (where for simplicity we assume this means share a boundary, but could be any measure of spatial locality), and N_i is the number of neighbours of *i*. Samples of h_i are taken by sampling from r_i and r_j . As such, h_i provides a measure of local covariance, with its sign reflecting whether it has higher or lower growth than its neighbours. Moreover, we can estimate other quantities that allow us to compare the heterogeneity measure of different spatial elements. For instance, we estimate $Prob(h_i > 0)$, allowing us to identify elements with considerably high heterogeneity.

3 Case-study

3.1 Data

We apply the models described above to data on daily counts of SARS-CoV-2 cases in England and in LTLAs between 1 September 2020 and 6 December 2021, dataset provided by Public Health



Figure 1. Validation of the 'positives model' and the 'proportions model'. Posterior distributions of the growth rate for simulated data under two scenarios (top—constant testing; bottom—increasing testing) and two models (left—positives model; right—proportions model). We display the median (solid line), 50% credible interval (dark shaded ribbon), and 90% credible interval (light shaded ribbon). The dotted lines indicate the true growth rate.

England (now UK Health Security Agency (UKHSA)). The data correspond to the count of people from the wider population (Pillar 2 of the UK government testing programme, UK Health Security Agency, 2021) with at least one positive PCR test, reported by specimen date and residence location (by LTLA). The data also include the count of negative tests. The dataset includes a total of 4.51 million positives cases (time series shown in Figure 2, Panel b), with test positivity ranging between 0 and 0.3 (Figure 2, Panel c), and a total test count of 62.8 million (time series shown in Figure 2, Panel a). We applied the models at a national level, to case counts in England (Section 3.2), and at a local level, to cases per LTLA in England (Section 3.3).

3.2 Growth rate estimation of SARS-CoV-2 in England from 1 September 2020 to 6 December 2021

We apply the positives and proportions models to the count of cases of SARS-CoV-2 in England between 1 September 2020 and 6 December 2021. For both models, we chose $l_0 = 50$ days and $\tau_0 = 1$ as the baseline value for the length-scale and precision, respectively, and B = I. We replicate the same choices for *h* for the approximation of the derivative as in Section 2.4. Following the implementation details in Section 2.3, the model takes less than 3.0 s of CPU time to estimate the posterior distribution of the parameters (using the package INLA 21.07.10 in R 4.1.0 using a MacBook Pro with the Apple M1 chip) and an additional 4.7 s to generate 1000 samples of the parameters.

Our time period of study contains both the second wave of infections (punctuated by a shortterm imposition of strong non-pharmaceutical interventions from 5 November 2021 to 2 December 2021) and the protracted third wave. It is clear from the data that there is a pronounced effect of weekends on the testing patterns, with lower testing but a higher proportion of positives on a weekend (shown as filled circles in Figure 2).

Both the fitted positives and proportions models had reasonable correspondence with the empirical data (Figure 2, Panels b and c). The dark-shaded ribbon shows the credible interval of the underlying Gaussian process, while the light-shaded ribbon shows the model fit including day-of-the-week effects. Our posterior distributions for the hyperparameters of the Gaussian process were confined to a narrow region of the prior distribution, showing we had garnered knowledge from the available data (Figure 3, Panel a). For the positives model, the standard deviation σ had a posterior median of 5.97 (95% credible interval 3.93–10.88) and the length-scale *l* had a posterior median of 120.97 (95% credible interval 89.12–187.73). The proportions model had a similar pattern with lower values, where the standard deviation σ had a posterior median of 2.26 (95% credible interval 1.56–3.81) and the length-scale *l* had a posterior median of 59.34 (95% credible interval 43.71–91.34).

There was usually a high level of concordance in the qualitative relationship between the growth rate estimates from the positives model and proportions model, with the models particularly well-agreeing whether the growth rate was positive or negative (Figure 2, panel d). This agreement provides additional confidence that we are seeing a robust signal from the data. Nevertheless, there were sustained periods with the two models producing dissimilar quantitative estimates, such as during December 2020. Higher differences in testing correspond to higher differences in growth rate estimation (Figure 3, panel b). This helps explain the discrepancies observed in December 2020, when testing practices are likely to be affected by the holiday period.

3.3 Spatial heterogeneity in cases of SARS-CoV-2 in the North–West region in England, April 2021

We applied the proportions model to the count of positive cases of SARS-CoV-2 in England for each of the 317 LTLAs. Since data at a lower resolution can be noisy, setting weak priors for the hyperparameters of the Gaussian process can lead to unrealistic length scales to account for the noise. To overcome that issue, we assume that the covariance function of the underlying Gaussian process at a local authority level has a similar shape to the national data. Therefore, we set the baseline values σ_0 and l_0 for the LTLA level to be the posterior median of σ and l obtained with the national data in Section 3.2, respectively, with precision B = 101.

We focus on the results from 23 April 2021, when infections with the Delta variant were increasing in the North–West of England, particularly in Bolton where our proportions model gave an

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Figure 2. Model fitting and posterior distribution of the growth rate for SARS-CoV-2 cases in England from 1 September 2020 to 6 December 2021. (Panel a) Number of tests conducted. Triangle markers correspond to reported test counts on weekdays, whilst circle marker correspond to reported test counts on weekends. (Panel b) Median (lines) and 95% credible interval (shaded ribbons) of the model fitting (light-coloured solid line, which includes a day-of-the-week effect) and the Gaussian process (dark-coloured dashed line) for the positives model. Circles and triangles correspond to the daily count of positives on weekdays and weekends respectively. (Panel c) Median (lines) and 95% credible interval (shaded ribbons) of the model fitting (light-coloured solid line) and the Gaussian process (dark-coloured dashed line) for the proportions model. Circles and triangles correspond to the proportions of positives per day. (Panel d) Median (lines) and credible interval (darker shaded ribbons for 50%, lighter shaded ribbons for 95%) for the growth rate estimations in the positives model (dashed line) and proportions model (solid line).



Figure 3. (Panel a) Comparison of the prior and posterior distributions of length-scale *I* (in days) and standard deviation σ for the positive and proportion models when applied to SARS-CoV-2 cases in England. Filled circles correspond to samples from the prior distribution. Triangles correspond to samples from the posterior distribution in the positives model. Squares correspond to samples from the posterior distribution in the proportions model. The dashed ovals represent the 95% posterior density region of each distribution. (Panel b) Comparison between the difference in testing (change in the count of tests in seven days, *y*-axis) and the difference between the growth rate estimations of the positives and the proportions model (*x*-axis). The filled circle markers correspond to the median growth rate difference between the two models, with horizontal bars representing the 95% credible interval of the difference between growth rates.

estimated positivity of 3.25% (95% PI: 2.65%–3.90%) (Figure 4, Panel a). Multiple neighbouring LTLAs in the North–West region had median estimates for proportion of tests being positive above 2%. In other regions at that time, some urban centres had a similar high incidence of 2% or above, which included Manchester and Sheffield. However, we generally measured incidence to be lower in other regions compared to the North–West. For example, all LTLAs in the South–West and along the southern coast had low median incidence estimates (below 1%).

Though there was regional structure to the magnitude of test positivity, for growth rates we observed spatial variability in areas experiencing high growth in cases and those where incidence was declining (Figure 4, Panel b). Areas expressing the greatest heterogeneity were regionally disconnected (Figure 4, Panel c). LTLAs whose probability of positive heterogeneity exceeded 0.95, thereby indicating high growth rates larger than the surrounding areas, included Erewash in the East (median heterogeneity: 1.42), Sefton in the North–West (median heterogeneity: 1.00), Bedford in the East (median heterogeneity: 0.79), and Bolton in the North–West (median heterogeneity: 0.50). For Bolton, our heterogeneity measure suggested that area was having a localised increase (>99% probability of heterogeneity being greater than zero) rather than a regionallydriven event.

Through concurrently considering the growth rate and the proportion of tests with a positive result, we could discern those LTLAs suffering from both high prevalence and high growth rates (thereby possibly requiring further support), such as Bolton and Blackburn with Darwen, and LTLAs to monitor closely due to having low prevalence but high growth rates, including Erewash, Bedford, and Sefton (Figure 5, Panel a). Although Selby has the highest estimated proportion testing positive (4.19%), the growth rate had been decreasing in the prior week (Figure 5, Panel b).

4 Discussion

In this paper, we have proposed two model structures, the positives model (which only uses data on confirmed positive cases) and the proportions model (which uses both positive and negative test information), to estimate the instantaneous growth rate of cases. We note that any measure based on cases is necessarily a lagged indicator of infectious processes due to the delay between infection



Figure 4. Epidemiological trends at the LTLA level in England on 23 April 2021. (Left) Estimated median of posterior of incidence, with lighter shading corresponding to higher incidence estimates. (Centre) Estimated median of posterior of growth rate. Regions with thicker borderlines correspond to LTLAs where the probability that the growth rate is greater than zero exceeded 95%. (Right) Median of heterogeneity. Regions with thicker borderlines correspond to LTLAs where the probability that the heterogeneity is greater than zero exceeded 95%.



Figure 5. (Left) Smooth estimation of positivity (*y*-axis) and growth rate (*x*-axis) of every LTLA in England on 23 April 2021 coloured by probability that the heterogeneity is greater than zero (darker shade for high probability, lighter for low probability). On the top axis, we state the doubling time associated with the corresponding growth rate. Vertical bars correspond to the 95% credible interval of positivity. Horizontal bars correspond to the 95% credible interval of the estimated growth rates. (Right) Trajectory of incidence-growth rate for LTLAs with high prevalence (top 2.5%) or high growth rate (top 5%) from 15 April 2021 to 23 April 2021 (squares correspond to 23 April).

and notification of disease, which generally only occurs once symptoms arise. However, as we show for simple models, our methods can robustly estimate both the growth rate and temporal changes in the growth rate, which are often related to external epidemiological factors of public-health interest.

The latent structure of both models includes a GP that interpolates the epidemic curve and approximates the underlying process that generates the disease incidences. We then take samples of the derivative of the GP to estimate the growth rate. We use a Matérn covariance function in the Gaussian process as a default choice due to its properties for simulating natural phenomena (Stein, 1999). Other covariance functions can accommodate additional assumptions, such as noise models with dependencies or seasonal effects. The models are implemented using the Laplace approximation incorporated in the INLA package in R.

Both positives and proportions models use data on positive reported infections, while the proportions model also incorporates testing counts, enabling us to account for changes in test-seeking behaviour. We believe our approach has four benefits over existing methods. First, it is rapid, robust, and computationally efficient—all of which are considerable advantages when dealing with a rapidly changing epidemic in multiple spatial locations. Second, by focusing on growth rate rather than the reproduction number, we by-pass the complexities of estimating generation-time distributions that can substantially hinder other methods early in an outbreak. Third, the combined use of positive reported infections and number of tests allows us to deal with the proportion of tests that are positive, a measure that is relatively insensitive to changes in testing behaviour. Finally, the use of Gaussian processes means that the method is also relatively robust to missing data, allowing us to provide continuous estimates even if some of the data streams are considered unreliable (for instance, the high rate of false negatives reported by the Immensa Health Clinic in some regions in the UK in September 2021, Torjesen, 2021).

Throughout we applied our method to reported cases of SARS-CoV-2 infection in England as confirmed by PCR testing. We perform our analysis both at a national scale (Figure 2) and at a small regional scale (Figures 4 and 5). Our choice of pathogen was determined by the need to quantify and explain the ongoing pandemic, feeding our findings through SPI-M-O (Scientific Pandemic Influenza Group on Modelling, Operational sub-group) to policy advisers. England has seen three major waves of infection, broadly associated with the wild-type, Alpha and Delta variants. The first wave which began in March 2020 led to large numbers of hospital admissions and deaths, but was poorly quantified in terms of infection due to the low level of community testing. The second wave began in September 2020 and peaked in late December 2020 or early January 2021 with over 60,000 cases reported on 29 December 2020. The third wave from June 2021 has been characterised by a prolonged period (over five months) of high cases, but with relatively low hospital admissions and deaths due to high vaccine uptake.

The national trends in growth rate highlight the complex pattern of growth (r > 0) or decay (r < 0) over time (Figure 2). Some notable changes that correspond to mitigation activities include: a pronounced negative growth rate during November 2020 due to the National four-week lock-down, although the growth rate had been lower in October 2020 than in September 2020; the negative growth rate during January–April 2021, during which time England was in lockdown followed by Steps 1 and 2 of the Government's COVID-19 response (UK Cabinet Office, 2021), which transitioned into high growth rates by late May 2021; a sharp drop in growth rate (especially as estimated by the positives model) in July 2021 which has been labelled as the 'pingdemic' due to the large number of individuals contacted through the Test-and-Trace App, and the potential changes in behaviour to avoid this; finally, we observe that much of August–November 2021 is characterised by growth rates close to zero, reflecting the high level of cases that have been maintained through this period.

Both the positives model and proportions model aim to capture the instantaneous growth rate of new cases and, if the efforts in testing are constant, both methods provide equivalent results. However, the estimations can differ when testing behaviour has a temporal trend - as seen during the COVID-19 outbreak in England. For instance, if the testing rate increases, the positives model can underestimate the actual growth rate (Figure 3, Panel b). In contrast, the proportions model accounts for changes in the number of tests and can give more reliable estimates. However, both models can be affected by more nuanced changes in testing behaviour; our proportions model assumes that any change in test-seeking behaviour affects all sections of the population equally—if this is not true (such as the introduction of twice-weekly lateral flow testing for secondary school children) then there can be biases. We propose to include both approaches into routine analysis since they give different perspectives to the same data, particularly when there is little knowledge of the processes driving testing behaviour in the population.

Another strength of our growth rate estimation method is the relatively low computational expense and run time, using the Laplace approximation implemented in INLA (Rue et al., 2009), permitting the application of the model at a local level (to each of the 317 LTLAs in England). Spatially, the English COVID-19 case data are either broken into seven National Health Service regions, or into 317 LTLA. LTLAs range in size from just over 2,000 people (Isles of Scilly) to well over a million (Birmingham), but most contain around 140,000 inhabitants. Performing our analysis at this spatio-temporal scale allows us to identify both highly

localised outbreaks (as shown in the maps in Figure 4) or wider regional trends, enabling scrutiny of locations exhibiting atypical data patterns. Furthermore, introducing a heterogeneity measure enabled comparisons of the growth rates between neighbouring LTLAs. The heterogeneity measure has been used during the pandemic to highlight places with abnormal growth patterns, generally identifying LTLAs with significantly higher growth. The process has also be extended (by considering S-gene target failure) to quantify the spread of new variants (e.g. Alpha and Delta) to pinpoint localities that were increasing above mere noise (Challen et al., 2021).

Our analysis of LTLAs was focused around 23 April 2021; at this time the Delta variant had begun to establish across England (with about 20% of cases attributed to Delta), hospital admissions and deaths were continuing to decline, but community cases had reached a nadir. Understanding the spatial patterns of growth at this time, and linking it to the prevalence of the Delta variant, was important for assessing the invasion of the new variant. We observe a mixed mosaic of growth rates across England (Figure 4) with a few regions where the growth rate is significantly above zero. Many of these regions also appear in the heterogeneity map as islands of growth amid a sea of declining cases; which suggests a rapid localised growth in these areas. Focusing on LTLAs that either have high growth rates or high prevalence (Figure 5) we identify three main groupings that may require further epidemiological investigation. First, there are four LTLAs (South Hams, South Northamptonshire, Erewash, and Hyndburn) that have high positive growth rates and where we expect cases to continue to rise. Second, there is a group of 15 LTLAs where a high proportion of tests (between 2% and 4%) are positive; of these Bolton, Trafford, and again Hvndburn (all in the North-West of England) are of the greatest concern due to their positive growth rate. Finally, Selby in the North-East of England (clearly identifiable on the incidence map of Figure 4) has an extremely high proportion of tests that are positive, and while the mean growth rate is slightly below zero this is not statistically significant suggesting that cases will remain high over the short term.

Our approach for estimating the growth rate is a purely statistical method and therefore has limitations. First, the model is non-mechanistic and does not incorporate any epidemiological assumptions. Therefore, it is not suitable for predicting future changes in infections or making long-term forecasts, particularly as it cannot account for the depletion of susceptible through infection or vaccination. Second, we assume that the spatial regions investigated are independent and homogeneous, we do not account for the movement of infection between regions (Kraemer et al., 2021) nor the spatial and social structure within a region. A lack of internal structure could be important for public-health concerns; for example, an outbreak that is primarily increasing in the young has very different health implications compared to one that is increasing in the elderly. There is no reason why richer data structures cannot be incorporated within our methodology (for example looking at the growth rate in a set of age-groups), but such an analysis requires large amounts of data and is increasing complex to interpret. Third, the data analysed in this study come from PCR testing (or individuals that have performed a lateral flow test followed by PCR). Therefore, there are limitations due to specificity and sensitivity of the test and the ability of individuals to swab reliably. Associated with this, and discussed above, changes to test-seeking behaviour beyond a simple increase in testing could introduce a range of biases. It is important to stress that throughout we are fitting to positive tests not infections, although we believe the two are highly correlated. Finally, though Gaussian processes provide a flexible tool, some prior knowledge of the patterns of the disease is required to inform the subjective choice of the covariance function and its priors. If the data sources are not consistent over the time course of the study, it will affect both models. Moreover, abrupt changes in the epidemic curve are harder to pick for certain covariance functions (e.g. smooth covariance functions). This highlights the need for further studies around how to design more complex covariance functions that allow such abrupt changes to be captured.

In summary, we have presented a general structure for estimating instantaneous growth rates that uses a Bayesian hierarchical model to fit a Gaussian process to the epidemiological data. Applied to high-resolution spatio-temporal SARS-CoV-2 case and testing data from England, we have demonstrated the ability of parsimonious models estimating instantaneous growth rate to both determine long-term patterns of growth at a national scale, and highlight growth and spatial heterogeneity at a regional scale.

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Code and data availability

The code used for generating the results in this paper is available at GitHub/juniper-consortium/ growth-rate-estim. Data are available to researchers with data sharing agreements in place with UKHSA.

Appendices

Appendix A. Directed acyclic graph of the models

We display the directed acyclic graph and the description of the parameters corresponding to the positives model described in Section 2.1.1 (Figure A1 and Table A1) and the proportions model described in Section 2.1.2 of the main document (Figure A2 and Table A2), respectively.

Appendix B. Growth rate comparison

Let w, n, and z be continuous functions on $\mathbb{R} \cup \{0\}$, such that at a given time $t \in \{0, 1, ...\}, w(t)$ denotes the number of new cases, n(t) denotes the number of tests, and z(t) denotes the number of positive tests. Note that although w, n, and z are continuous functions, their values have an interpretable meaning only on discrete times (for instance, daily counts). Our goal is to estimate the growth rate r(t), defined as the per capita change in the number of new cases per time; that is $r(t) = \partial_t (w(t))/w(t)$.

In the positives model, we approximate r(t) as the growth rate of observed positive tests z(t), denoted $r_z(t) = \partial_t(z(t))/z(t)$. We describe z(t) in terms of a latent function x(t) such that $z(t) = \exp(x(t))$, which simplifies the growth rate as $r(t) \approx r_z(t) = \partial_t(x(t))$.

In the proportions model, we describe the proportion of positive tests z(t)/n(t) in terms of a latent function x(t) such that $z(t)/n(t) = \text{logit}^{-1}(x(t))$. The derivative of x_t is not directly related to $r_z(t)$ as in the positives model; however, we show below it is related to $r_z(t)$ and $r_n(t)$,



Figure A1. Directed acyclic graph describing the hierarchical conditional independence structure of the positives model, described in Section 2.1.1. The parameters η , μ_t , x_t , and $w_{d(t)}$ and the hyperparameters l, σ , and τ_w are enclosed in circles. Inputs of the model are enclosed in squares.

Table A1.	Description	and the prio	r distribution	of the parar	meters of the	positives model
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	Parameter description	Prior distribution
y_t	Number of positive tests on day t	$y_t \mu_t, \eta \sim \text{Negative binomial } (\mu_t, \eta)$
η	Overdispersion of the negative binomial distribution	$\log\left(\eta\right) \sim \mathcal{N}(m_{\eta}, \tau_{\eta}^{-1})$
μ_t	Median of the negative binomial distribution	$\mu_t = x_t + w_{d(t)}$
x_t	Observations of the Gaussian process $f(t)$	$\begin{aligned} x_t &= f(t), \\ f(s) \boldsymbol{\theta} \sim \mathcal{GP}(0, k_{\boldsymbol{\theta}}(f(s), f(s'))), \end{aligned}$
l σ	Length - scale parameter of the Gaussian process Standard deviation of the Gaussian process	$\boldsymbol{\theta} = (l, \sigma)$ (log (l), log (\sigma)) ~ $\mathcal{N}((\log (l_0), \log (\sigma_0)), B^{-1})$
$w_{d(t)}$	Day-of-the-week effect (d : day of the week on day t)	$w_{d(t)} \sim \mathcal{N}(0, \tau_w^{-1})$
$ au_w$	Precision parameter of the day-of-the-week effect	$\tau_w \sim \Gamma(a_w, b_w)$

where $r_n(t) = \partial_t(n(t))/n(t)$ is the growth rate of number of tests performed. First, we compute the derivative of the function x(t)

$$\begin{aligned} \partial_t(x(t)) &= \partial_t \left[\log \left\{ \frac{z(t)}{n(t) - z(t)} \right\} \right] \\ &= \left(\frac{n(t) - z(t)}{z(t)} \right) \left(\frac{n(t)\partial_t(z(t)) - z(t)\partial_t(n(t))}{(n(t) - z(t))^2} \right) \\ &= \{r_z(t) - r_n(t)\} [1 + \exp\{x(t)\}] \end{aligned}$$

Then, we approximate r(t) as the growth rate of positive tests minus the growth rate of number of tests: $r(t) \approx r_z(t) - r_n(t) = \partial_t(x_t)/[1 + \exp{\{x(t)\}}].$



Figure A2. Directed acyclic graph describing the hierarchical conditional independence structure of the proportions model, described in Section 2.1.2. The parameters ρ , μ_t , x_t , and $w_{d(t)}$ and the hyperparameters l, σ , and τ_w are enclosed in circles. Inputs of the model are enclosed in squares.

Table A2.	Description and the prior	distribution of the	e parameters of th	he proportions m	odel, described in Section
2.1.2					

	Parameter description	Prior distribution
y_t	Number of positive tests on day <i>t</i>	$y_t \mu_t, \rho, n_t \sim \text{Beta}-\text{binomial} (\mu_t, \rho, n_t)$
n_t	Number of tests on day t	
ρ	Overdispersion of the beta-binomial distribution	$\operatorname{logit}^{-1}(\rho) \sim \mathcal{N}(m_{\rho}, \tau_{\rho}^{-1})$
μ_t	Median of the Beta-Binomial distribution	$\mu_t = x_t + w_{d(t)}$ $x_t = f(t).$
x_t	Observations of the Gaussian process $f(t)$	$f(s) \boldsymbol{\theta} \sim \mathcal{GP}(0, k_{\boldsymbol{\theta}}(f(s), f(s'))),$
$l \sigma$	Length-scale parameter of the Gaussian process Standard deviation of the Gaussian process	$\theta = (l, \sigma)$ (log(l), log(\sigma)) ~ $\mathcal{N}((\log(l_0), \log(\sigma_0)), B^{-1})$
$w_{d(t)}$	Day-of-the-week effect (d : day of the week on day t)	$w_{d(t)} \sim \mathcal{N}(0, \tau_w^{-1})$
$ au_w$	Precision parameter of the day-of-the-week effect	$\tau_w \sim \Gamma(a_w, b_w)$

Appendix C. MCMC model fitting

Model fitting in the main manuscript is performed using the integrated nested Laplace approximation implemented in the R package INLA. This approximation is suitable if the posterior distribution is unimodal. However, for the proposed models, there is no guarantee this condition is fulfilled, which might lead to the incorrect inference of the posterior distribution of the model parameters. As an alternative to model fitting, we have implemented both positives and proportions models using the Hamiltonian Monte Carlo algorithms in the software STAN and compared them to the results in Section 3.2 of the main manuscript. The MCMC was started at random initial values, with 1,000 burn-in iterations and a total of 2,000 iterations. For both models, the posterior distribution of the hyperparameters agrees with the results of the Laplace approximation implemented in INLA. Figure C1, Panel a, shows the trace plot of the hyperparameter of the positives model. Panel b shows the posterior distribution of the hyperparameters for both MCMC and the Laplace approximation. Figure C2 shows similar results for the proportions model.



Figure C1. (Panel a) Traces of the hyperparameters of the positives model (overdispersion, precision of the day-of-the-week effect, standard deviation of the GP, and length-scale of the GP) using MCMC and three different initial values. (Panel b–d) Comparison of the posterior distribution of the hyperparameters of the positives model obtained from the MCMC vs the Laplace approximation (INLA).



Figure C2. (Panel a) Traces of the hyperparameters of the proportions model (overdispersion, precision of the day-of-the-week effect, standard deviation of the GP, and length-scale of the GP) using MCMC and three different initial values. (Panel b–d) Comparison of the posterior distribution of the hyperparameters of the proportions model obtained from the MCMC vs the Laplace approximation (INLA).

The code to estimate the posterior distributions of the parameters of the model using STAN is available in the repository GitHub/juniper-consortium/growth-rate-estim. The estimates in Figures C1 and C2 take 5–7 hr of CPU time for both models (using a MacBook Pro with the Apple M1 chip).

Appendix D. Sensitivity analysis of the Gaussian process parameters

In this section, we evaluate the sensitivity of the results presented in Sections 3.2 and 3.3 of the main manuscript under different prior assumptions of the Gaussian process parameters (lengthscale l and standard deviation σ). First, we fit the positives model using the same data in Section 3.2 the for England as of main manuscript, with prior $(\log(l), \log(\sigma)) \sim \mathcal{N}((\log(l_0), \log(\sigma_0)), B^{-1}),$ under three scenarios:

- Scenario 1: same prior as before, with $l_0 = 50$ days, $\sigma_0 = 1$, and $B = \mathbb{I}$,
- Scenario 2: $l_0 = 100$ days, $\sigma_0 = 0.5$, and B = 101,
- Scenario 3: $l_0 = 25$ days, $\sigma_0 = 2$, and B = 101.

Scenarios 2 and 3 represent more restrictive priors than scenario 1. Figure D1, Panel a, shows samples and the 95% density region for the prior (light-coloured lines) and posterior distributions (dark-coloured lines) of the Gaussian process parameters under these scenarios. Changes in the prior distributions did not greatly impact the posterior. Moreover, the growth rate estimates were not considerably changed under the different scenarios (Figure D1, Panel b).

A similar analysis was performed for a dataset with noisier data. We chose to analyse Rutland, the English Local Authority with the lowest number of cases in our dataset. We fit the proportions model with the same prior shape $(\log (l), \log (\sigma)) \sim \mathcal{N}((\log (l_0), \log (\sigma_0)), B^{-1})$, under three scenarios:

- Scenario 1: same prior as before, with $l_0 = 59.34$ days, $\sigma_0 = 2.26$, and $B = \mathbb{I}$,
- Scenario 2: $l_0 = 50$ days, $\sigma_0 = 1$, and $B = \mathbb{I}$,
- Scenario 3: $l_0 = 15$ days, $\sigma_0 = 0.5$, and $B = 10\mathbb{I}$.

Scenario 2 represents a non-informative prior, while the prior in scenario 3 has a low length-scale, and low standard deviation compared to the original prior in scenario 1. The results are shown in Figure D2. Panel a in figure shows the samples and the 95% density region for the prior (light-



Figure D1. (Panel a) Prior and posterior distributions of the Gaussian process parameters under three different prior choices. Results were obtained by applying the positives model to the positive cases of England, as presented in Section 3.2 of the main manuscript. (Panel b) Comparison between the median and 95% CI of the posterior of the growth rates obtained under scenario 1 (*x*-axis) and the median and 95% CI of the posterior of the growth rates obtained under scenarios 2 and 3 (*y*-axis).

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Figure D2. (Panel a) Prior and posterior distributions of the Gaussian process parameters under three different prior choices. Results were obtained by applying the proportions model to cases in Rutland, a UK Local Authority, as presented in Section 3.2 of the main manuscript. (Panel b) Median and 95% CI of the posterior of the growth rates obtained under three scenarios for the prior of the Gaussian process hyperparameters.

coloured lines) and posterior distributions (dark-coloured lines) of the Gaussian process parameters under these scenarios. The posterior distributions under scenarios 2 and 3 differ from the posterior obtained under the original prior setting. The model relies on the prior to obtain information about the Gaussian process when data are poor. However, the effect in the growth rate posterior is not as pronounced. Figure in Panel b shows the median and 95% confidence interval of the growth rate posterior distribution for each case. The median does not differ greatly although the confidence intervals under the prior in scenario 3 are larger.

Appendix E. Exploring other covariance functions

The models presented in the main manuscript are based on a Gaussian process with the Matérn covariance function and smoothing parameter v = 3/2. In this section, we explore the effect of changing the smoothing parameter or using a squared exponential covariance function in the posterior distribution of the growth rate. We consider only covariance functions k(x, x') (or k(r)) that are strictly decreasing with respect to r = |x - x'|. For other applications, terms can be added to the covariance function to obtain different model structures, such as stationary autoregressive processes to incorporate noise models with dependencies (Murray-Smith & Girard, 2001) or terms with a seasonal component.

We repeated the analysis in Section 3.2 of the main manuscript and estimated the posterior distribution of the growth rate in England using the positives model and different covariance functions:

Matérn covariance function with v = 3/2, as in Section 3.2, of the main manuscript, with parameters length-scale *l* and standard deviation σ (Stein, 1999, Ch. 2):

$$k_{(l,\sigma)}^{M32}(f(s),f(s')) = \sigma^2 \left(1 + \sqrt{3} \frac{|s-s'|}{l}\right) \exp\left(-\sqrt{3} \frac{|s-s'|}{l}\right)$$

• Matérn covariance function with v = 5/2, with parameters length-scale *l* and standard deviation σ

$$k_{(l,\sigma)}^{M52}(f(s), f(s')) = \sigma^2 \left(1 + \sqrt{5} \frac{|s-s'|}{l} + \frac{5}{3} \frac{(s-s')^2}{l^2} \right) \exp\left(-\sqrt{5} \frac{|s-s'|}{l}\right)$$

• Squared exponential covariance function, with parameters length-scale l and standard deviation σ

$$k_{(l,\sigma)}^{SE}(f(s), f(s')) = \sigma^2 \exp\left(-\frac{(s-s')^2}{l^2}\right)$$

Figure E1 shows the comparison of the growth rate posterior distribution using a Matérn covariance function with v = 3/2 and a smoother version of the Matérn function with v = 5/2 (Panel a). A similar analysis comparison is shown for the Matérn v = 3/2 and the squared exponential covariance function (Panel b). Samples of the process with Matérn v = 3/2 are less smooth since they are one-differentiable. Samples of the process with Matérn v = 5/2 are smoother as they are two-differentiable, while samples using a squared exponential covariance are infinitely



Figure E1. Comparison of the posterior distribution of growth rates in England using different covariance functions for the positives model. (Panel a) Comparison between the squared exponential function (solid line, dark ribbon) and the Matérn function with $\mu = 3/2$ (dashed line, light ribbon). (Panel b) Comparison between the Matérn function with $\mu = 1/2$ (solid line, dark ribbon) and the Matérn function with $\mu = 3/2$ (dashed line, light ribbon).



Figure F1. Median and 95% Cl of samples of the posterior of the growth rates in England using two methods: samples of the derivative taken using finite differences (solid line, light ribbon) and samples taken from the derivative of the Gaussian process (dashed line, dark ribbon).

differentiable, and therefore unrealistic. Samples from the Matérn v = 3/2, as used in the main manuscript, are more realistic to simulate natural phenomena than processes from the squared exponential class (Stein, 1999).

Appendix F. Estimation of the growth rate using Gaussian processes

In Section 2.2, the derivative of the Gaussian process f is estimated by taking numerical approximations of the derivative from samples of the process f. Alternatively, samples of the derivative can be directly obtained by sampling from the derivative of the Gaussian process. Let $\frac{d}{dt}f(t)$ be the derivative of the process f with covariance function $k_{\vec{\theta}}(\cdot, \cdot)$ and parameters $\vec{\theta}$. If we observe n points of the original process f for t_1, \ldots, t_n , then the derivative of the process at t_1, \ldots, t_n is given by

$$\left(\frac{d}{dt}f(t_1), \ldots, \frac{d}{dt}f(t_n)\right)^T | f(t_1), \ldots, f(t_n) \sim MVN(\Lambda \Sigma^{-1}\vec{f}, \Delta - \Lambda^T \Sigma^{-1}\Lambda)$$

where $\vec{f} := (f(t_1), \ldots, f(t_n))^T$, and Σ , Λ , and Δ are $n \times n$ matrices such that $\Sigma_{ij} = k(f(t_i), f(t_j))$, $\Lambda_{ij} = \frac{\partial}{\partial t_i} k(f(t_i), f(t_j))$, and $\Delta_{ij} = \frac{\partial^2}{\partial t_i \partial t_j} k(f(t_i), f(t_j))$ (Rasmussen & Williams, 2006). We repeated the analysis in Section 3.2 of the main manuscript using the positives model in

We repeated the analysis in Section 3.2 of the main manuscript using the positives model in England, sampling from the posterior distribution of the growth rate using two methods: finite differences and direct samples from the process $\frac{d}{dt}f(t)$. Figure F1 shows the comparison of the posterior distribution of growth rates by approximating the derivative with finite differences and taking samples of the process $\frac{d}{dt}f(t)$. The 95% confidence interval for direct samples from the derivative is less wide. However, it involves sampling from the multivariate Normal distribution and the inverse of the matrix Σ , which increases with *n*. Producing the results in Figure F1 takes around 5–10 min (using a MacBook Pro with the Apple M1 chip).

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