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Strategies to Mitigate COVID-19 Resurgence Assuming Immunity Waning: A Study for Karnataka, India

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Abstract

COVID-19 vaccination is being rolled out among the general population in India. Spatial heterogeneities exist in seroprevalence and active infections across India. Using a spatially explicit age-stratified model of Karnataka at the district level, we study three spatial vaccination allocation strategies under different vaccination capacities and a variety of non-pharmaceutical intervention (NPI) scenarios. The models are initialised using on-the-ground datasets that capture reported cases, seroprevalence estimates, seroreversion and vaccine rollout plans. The three vaccination strategies we consider are allocation in proportion to the district populations, allocation in inverse proportion to the seroprevalence estimates, and allocation in proportion to the case-incidence rates during a reference period.

The results suggest that the effectiveness of these strategies (in terms of cumulative cases at the end of a four-month horizon) are within 2% of each other, with allocation in proportion to population doing marginally better at the state level. The results suggest that the allocation schemes are robust and thus the focus should be on the easy to implement scheme based on population. Our immunity waning model predicts the possibility of a subsequent resurgence even under relatively strong NPIs. Finally, given a per-day vaccination capacity, our results suggest the level of NPIs needed for the healthcare infrastructure to handle a surge.

1 Introduction

India has witnessed a significant resurgence in the number of COVID-19 reported cases since the beginning of March 2021. The number of confirmed active cases which was 165,412 on 01 March 2021 increased thirteen-fold to 2,284,411 by 21 April 2021. The state of Karnataka, a large state in South India with a population of about 70 million (estimated for 2020) and an area of approximately 192,000 km², also experienced this resurgence: from 5,945 confirmed active cases on 01 March 2021 to 196,236 confirmed active cases on 21 April 2021; the rate of positive tests increased from 0.68% to 15.87%, the number in intensive care units increased from 116 to 985, and the cumulative deaths increased from 12,343 to 13,885 during the same period [17, 18]. This resurgence has significantly stressed the healthcare system. Further, the resurgence has been heterogeneous across the districts of Karnataka, with Bengaluru Urban, Bidar, Kalaburagi, Udupi, Dakshina Kannada, and Tumakuru showing early resurgence.

India approved the use of two vaccines whose effectiveness from Phase 3 trials are now well-documented [6, 47]. To vaccinate 70%-90% of the population, Karnataka needs 98 – 126 million doses. This will likely take 6-24 months given current vaccine production levels in India (approximately 65 million doses per month

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NOTE: This preprint reports new research that has not been certified by peer review and should not be used to guide clinical practice.

during April 2021), production ramp-up plans, and all India requirements (Karnataka has about 1/20 of India's population). The strategy has thus far prioritised (*i*) healthcare workers and front-line workers from 16 January 2021; (*ii*) those 60 years and above plus those 45 years and above with co-morbidities, from 01 March 2021, given that COVID-19 disproportionately affects older adults and individuals with co-morbidities; and (*iii*) those 45 years and above from 01 April 2021. From 01 May 2021, all individuals 18 years and above became eligible for vaccination [35]. The challenges involved in vaccinating nearly a billion adults at the national level are outlined in [29].

Given the limited supply of vaccines, the eligibility of the entire adult population for vaccines, and Karnataka's heterogeneous spread of COVID-19 across its districts, a study of *vaccination allocation policies across districts* is needed to design effective vaccination strategies. There are multiple possible criteria for vaccine allocation across districts – based on population, on seroprevalence, or on case-incidence rates. It is crucial to understand how these strategies compare in terms of minimising the number of cases. Assessing the effectiveness of these strategies is complicated due to many factors, including immunity waning, new variants, mobility, non-pharmaceutical interventions (NPIs)¹, and delayed onset of vaccine effectiveness. To account for these factors and assess the overall impact of the vaccination strategies, we resort to a modeling study using a spatially explicit, age-stratified epidemiological model.

There are many district-level epidemiological models already available for Karnataka (See [1,3,25,40,43]). These models use time-series data on reported cases, confirmed active cases, tests conducted (some models), and deaths across the districts in the state. Age-stratified models have been used to understand vaccine allocation strategies for India around the time of the first wave [24]. While spatial allocations of COVID-19 vaccines have been studied in the recent literature [10, 13, 32], such a study has not been undertaken in the Indian context. We extend prior work by modelling the following additional factors.

- Karnataka conducted an extensive state-wide serological survey [4,5] during 02-16 September 2020 near the first peak. This information on the state of the pandemic at that time was used to initialise our model.
- Though more than 98% of the infected participants in a study seroconverted² [48], it is well-documented that the antibody titres decay and the decay rate is robust across asymptomatics, mild symptomatics, age groups, sex, etc. [36] at approximately 25% reduction every two weeks. Antibody titres seem to be correlated with the severity of the infection [12, 22, 27, 33]. Data seems to suggest significant seroreversion among asymptomatics [30] who constitute a sizeable majority of the COVID-19 infected population in India [28]. We account for this seroreversion in our epidemiological model.
- As of 22 April 2021, about 6.7 million people, comprising healthcare workers, front-line workers, and people over 45 years of age, or just under 10% of the population of Karnataka, received at least one dose of vaccination in phases since 16 January 2021. We account for these vaccinations and their effectiveness in our model.
- There has been an increase in transmission during the resurgence phase, perhaps due to new variants in circulation. NPIs are being imposed for six weeks during April-June 2021 to break the chain of transmission [26]. We re-calibrate the contact rates to account for this resurgence and the NPIs.

By accounting for the above aspects in our epidemiological modelling, we study the following:

- Effectiveness of three possible vaccination allocation strategies across districts: in proportion to the population, in inverse proportion to seroprevalence estimates, and in proportion to case-incidence rates;
- Possibility of a subsequent resurgence in the latter half of 2021: under an immunity waning model along with various daily vaccination capacities and NPI scenarios; and
- (iii) The target level of NPIs required: to ensure total caseload in a four-month horizon is within the healthcare infrastructure capacity given a daily vaccination budget.

The situation is fluid on the ground with significant ramping up of testing³, renewed telemedicine campaigns [20], and non-pharmaceutical interventions [26]. These will impact people's behaviour, mobility patterns, and transmission dynamics. Nevertheless, we envisage that our study will provide epidemiological insights for better public health response and can serve as a framework that can be updated and expanded to a national scale based on data availability.

 $^{^{1}}$ These include the closure of schools and colleges, workplaces, community spaces, mobility restrictions, and varying degrees of lockdowns.

 $^{^{2}}$ But see [44] which is an Indian cohort study that reported only 85% seroconversion rate.

 $^{^{3}}$ There has been a 50% increase in testing from 118,933 tests on 01 April 2021 [19] to 162,534 tests on 21 April 2021 [17].



Figure 1: Datasets for our epidemic simulator. The epidemic simulator takes data from several sources. Disease parameters come from the current state of knowledge of how COVID-19 affects an infected individual. Case surveillance data include tests, reported cases, discharges, and deaths. Serosurveillance data comes from a state-wide survey done in September 2020 during the first peak. The timings and the duration of the NPIs are based on when they come into effect in the state. A mobility model captures the spread of infection in the units before the April 2021 resurgence. Karnataka's actual geography enables better visualisation of the simulation outcomes. Demographic data for the units are taken from the 2011 census and extrapolated to 2020. Vaccine supply assumptions are based on historical vaccination data. Experimental vaccination allocation policies are based on these data elements. The epidemic simulator takes all these and generates outcomes for a comparative study.

2 Methods: Data, Modelling, and Experiment Design

Karnataka has 30 administrative zones called districts. We treat all districts other than the capital Bengaluru Urban district as individual spatial units. Since the Bengaluru Urban district is heavily populated, we further subdivide it into nine units. This yields a total of 38 units which were also the units used in the state-wide serosurvey conducted in September 2021 [4,5].

We use a spatial and age-stratified compartmental model to study the disease spread within Karnataka. There are five compartments: susceptible (S), exposed (E), infected (I), recovered/deceased (R) and vaccinated (V). The infected compartment includes both detected and undetected infections. The population is divided into sub-units, with each sub-unit comprising an (age-group, unit) pair, where a unit is one of the 38 units described above. The model keeps track of the numbers in the five compartments in every sub-unit and lets them evolve over time. The trajectory is obtained by time-discretisation and solutions to difference equations [45, 46], which we shall soon describe.

Datasets: We use the following datasets in our model. The datasets used are portrayed in Figure 1 and summarised in Table 1.

- Population data is based on the official projected population data for 2020 [21]. The age distribution is based on a 2017 projection from [37].
- Age-stratified contact rates [38] modulate the number of contacts made between people of various age groups.
- Disease parameters (incubation period, mean recovery time) are based on [11, 23, 34].
- We leverage the unit-wise active infection and past infection data from the Karnataka-wide round-1 seroprevalence study [4,5] to initialise our simulation on 11 October 2020.
- Time series of daily reported cases data and tests conducted in each unit are taken from the official daily bulletins (e.g., [18], summarised in [42]) starting from 11 October 2020. The data until 22 April 2021 is used for model calibration, and the data until 11 May 2021 is used for validation and uncertainty quantification.
- To move from reported cases to infections, we use the unit-wise reported cases-to-infections ratio (CIR) estimated from the Karnataka round-1 seroprevalence study [4,5]. Further, we use testing data



Figure 2: Compartments, disease progression, vaccination, and immunity waning (antibody decay) components of the model.

from March 2021 to modify the CIR on a weekly basis. This is to account for significantly increased testing since March 2021. The testing data are taken from Karnataka's daily COVID-19 bulletins.

• We also use the daily number of vaccinations administered in each unit and assume, for simplicity, a conservative 66% efficacy 14 days after one dose⁴.

Model: Let \mathcal{D} denote the set of units and let \mathcal{A} denote the set of age groups. A sub-unit (also referred to as a *patch*) of the population is identified with a given unit and an age group; let $u = (i, x) \in \mathcal{D} \times \mathcal{A}$ be a patch of the population. Let $N_{(i,x)}$ denote the population of patch (i,x). Let us consider the evolution of the exposed population at u. People from patch u get exposed to the virus from the infected population in u as well as from the infected population in other patches v owing to mobility across units and interactions across age-groups. We model the impact of mobility on infection spread by keeping the population of the patches constant and by adjusting the within-unit and cross-unit contact rates based on a mobility matrix. Let $\Theta(t)$ denote the unit-wise travel matrix at time t, i.e., $\theta_{j,i}(t)$ denotes the probability that an individual in unit j spends time in unit i at time t. Let M denote the matrix of age-stratified interaction rates, i.e., $M_{x,y}$ is the number of contacts made by individuals of age group x with age group y on a typical day (time unit being one day). We also define $\beta_i(t)$, a piece-wise constant function, to be a contact rate modulating parameter at time t. Let $S_{(i,x)}(t), E_{(i,x)}(t), I_{(i,x)}(t), R_{(i,x)}(t)$ and $V_{(i,x)}(t)$ denote the number of people in susceptible (S), exposed (E), infected (I), recovered/deceased (R), and vaccinated (V) states, respectively, in patch (i, x) at time t; see Figure 2. Assuming that a tagged individual in patch u = (i, x) comes in contact with a tagged individual in patch v = (j, y) at rate $\beta_i(t)\theta_{j,i}(t)M_{x,y}/N_{(j,y)}$ effective contacts per unit time, the change in the exposed population at time t in patch u = (i, x), defined as $\Delta E_{(i,x)}(t) = E_{(i,x)}(t+1) - E_{(i,x)}(t)$, is given by

$$\Delta E_{(i,x)}(t) = \beta_i(t) S_{(i,x)}(t) \sum_{j \in \mathcal{D}} \theta_{j,i}(t) \sum_{y \in \mathcal{A}} M_{x,y} \frac{I_{(j,y)}(t)}{N_{(j,y)}} - \alpha E_{(i,x)}(t)$$
(1)

where α is the rate at which individuals move from the exposed state to the infected state (i.e., $1/\alpha$ is the mean incubation period). Let γ denote the recovery rate, i.e., $1/\gamma$ is the mean time to recovery or death.

 $^{^{4}}$ At the time of writing, as indicated in the Introduction, two vaccines were being administered in India. The guidelines recommend two doses of the vaccine spaced twelve weeks apart. For effectiveness studies, see [47] and the interim phase 3 results [6].

Dataset	Description	Reference
Population	The population of each unit of Karnataka (estimated for 2020).	[21]
Age distribution	The age distribution of the population in Karnataka. This is used	[37]
	to find out the population of each age group in each unit.	
Age stratified contact	The number of contacts made by an individual of a given age	[38]
rates	group with another individual of another age group. This is used	
	to modulate the contact rates based on age-groups.	
Disease parameters	The mean incubation time in and the mean recovery time of	[11, 23, 34]
	COVID-19 infected individuals. These are used to model the	
	spread of the disease.	
Seroprevalence data	Unit-wise antibody (IgG) prevalence and active prevalence of	[4, 5]
	COVID-19 from the Karnataka-wide round-1 serosurvey con-	
	ducted during 02–16 September 2020. The round-1 data is used	
	to initialise the simulator. We then use a Weibull model for im-	
	munity waning.	
Daily new reported	The time-series of unit-wise daily new reported cases since	[42]
cases	11 October 2020. This is used to calibrate the contact rates.	
Tests	The time-series of tests done in the state of Karnataka since	[42]
	19 February 2021. This is used to modify the cases to infections	
	ratio.	
Vaccinations	The time-series of unit-wise vaccine doses administered since	[42]
	16 January 2021.	

Table 1: Summary of datasets used in the simulator, their brief descriptions, and references.

Then the changes in the variables S, I, and R for the patch (i, x) at time t are modelled as

$$\Delta S_{(i,x)}(t) = -\beta_i(t) S_{(i,x)}(t) \sum_{j \in \mathcal{D}} \theta_{j,i}(t) \sum_{y \in \mathcal{A}} M_{x,y} \frac{I_{(j,y)}(t)}{N_{(j,y)}} - \Delta V_{(i,x)}(t) + \Delta W_{(i,x)}(t), \tag{2}$$

$$\Delta I_{(i,x)}(t) = \alpha E_{(i,x)}(t) - \gamma I_{(i,x)}(t),$$
(3)

$$\Delta R_{(i,x)}(t) = \gamma I_{(i,x)}(t) - \Delta W_{(i,x)}(t).$$

$$\tag{4}$$

Additionally, $V_{(i,x)}(t)$ is incremented based on (for simplicity) the number of first-dose vaccines administered fourteen days prior to t in that unit, after taking into account the efficacy of the vaccine. This data is extracted from bulletins like [18] and is summarised in [42]. The 14-day delay accounts for the time for antibodies and immunity to develop. The increment to the vaccinated pool $\Delta V_{(i,x)}(t)$ depends on the number vaccinated in age group x of district i and the effectiveness of the vaccine, which is conservatively set to 66%, the lower of the two effectiveness numbers reported in [6, 47] after two doses.

In addition to the disease progression dynamics (1)–(4), we also consider an agent-based immunity waning factor in our model. The feature, also known as seroreversion, is motivated by the observed reduction in seroprevalence in the densely populated areas of Mumbai [41]. Both reinfections from the original strain due to antibody waning and cross-strain infections are possible factors driving the increasing trend in the daily number of reported cases in Karnataka starting early March 2021. In our implementation of immunity waning and seroreversion, individuals in the recovered state are assumed to lose their immunity after a random amount of time which has the Weibull distribution with shape $\kappa = 3.67$ and scale $\lambda = 120$ days, a model inspired by that in [7] but with a faster median seroreversion period to account for the resurgence. We sample a duration from this Weibull distribution for each recovered individual and move that individual from the recovered state to the susceptible state after this duration. This is the $\Delta W_{(i,x)}(t)$ in (2) and (4).

Calibration: Certain parameters are kept fixed throughout the simulation duration. These are the mean incubation period $1/\alpha$ (assumed to be 5.8 days) and the mean infectious period $1/\gamma$ (assumed to be 5 days). These are based on [11,23,34]. The piece-wise constant contact rate parameters $\beta_i(\cdot)$ are tuned to minimise the per day squared error between the logarithm of daily-new-reported-cases time-series and that of the model (see Appendix 4.2). We aim to match the time-series for each unit by calibrating its contact rate. Since our simulator tracks infections, we multiply the reported cases by the estimated cases-to-infections ratio available from the Karnataka-wide round-1 seroprevalence study [4, 5] as the total infections that the

simulator should track. To account for the increased testing since March 2021, we modify the CIR every week based on the ratio of the average number of daily tests done in a given week to the average number of daily tests done during 18–28 February 2021. Finally, we consider a 7-day moving average for the daily infected time-series to smooth the curves for tracking.

In summary, the following steps are performed during calibration:

- Start the simulator with susceptible, infected and recovered fractions matched to the round-1 seroprevalence data projected to 11 October 2020. Tune the contact rate parameters during 11–31 October 2020 to match the number of reported cases on 01 November 2020 within 10%.
- Tune the unit-wise contact rate parameters (one scalar per unit) during the period 01 November 2021 28 February 2021 to bring the per day squared error to within 0.1 for each unit. We use the identity matrix for mobility during this period, so the disease evolves independently in each unit. We update the contact rates in each iteration after evolving the time-series over the duration.
- Tune the unit-wise contact rate parameters during 01–15 March 2021 to match the number of modelreported cases on 15 March 2021 within 10% of the actual reported cases. Then tune the unit-wise contact rate parameters during 16 March – 07 April 2021 to minimise the per day squared error during this period. To account for the stochasticity due to a low number of reported cases in many units, we introduce moderate uniformly mixing mobility, i.e., $\Theta(t) = (1 - \varepsilon)I + \varepsilon J$, where I denotes the identity matrix of size $|\mathcal{D}|$, J denotes the all-one matrix of size $|\mathcal{D}|$, and $\varepsilon = 0.01$.
- Repeat the above for the duration 08–22 April 2021 to arrive at the corresponding calibrated contact rates for this period. From this period onward, since the infections have been seeded, we go back to diagonal mobility, i.e., $\Theta(t) = I$. If mobility data is available, then the mobility matrix $\Theta(t)$ can be appropriately set.
- For the projections, we redo the calibration for the period 08 April–01 May 2021 and hold the resultant contact rate parameters constant into the future.
- Finally, to account for the delay in sample collection and test outcomes, we delay the trajectory of reported cases by one week.

The immunity waning Weibull distribution is fixed to have shape $\kappa = 3.67$ and scale $\lambda = 120$ and is common across all units. This yields a median seroreversion time of 109 days.

Validation: To validate the performance of our calibrated simulator, we use the tuned contact rates during 08-22 April 2021 and run the simulator with these contact rates during 23 April – 11 May 2021. We validate the simulator by comparing the simulator's projections with the actual reported cases during 23 April – 11 May 2021. To quantify the uncertainty in our predictions, we compute the root mean squared error between the projected number of cases and the actual reported cases between 23 April – 11 May 2021.

Experiment design: We use our simulator to estimate the number of reported cases for each unit in Karnataka until 31 December 2021. However, we caution that we do not have uncertainty quantifications for the longer-term projections (beyond two weeks). The experiments are described below.

We first consider two sets of scenarios to study the effect of non-pharmaceutical interventions (NPIs). These are the following.

- The first set is the setting of No-NPI. Here the mobility matrix during 02 May 2021 31 December 2021 is assumed to be $\Theta(t) = I$, i.e., the same as that of the period 01 April 2021 01 May 2021 (violet extrapolation in Figure 3).
- The second set consists of three scenarios with varying levels of NPI during 02 May 2021 31 December 2021:
 - (i) 1/3-NPI with mobility matrix $\Theta(t) = \frac{2}{3}I$;
 - (ii) 1/2-NPI with mobility matrix $\Theta(t) = \frac{1}{2}I$;
 - (iii) 2/3-NPI with mobility matrix $\Theta(t) = \frac{1}{3}I$.

The above scenarios are depicted in Figure 3. Note that $\Theta(t) = \alpha I$ implies that the number α is the reduction in the effective contact rate from a reference level of 1. We choose to study three different levels to account for uncertainties in the actual level of NPI and compliance.



Figure 3: Mobility scenarios considered for the simulation experiment design. The purple band indicates a period when the mobility matrix is $(1 - \varepsilon)I + \varepsilon J$. The mobility is I at all other times.

We also design three vaccination policies. Starting from 23 April 2021, vaccines are allocated to units in proportion to:

- The population of the units;
- The inverse of the model-predicted seroprevalence on 18 February 2021; and
- The case-incidence rates during 01–15 April 2021.

We have chosen 18 February 2021 as the reference date for the inverse-seroprevalence strategy to match the second seroprevalence study, which is in progress. For each policy above, starting from 23 April 2021, we assume that the daily budget of available vaccine doses is 167000 (which is the average daily number of first doses of vaccines administered in the state of Karnataka during 01–15 April 2021). Until 22 April 2021, we use the actual number of the first dose of the vaccine administered; this data is available in [16]. Since the vaccination drive started with healthcare workers and subsequently extended to people above 60, 45 and then 18 years of age, we assume that the vaccines allocated to a unit get equally distributed among the age groups of 18+ during 16 January 2021 – 15 March 2021 (healthcare workers), equally distributed among the age groups of 40+ during 16 March 2021 – 30 April 2021, and equally distributed among the age groups of 18+ after 01 May 2021, respectively. Even though India has a significantly younger population, in our simulator, the distribution of vaccines across age groups has a desirable natural bias towards the elderly, who are more susceptible to COVID-19.

Finally, we also study two decreased and seven increased daily vaccination capacities where the 167000 is replaced by 100000, 133000, 200000, 233000, 267000, ..., 400000 doses per day (first dose). See Table 2.

The outcomes of our experimental study are summarised using:

- The variation in the occurrence of the peak of daily reported cases across the units of Karnataka under various NPIs;
- The short-term and long-term cumulative number of reported cases in the units of Karnataka under the three vaccination policies; and
- The reduction in the number of projected cases with increase in the daily budget of available vaccine doses for various NPIs.

The outcomes will quantify the public health benefit of NPIs over No-NPI, study the effectiveness of the three vaccination policies, and highlight the limitations/benefits of reduced/increased vaccine allocations across various levels of NPIs.



Figure 4: Validation plots for the state of Karnataka. The vertical line in both plots separates the calibration and the validation periods. The root mean squared error during the validation period is at most 9% for the state of Karnataka.



Figure 5: Uncertainty quantification for the validation period in each unit of Karnataka. More than one-half of the units have a root mean squared error of less than 20%. Five districts have errors between 40-50% which are due to the sudden change in trends in the reported cases not captured by the model.

3 Results

Validation and uncertainty quantification: The prediction errors in the simulator's projections for the entire state of Karnataka are visualised in Figure 4. The projections are in blue and should be compared with the actual reported cases in red on the right-hand side of the vertical line marking 22 April 2021. Similar validation plots for the individual units can be found in Figure 13. For the state of Karnataka, we find that the root mean squared error between the projected number of cases and the reported number of cases is about 9%. The percentage error for each unit is visualised in Figure 5. While the short-term prediction errors for more than half of the units are within 20%, the error for some units such as Raichur and Vijayapura are close to 50%. Longer-term prediction errors will only be larger. Further, there is a steep rise in the prediction errors for the units Chikkaballapur, Kodagu, Kolar, Vijayapura and Raichur.

Future projections and peaks: Figure 6 depicts the daily and the cumulative number of reported cases in Karnataka until 31 December 2021 under various NPIs. In these plots, we assume that a daily budget of 167000 vaccine doses is available and distributed in proportion to each unit's population. Note that these are long term projections and should be interpreted with caution.

The units are classified as E1 < E2 < M1 < M2 < L1 < L2 with E standing for early peaks (May 2021), M standing for mid-range peaks (June 2021), and L standing for late peaks (July 2021 and after). E1 and E2



Figure 6: The projected number of new infections for the state of Karnataka under various NPIs. These projections are for the vaccination policy that allocates 167000 vaccine doses per day proportional to each unit's population. Uncertainty bands are not indicated to reduce clutter. Recall the 9% root mean squared error for a two-week ahead projection. The left figure is for the daily new reported cases, and the right figure provides the cumulative reported cases.



Figure 7: The evolution of susceptible fraction in the state of Karnataka over time. These plots are for the vaccination policy that allocates 167000 vaccine doses per day proportional to each unit's population. Observe that the susceptible fraction increase during the period August-November 2021 due to antibody waning.

stand for the first and second half of May 2021, respectively, with similar interpretations for M1, M2, and L1, L2.

Peaks – The counter-factual case of No-NPI: Under the counter-factual No-NPI scenario, the state-level peak may have occurred during mid-May 2021. For the unit-wise daily and the cumulative number of reported cases, see Figure 11. The model suggests that the units would have peaked at different times. Most of the Bengaluru units are in the E category (either E1 or E2) and Kalaburagi, Bidar, Kodagu, and Tumakuru. Except for Kodagu, these units saw an early resurgence. Kodagu saw a steep rise in infections, which suggests an early peak. Bagalkote, Chitradurga, Haveri, Dharwad and Gadag are in the L category. See the top two subfigures in Figure 9 for the No-NPI setting. The left-hand side figures present the E-M-L categorisation, while the right-hand side figures present a spatial view of this categorisation.

Peaks – The various NPIs: Under 1/3-NPI, the model predicts that only about a third of the units will have peaks after mid-May 2021 (E2 and beyond). Under 1/2-NPI and 2/3-NPI, the model predicts that cases reduce from as early as mid-May 2021 in every unit and also in the state.

Projected cumulative cases on 31 August 2021 – A comparison: The projected cumulative reported cases at the end of August 2021 seem to have significant reduction due to these NPI assumptions, see Figure 6 and Tables 4–7. These assume 167000 vaccination doses per day. Compared to No-NPI, on 31 August 2021, the model projects that 1/3-NPI may reduce the (cumulative reported) cases by roughly 926,000, 1/2-NPI may reduce it by another 431,000 cases, and 2/3-NPI may reduce it by another 306,000 cases. This order of magnitude reduction appears robust across all three vaccination policies, which we discuss next.

Comparison of the three vaccination policies: Table 5 provides a summary of the cumulative modelprojected reported cases on 15 May 2021 and 31 August 2021, respectively, starting from a reference level taken to be zero on 01 November 2020 (which is the reference date)⁵. This is under the three vaccination policies assuming that (i) the daily budget of available vaccine doses is 167000 and (ii) 1/3-NPI is in force starting from 02 May 2021. The (short-term) model-projected cumulative number of reported cases in the state on 31 May 2021, starting from 01 November 2020, is 1.718, 1.720 and 1.705 million under the vaccination policies of the population proportional, inverse seroprevalence proportional and case-incidence rate proportional strategies, respectively. This suggests that all three vaccination strategies are similarly effective in the short term according to the model. Similarly, the model-projected cumulative reported cases on 31 August 2021 are 2.222 million, 2.249 million and 2.254 million under the same three policies, respectively.

While the above was for 1/3-NPI, Table 4, Table 6 and Table 7 summarise the model-projected cases under No-NPI, 1/2-NPI and 2/3-NPI, respectively. All of these are within 2% of each other. However, a finer observation indicates that the incidence-rate proportional allocation is more effective in the short term, but the population-proportional allocation is effective in the long-term when viewed at the level of the state⁶. In the BBMP units, the incidence-rate proportional allocation continues to remain the most effective among the three.

The benefit of increased daily vaccinations: Table 2 summarises the effect of increased/decreased number of daily vaccinations under various NPIs, assuming that vaccines are allocated in proportion to population. Under No-NPI, the second column in Table 2, we find that the cumulative number of projected cases on 31 August 2021 reduces from 3.27 million to 2.82 million as the number of daily vaccinations increases from 100000 to 400000. Reductions are observed in the other NPIs as well. Going row-wise, for a fixed daily vaccination budget, we find a significant reduction in the number of cumulative cases as NPIs are strengthened, as expected. Going column-wise, an increase of vaccination capacity is most effective when the intensity of NPI is low. Table 3 summarises the reductions in percentage with respect to the reference number of cases under No-NPI and 100000 vaccinations per day.

Resurgence: Table 8 shows the susceptible population percentage (model-predicted) in each unit. We note that with 100000 vaccination capacity per day, 15/38 units have more than 60% susceptible on 31 August 2021. See also Figure 7 which shows that the susceptible fraction increases during August-November 2021 due to antibody waning. Figure 6 shows a future peak between mid-November and mid-December 2021. Figure 11 shows the resurgence in the units. Except for Bidar and Kalaburagi, all show

 $^{^{5}}$ To compare with the actual cumulative COVID-19 cases, one must add our increments from the reference date to the actual cumulative cases of 827,064 on 01 November 2020.

 $^{^{6}}$ In the case of 2/3-NPI, the incidence rate proportional allocation does marginally better.

Table 2: The projected number of cumulative reported cases in the state of Karnataka on 31 August 2021 under the four NPI intensities and various daily budgets of available vaccine doses. Vaccinations are assumed to be allocated in proportion to the population of each unit.

Daily budget	No-NPI	1/3-NPI	1/2-NPI	2/3-NPI
100000	3266600	2278100	1815200	1490700
133000	3206300	2249400	1803000	1487500
167000	3147900	2221800	1790400	1484600
200000	3094100	2196100	1779400	1480900
233000	3043300	2172400	1769400	1477400
267000	2993600	2148800	1758400	1474200
300000	2948300	2127600	1749200	1470900
333000	2905200	2107200	1740000	1468000
367000	2863000	2087000	1730900	1465500
400000	2824200	2069100	1722400	1462800

Table 3: Reduction in the number of cumulative projected cases on 31 August 2021 under various NPI intensities and daily vaccination budgets. Vaccinations are assumed to be allocated in proportion to the population of each unit.

Daily budget	No-NPI	1/3-NPI	1/2-NPI	2/3-NPI
100000	Reference	30.3%	44.4%	54.4%
133000	1.8%	31.1%	44.8%	54.5%
167000	3.6%	32.0%	45.2%	54.6%
200000	5.3%	32.8%	45.5%	54.7%
233000	6.8%	33.5%	45.8%	54.8%
267000	8.4%	34.2%	46.2%	54.9%
300000	9.7%	34.9%	46.5%	55.0%
333000	11.1%	35.5%	46.7%	55.1%
367000	12.4%	36.1%	47.0%	55.1%
400000	13.5%	36.7%	47.3%	55.2%

resurgence before December 2021. In our model, Bidar and Kalaburagi will show resurgence since the infection is never wiped out in the simulator. The resurgence in our model can be attributed to residual levels of infection in the unit, which then begins to increase as immunity wanes. Mobility and variants that can escape immunity are not taken into account.



Figure 8: Surface plot (top figure) and contour plot (bottom figure) summarising the reduction in the number of model-projected cases on 31 August 2021 for varying daily vaccination budgets and NPI intensities. The surface plot and the contours are based on linear interpolation from the data points in Table 3.



Figure 9: Left subfigure: The time of occurrence of the peak number of reported cases for each unit under No-NPI, 1/3-NPI, 1/2-NPI and 2/3-NPI, respectively. E, M, L stand for early, mid, late peaks, and the ordering is: E1 < E2 < M1 < M2 < L1 < L2. Right subfigure: Geo-spatial representations of the two scenarios. Spatial heterogeneity manifests in the No-NPI setting. In the 1/2- and 2/3-NPI settings, the cases start to recede from mid-May 2021. Note that the NPI intensities are held constant until 31 December 2021 in our simulations.

Unit	Population-	InverseSero-	IncidenceBate-	Population-	InverseSero-	IncidenceRate-
	proportional	proportional	proportional	proportional	proportional	proportional
	31 May	31 May	31 May	31 August	31 August	31 August
BBMP-Bommanahalli	93000	93200	89300	112500	113500	99200
BBMP Deceraballi	51100	50800	46800	60000	67800	53200
BBMP Fast	148000	148800	146100	175800	181000	167500
BBMP Mahadownura	120400	120400	11/100	151300	151000	107500
BBMP BB Nagar	81600	81100	76000	85700	84300	78200
BBMP-South	13/100	13/000	132900	1/1900	145100	138800
BBMP_West	133000	133700	128600	130300	141700	131000
BBMP-Velahanka	79000	78600	72100	10/100	102400	80700
Best of Bengaluru-	103500	103500	99900	171000	173100	1/3000
Urban	105500	105500	55500	111500	175100	140500
Bagalkot	32000	33200	33500	106700	11/100	122800
Ballari	86800	87700	88500	122600	127400	122000
Belgaum	40000	40600	40800	93000	103000	105100
Bengaluru-Bural	37600	37700	37600	44100	44500	44400
Bidar	18900	18700	18900	25800	24200	26400
Chamarajanagar	26100	26200	26500	31200	31600	32600
Chikkaballapur	60600	60600	61600	95800	95600	102200
Chikmagalur	24900	24700	25200	35900	35100	38000
Chitradurga	11500	11600	11700	30000	31800	34400
Dakshina-Kannada	52400	52900	53000	94400	101400	103700
Davanagere	28200	28500	28800	59600	62600	66200
Dharwad	42400	42100	42900	175300	160700	207000
Gadag	8900	8900	9000	47400	46000	60600
Hassan	47800	47500	48200	69100	67400	72900
Haveri	9600	9300	9700	35000	27200	41200
Kalaburagi	51100	51100	51500	68800	68900	72800
Kodagu	21200	21100	21300	22100	21900	22400
Kolar	44400	44800	45100	56600	58400	59600
Koppal	42100	41800	43000	76600	74300	83600
Mandva	54600	55300	55300	65300	68000	68300
Mysuru	128400	130300	130100	177600	190300	188800
Raichur	46900	47100	47900	57000	57600	60500
Ramanagar	21600	21500	21900	29100	28700	30800
Shivamogga	37000	37100	37500	90700	93500	102100
Tumakuru	61200	61500	61900	67300	68500	69800
Udupi	33600	33700	33700	54200	55400	56400
Uttara-Kannada	35400	35300	35900	78100	76400	84800
Vijayapura	28700	28900	29100	40600	42100	43900
Yadgir	30300	29300	30900	45700	41000	49000
Karnataka	2109100	2113900	2087600	3147900	3177300	3202500

Table 4: Comparison of unit-wise cumulative reported cases (since 01 November 2020) on 31 May 2021 and 31 August 2021 under No-NPI with a daily budget of 167000 vaccine doses.

Unit	Population-	InverseSero-	IncidenceRate-	Population-	InverseSero-	IncidenceRate-
	proportional	proportional	proportional	proportional	proportional	proportional
	31 May	31 May	31 May	31 August	31 August	31 August
BBMP-Bommanahalli	79900	79900	77900	90400	90900	83900
BBMP-Dasarahalli	42100	41900	39900	50800	50000	43300
BBMP-East	129100	129500	128100	143400	145600	139800
BBMP-Mahadevpura	102900	102800	99700	117300	117300	106900
BBMP-RR-Nagar	76500	76100	73600	79000	78300	74400
BBMP-South	125400	125900	124700	129800	131200	128300
BBMP-West	125400	125800	122700	129000	130100	124400
BBMP-Yelahanka	65700	65400	62200	77900	77200	66900
Rest of Bengaluru-	79500	79500	77900	111500	112300	99200
Urban						
Bagalkot	21300	21400	21500	51700	56000	60700
Ballari	63600	64000	64400	93400	97500	100900
Belgaum	26900	27100	27200	54400	61000	62400
Bengaluru-Rural	32500	32600	32600	36300	36500	36400
Bidar	16600	16500	16600	18400	18000	18500
Chamarajanagar	21000	21000	21200	26000	26300	27100
Chikkaballapur	42400	42400	42900	70600	70800	76200
Chikmagalur	19100	19000	19200	26000	25500	27200
Chitradurga	8500	8500	8500	14800	15600	16600
Dakshina-Kannada	40800	41000	41000	56500	59000	59800
Davanagere	18900	19000	19200	37600	39900	42500
Dharwad	30100	30000	30300	56500	53900	62600
Gadag	6500	6500	6600	11000	10900	12000
Hassan	37700	37600	37900	49200	48400	51200
Haveri	6300	6200	6400	14900	12100	17700
Kalaburagi	43400	43400	43600	50000	50100	51200
Kodagu	19500	19400	19500	20400	20300	20600
Kolar	35400	35500	35700	44600	45900	46700
Koppal	28000	27800	28300	52800	51400	58600
Mandya	45300	45600	45700	53300	55100	55300
Mysuru	100000	101000	100800	132300	140300	139400
Raichur	37100	37100	37600	47000	47500	49900
Ramanagar	16100	16000	16300	22700	22500	24100
Shivamogga	25900	25900	26100	47700	49300	53200
Tumakuru	54500	54600	54900	58900	59500	60300
Udupi	26500	26500	26600	34700	35200	35600
Uttara-Kannada	24000	23900	24200	47400	46700	52100
Vijayapura	22500	22600	22700	29400	30300	31200
Yadgir	21300	20800	21600	34300	30800	37200
Karnataka	1718000	1720000	1705400	2221800	2249100	2254200

Table 5: Comparison of unit-wise cumulative reported cases (since 01 November 2020) on 31 May 2021 and 31 August 2021 under 1/3-NPI with a daily budget of 167000 vaccine doses.

Unit	Population-	InverseSero-	IncidenceRate-	Population-	InverseSero-	IncidenceRate-
	proportional	proportional	proportional	proportional	proportional	proportional
	31 May	31 May	31 May	31 August	31 August	31 August
BBMP-Bommanahalli	73700	73800	72500	79500	79700	76100
BBMP-Dasarahalli	38100	38000	36700	42300	42000	38700
BBMP-East	120300	120500	119600	128100	129100	126300
BBMP-Mahadevpura	95000	95000	93000	102400	102400	97200
BBMP-RR-Nagar	73800	73600	71700	75400	75000	72400
BBMP-South	120900	121300	120500	123700	124500	122800
BBMP-West	121300	121600	119500	123700	124300	120700
BBMP-Yelahanka	59800	59800	57700	66000	65800	60400
Rest of Bengaluru-	69700	69800	68900	83800	84200	79000
Urban						
Bagalkot	17200	17300	17300	27200	28400	29700
Ballari	52600	52900	53200	71200	73700	75700
Belgaum	22000	22200	22200	33000	35400	35900
Bengaluru-Rural	30100	30200	30200	32300	32400	32300
Bidar	15600	15600	15600	16400	16300	16400
Chamarajanagar	18100	18200	18300	21800	22000	22500
Chikkaballapur	34500	34500	34700	50900	51100	54200
Chikmagalur	16500	16500	16600	20100	19900	20700
Chitradurga	7400	7400	7400	9500	9700	9900
Dakshina-Kannada	36200	36300	36400	42500	43300	43600
Davanagere	15400	15400	15500	23600	24600	25600
Dharwad	25800	25800	26000	33400	32900	34700
Gadag	5700	5700	5700	6900	6900	7200
Hassan	33400	33300	33500	39000	38700	39900
Haveri	5200	5200	5300	7800	7100	8400
Kalaburagi	40200	40200	40300	43300	43300	43700
Kodagu	18300	18300	18400	19100	19000	19200
Kolar	30900	31000	31100	36600	37300	37700
Koppal	22200	22200	22400	35200	34600	38300
Mandya	40500	40800	40800	45600	46600	46700
Mysuru	87300	87800	87700	104900	108700	108200
Raichur	31600	31700	32000	38900	39300	40800
Ramanagar	13400	13400	13500	17700	17600	18600
Shivamogga	21800	21800	21900	29700	30200	31300
Tumakuru	50900	51000	51200	53800	54200	54600
Udupi	23700	23700	23700	27200	27400	27500
Uttara-Kannada	19700	19600	19800	29400	29200	31200
Vijayapura	19700	19800	19900	23300	23700	24100
Yadgir	17200	16900	17400	25200	23200	27000
Karnataka	1546100	1548000	1538200	1790400	1803300	1799100

Table 6: Comparison of unit-wise cumulative reported cases (since 01 November 2020) on 31 May 2021 and 31 August 2021 under 1/2-NPI with a daily budget of 167000 vaccine doses.

Unit	Population-	InverseSero-	IncidenceRate-	Population-	InverseSero-	IncidenceRate-
	proportional	proportional	proportional	proportional	proportional	proportional
	31 May	31 May	31 May	31 August	31 August	31 August
BBMP-Bommanahalli	68200	68200	67500	70700	70700	69200
BBMP-Dasarahalli	34800	34700	34000	36400	36200	34900
BBMP-East	112400	112500	112000	115800	116100	115000
BBMP-Mahadevpura	88100	88100	86900	91200	91100	88900
BBMP-RR-Nagar	71200	71000	69800	72000	71800	70200
BBMP-South	116600	116800	116300	118100	118500	117600
BBMP-West	117400	117600	116200	118700	119000	117000
BBMP-Yelahanka	54900	54700	53600	57300	57000	54800
Rest of Bengaluru-	61800	61900	61400	66400	66500	65100
Urban						
Bagalkot	14200	14200	14200	16500	16600	16800
Ballari	43300	43400	43500	50300	51000	51500
Belgaum	18300	18400	18400	21100	21500	21600
Bengaluru-Rural	27900	27900	27900	28800	28900	28900
Bidar	14800	14800	14800	15100	15100	15100
Chamarajanagar	15400	15500	15500	17100	17300	17400
Chikkaballapur	28000	28000	28100	33400	33400	34200
Chikmagalur	14400	14400	14400	15700	15600	15800
Chitradurga	6600	6600	6600	7100	7100	7100
Dakshina-Kannada	32500	32600	32600	34500	34700	34800
Davanagere	12600	12700	12700	14800	15000	15200
Dharwad	22600	22600	22600	24600	24400	24700
Gadag	5100	5100	5100	5400	5400	5400
Hassan	29700	29700	29700	31700	31600	31900
Haveri	4400	4400	4400	5000	4800	5000
Kalaburagi	37400	37400	37500	38600	38600	38800
Kodagu	17100	17100	17100	17600	17500	17600
Kolar	26900	26900	27000	29200	29500	29600
Koppal	17700	17600	17800	21500	21300	22100
Mandya	36100	36200	36200	38400	38800	38800
Mysuru	76100	76400	76300	82700	83800	83600
Raichur	26600	26600	26700	29900	30100	30600
Ramanagar	11000	11000	11100	12700	12700	13000
Shivamogga	18700	18700	18700	20800	20900	21000
Tumakuru	47400	47400	47500	48800	49000	49200
Udupi	21300	21300	21300	22500	22500	22600
Uttara-Kannada	16400	16300	16400	18900	18800	19200
Vijayapura	17400	17500	17500	18700	18800	18900
Yadgir	13800	13700	13800	16600	16000	17000
Karnataka	1399200	1399800	1393300	1484600	1487900	1480300

Table 7: Comparison of unit-wise cumulative reported cases (since 01 November 2020) on 31 May 2021 and 31 August 2021 under 2/3-NPI with a daily budget of 167000 vaccine doses.

Table 8: The unit-wise percentage of susceptible population on 31 August 2021 with varying daily budgets of available vaccine doses under the 1/3-NPI, when vaccines are allocated in proportion to the population of each unit.

Unit	100K	133K	167K	200K	233K	267K	300K	333K	367K	400K
BBMP-	59.3	55.8	52.2	48.6	45.1	41.4	37.9	34.2	30.5	27.1
Bommanahalli										
BBMP-Dasarahalli	61.7	58.3	54.8	51.1	47.6	44.0	40.4	36.8	33.1	29.6
BBMP-East	60.6	57.1	53.4	49.9	46.3	42.5	38.9	35.4	31.6	28.1
BBMP-Mahadevpura	60.6	57.1	53.5	49.9	46.3	42.6	39.0	35.3	31.5	28.6
BBMP-RR-Nagar	55.2	51.5	47.7	44.1	40.5	36.7	32.9	29.6	27.5	26.1
BBMP-South	59.0	55.3	51.6	47.9	44.3	40.5	36.8	33.1	29.4	26.9
BBMP-West	56.3	52.7	49.0	45.3	41.5	37.8	34.1	30.5	28.1	26.7
BBMP-Yelahanka	60.0	56.4	52.9	49.3	45.8	42.1	38.5	35.0	31.2	28.3
Bagalkot	60.5	57.6	54.6	51.6	48.5	45.3	42.1	38.9	35.6	32.3
Ballari	47.2	44.2	41.1	37.9	34.8	31.5	28.3	25.0	22.1	20.1
Belgaum	56.8	53.9	51.0	48.0	45.0	41.8	38.6	35.5	32.1	28.9
Bengaluru-Rural	57.6	54.1	50.5	47.0	43.3	39.6	36.0	32.5	28.8	25.9
Rest of Bengaluru-	62.0	58.8	55.4	52.0	48.7	45.2	41.7	38.3	34.6	31.2
Urban										
Bidar	74.0	70.3	66.4	62.7	58.9	55.1	51.3	47.5	43.7	39.9
Chamarajanagar	45.0	41.8	38.5	35.3	32.0	28.6	25.3	22.3	20.6	19.5
Chikkaballapur	45.4	42.5	39.5	36.5	33.4	30.3	27.2	24.3	22.0	20.4
Chikmagalur	52.5	49.2	45.9	42.6	39.2	35.8	32.4	29.3	26.9	25.1
Chitradurga	64.8	61.6	58.3	55.0	51.7	48.3	44.9	41.5	37.9	34.5
Dakshina-Kannada	64.4	61.0	57.5	54.0	50.5	46.9	43.4	39.8	36.2	32.9
Davanagere	53.7	50.9	48.0	45.1	42.1	39.0	35.9	32.8	29.5	26.3
Dharwad	73.8	70.3	66.7	63.1	59.6	55.9	52.3	48.7	44.9	41.3
Gadag	78.2	74.5	70.6	66.9	63.2	59.3	55.5	51.8	47.9	44.1
Hassan	56.7	53.4	49.9	46.6	43.1	39.6	36.1	32.6	29.5	27.2
Haveri	66.3	63.3	60.1	57.0	53.8	50.5	47.2	43.9	40.4	37.1
Kalaburagi	67.7	64.1	60.4	56.7	53.1	49.4	45.7	42.0	38.2	34.6
Kodagu	46.4	43.1	39.4	35.9	32.4	29.2	26.6	24.8	23.3	22.2
Kolar	51.1	47.9	44.5	41.3	37.9	34.5	31.1	27.7	24.4	22.0
Koppal	48.2	45.5	42.5	39.7	36.7	33.5	30.5	27.4	24.3	21.9
Mandya	50.4	47.0	43.5	40.0	36.6	33.1	29.6	26.5	24.1	22.3
Mysuru	50.9	47.7	44.3	41.0	37.7	34.1	31.0	28.4	26.3	24.5
Raichur	43.2	40.0	36.7	33.5	30.2	26.8	24.1	22.0	20.3	19.3
Ramanagar	44.5	41.4	38.2	35.1	31.9	28.6	25.5	23.0	21.2	19.5
Shivamogga	61.6	58.6	55.3	52.1	48.9	45.5	42.1	38.8	35.3	32.1
Tumakuru	53.3	49.8	46.3	42.7	39.1	35.5	31.9	28.3	25.0	23.7
Udupi	61.9	58.5	54.9	51.4	47.9	44.2	40.7	37.1	33.9	31.3
Uttara-Kannada	53.4	50.6	47.6	44.6	41.6	38.4	35.3	32.1	29.0	26.7
Vijayapura	57.1	53.8	50.5	47.1	43.7	40.2	36.8	33.3	29.8	26.5
Yadgir	46.8	44.0	40.9	37.9	34.9	31.7	28.6	25.5	22.1	19.5
Karnataka	57.2	53.9	50.5	47.2	43.9	40.4	37.0	33.7	30.6	27.9

Table 9: Projected mean \pm StdDev of active cases in Karnataka under various NPIs and vaccination allocation strategies during the months of June, July and August 2021 when the daily vaccination budget is 167000.

		Populati	on prop	ortional	Inverse	seropr	evalence	Incidence	e rate	propor-
					proportional			tional	tional	
NPI		June	July	August	June	July	August	June	July	August
	Mean	144900	45900	14500	147500	47200	14900	144000	51800	20100
No NPI	Mean + StdDev	185000	62300	18600	187500	64100	19000	181100	67600	24700
	Mean - StdDev	104800	29500	10400	107500	30300	10800	106900	36000	15500
	Mean	72000	22100	7400	73400	23500	8600	71500	25200	10900
1/3 - NPI	Mean + StdDev	94900	29700	9100	96100	31200	10400	92900	32400	12600
	Mean - StdDev	49100	14500	5700	50700	15800	6800	50100	18000	9200
	Mean	40700	9400	2600	41400	10000	3000	40200	10700	3900
1/2 - NPI	Mean + StdDev	57600	13300	3300	58200	13900	3700	56200	14500	4600
	Mean - StdDev	23800	5500	1900	24600	6100	2300	24200	6900	3200
	Mean	18900	2400	400	19200	2500	400	18500	2600	500
2/3 - NPI	Mean + StdDev	30000	3800	500	30300	3900	600	29200	3900	700
	Mean - StdDev	7800	1000	300	8100	1100	200	7800	1300	300

4 Discussion and Limitations

We first discuss the implications of our results. We then discuss limitations of our study arising from model errors, uncertainty in the factors driving the resurgence in Karnataka, policy changes in response to the local circumstances, and vaccine supply constraints.

4.1 Discussion of the results

The significant second surge has highlighted widespread susceptibility in early 2021. This could be due to waning immunity, or limited spread of the first wave, or novel variants, or a combination thereof. An expedited, effective, and equitable vaccine campaign remains the only feasible pathway to controlling COVID-19 in India. Given supply constraints and India's population size, tools for designing efficient vaccination campaigns are essential. In this paper, we developed a model (using serosurvey data and on-the-ground datasets) to study vaccination allocation strategies and the interplay of vaccination capacity and NPIs in achieving sufficient immunity levels. The main messages of this work are the following:

- Heterogeneity across units under various NPI-scenarios: Our model suggests that there is heterogeneity in the disease spread across units, e.g. under No-NPI the units would have peaked at different times. The statewide uniform NPIs that are in place currently restrict mobility and thus limit the transmission of the virus within/across units. Our model predicts that all units would see a downward trend in reported cases after 15 May 2021. Daily reported cases have indeed come down in most units. Further, compared to No-NPI, on 31 August 2021, the model projects that 1/3-NPI may reduce the (cumulative reported) cases by roughly 926,000. Under 2/3-NPI and 167000 daily vaccinations, the percentage of susceptible populations in the units varies from 36.7% (Raichur) and 70.6% (Gadag), indicating significant heterogeneity across the units (see Table 8).
- Effectiveness of vaccination strategies: As for the three vaccination policies, our model indicates that they are roughly equal in the short-term, but in the longer term, statewide allocation in proportion to population leads to fewer cases. This conclusion is robust at the state level across various NPI scenarios. If we focus on units with high incidence in April 2021, e.g. Bengaluru's BBMP units, these units benefit more from an allocation in proportion to the case-incidence rate. However, all vaccination allocation strategies are within a 2% margin of each other. Given the robustness of the results, population-based allocation strategies are likely to be as successful as any other strategy that may be harder to implement.
- The interplay of vaccination and NPI on healthcare infrastructure capacity: With 167000 available vaccine doses per day, we find that the number of cumulative cases on 31 August 2021 reduces from 3.14 million under No-NPI to 1.48 million under 2/3-NPI. Table 3 and Figure 8 quantify the percentage reduction in the number of cases with respect to the reference 100000 vaccine doses/day and No-NPI setting. We have thus identified the key interplay between the intensity of the NPIs and the daily vaccination capacity. Further Table 9 provides the mean ± StdDev of active cases in Karnataka under various NPIs and vaccination allocation strategies during the months of June, July and August 2021. These can be used either at the unit level or at the state level to design a desirable public health response (choice of NPI, vaccination strategy, capacity). For example, if a mean of more than 60,000 active cases in a month would imply that the healthcare infrastructure in Karnataka is operating at full capacity, then an appropriate public health response starting May 2021 could be to vaccinate at a daily rate of 167000 and choose a social interaction between 1/3-NPI and 1/2-NPI. This will help to minimise deaths and manage severity of illness occurring during the surge.
- *Resurgence:* Our model predicts a resurgence in cases in the form of a third wave during the months of November-December 2021. This prediction is a conservative one since it is based on only seroreversion and mobility being restricted to within units. In reality, the peak could be advanced or delayed due to stochasticity and mobility and could be more prominent due to more transmissible variants.

Our model makes use of several real-time data sources – population data, age-distribution, age-stratified contact rates, disease progression data, confirmed case trajectories across units, seroprevalence data, time series of tests conducted, vaccination time-series, and efficacy of vaccines. Our compartmental ODE model used patches that where made up of age and unit stratifications. We modelled mobility across subunits to be independent of age. A future extension could be to use age-stratified mobility data.

There are many models in the Indian context, e.g., [3] with four compartments and [1] with five compartments that are explicitly modelling asymptomatic spread, [40] with nine compartments and explicit modelling of age-stratification and serology-based cases-to-infections ratio factor, and the PDE model in [25]. Our SEIRV spatio-temporal model goes beyond these and includes mobility and seroreversion.

One could also consider agent-based models as in [2], but given the complexity of such a model for Bengaluru alone (12.4 million population), scaling it up six times to the level of Karnataka will involve significant (more-than-linear) complexity increase. Parsimonious SEIRV-based district-level age-stratified ODE models incorporating mobility and seroreversion do provide a good tradeoff point between complexity and the ability to gather insights for public health policy.

Recognising that we already have several parameters in our model – five phases of contact rates and one contact rate per unit in each phase – we used only one constant contact rate per unit and scalings thereof (NPI intensities) for the forward projections. The per-unit contact rates are estimated based on the trajectory of the case during 08 April – 01 May 2021. Further, we quantified uncertainty by doing an estimation during 08–22 April 2021 and by doing a validation from 23 April 2021 – 11 May 2021. Statewide root mean squared estimation error was 9% and included the estimation errors of 40-50% for the units Chikkaballapur, Kodagu, Kolar, Vijaypura, and Raichur. The large errors in these units seem to be due a change in the trend of the number of reported cases which could not be captured by the model.

In terms of the vaccination policy, one could also employ simulation optimisation approaches to derive, algorithmically, vaccination schedules that minimise various epidemiologically relevant objectives under budget constraints as done in [32,46]. Such models use simulation in the loop and can incorporate a forecast horizon at which the vaccination policies are evaluated. When combined with data on vaccine acceptance [31], population profile of co-morbidities [14] and logistical challenges in setting up mass/mobile vaccination sites, one can obtain a realistic spatial distribution and allocation strategy that is executable, effective and equitable.

4.2 Limitations

Our first limitation is on the calibration of parameters – we have many contact rate parameters, one for each unit and for each of the five different time horizons. Tuning all of them to match the ground truth may lead to overfitting and loss of predictive power but still provides sufficient ground for scenario-based comparisons.

The selection of time points for contact rate calibration is based on observed regime changes for the trajectory of the case in each unit (see Appendix 4.2). A more systematic approach to arrive at these change points automatically can make the projections scalable to other states and the districts of India. Further, the change in transmissibility, if attributable to an increase in the prevalence of the variant strains (e.g., B.1.1.7 or B.1.617.2), can be modelled using estimates of transmissibility advantage [15].

Despite NPIs, infections may continue to rise for some time. The impact of such interventions will manifest only after some time if there are reporting delays. Our model used a delay of 1 week, assuming a two-day sample collection delay and 5-day test-reporting delay. More data is needed to quantify these delays. Another reason for the delayed impact of NPI could be household spread due to ineffective home isolation. Until the home infections naturally go down, these infections may continue to rise. Such effects are best modelled in agent-based simulators.

The exact cause of the resurgence is yet unclear. It could be due to antibody decay, parameters for which are not yet fully understood, and associated reinfections. It could also be due to reinfections arising from variants having modified spike protein configurations offering immune escape or a combination of these reasons. As the recent studies using the Manaus, Brazil outbreak show [9,39], disentangling these factors may be complex without additional data. Our approach to handling these factors is to move recovered individuals into the susceptible pool based on best available information on waning immunity and to increase the contact rates to capture some of the effects of novel variants; these explain the resurgence to some extent. A more rigorous but also a more complex approach would be to bring in more compartments that represent disease states associated with variants and to model variants and their transmissibility.

Our study is predicated on a constant daily vaccination capacity (which varied from 100000 - 400000). However, there are significant vaccine supply shortages and day-to-day variations. This shortage had become particularly acute starting 01 May 2021 when those above 18 years of age became eligible for vaccination. Further, some of the vaccination capacity is set aside for the prescribed second dose, an aspect which we have not modelled in this work. Finally, the current vaccination campaign does not utilise the serostatus of individuals as in [8], and so some proportion of the capacity is used up for those already immune. In our implementation, for simplicity, vaccinations are focused on the susceptibles and optimistically moves 66% of the daily vaccinated number into the vaccinated compartment after 14 days. Further study is required to account for vaccine wastage and immunisation of those already infected.

References

- [1] M. Agrawal, M. Kanitkar, and M. Vidyasagar. SUTRA: An approach to modelling pandemics with asymptomatic patients, and applications to COVID-19. arXiv preprint arXiv:2101.09158, 2021.
- [2] S. Agrawal, S. Bhandari, A. Bhattacharjee, A. Deo, N. M. Dixit, P. Harsha, S. Juneja, P. Kesarwani, A. K. Swamy, P. Patil, et al. City-scale agent-based simulators for the study of non-pharmaceutical interventions in the context of the COVID-19 epidemic. *Journal of the Indian Institute of Science*, pages 1–39, 2020.
- [3] S. Ansumali, S. Kaushal, A. Kumar, M. K. Prakash, and M. Vidyasagar. Modelling a pandemic with asymptomatic patients, impact of lockdown and herd immunity, with applications to SARS-CoV-2. *Annual Reviews in Control*, 2020.
- [4] G. R. Babu, R. Sundaresan, S. Athreya, J. Akhtar, P. K. Pandey, P. S. Maroor, M. Padma, R. Lalitha, M. Shariff, L. Krishnappa, et al. The burden of active infection and anti-SARS-CoV-2 IgG antibodies in the general population: Results from a statewide survey in Karnataka, India. *medRxiv*, 2020.
- [5] G. R. Babu, R. Sundaresan, S. Athreya, J. Akhtar, P. K. Pandey, P. S. Maroor, M. R. Padma, R. Lalitha, L. Krishnappa, M. Shariff, et al. The burden of active infection and anti-sars-cov-2 igg antibodies in the general population: Results from a statewide sentinel-based population survey in karnataka, india. *International Journal of Infectious Diseases*, 2021.
- [6] B. Biotech. Bharat Biotech announces phase 3 results of COVAXIN: Indias first COVID-19 vaccine demonstrates interim clinical efficacy of 81%. https://www.bharatbiotech.com/images/press/ covaxin-phase3-efficacy-results.pdf, 03 March 2021.
- [7] N. Brazeau, R. Verity, S. Jenks, H. Fu, C. Whittaker, P. Winskill, I. Dorigatti, P. Walker, S. Riley, R. P. Schnekenberg, et al. Report 34: COVID-19 infection fatality ratio: estimates from seroprevalence. *Imperial College, London, Technical Report*, 2020.
- [8] K. M. Bubar, K. Reinholt, S. M. Kissler, M. Lipsitch, S. Cobey, Y. H. Grad, and D. B. Larremore. Modelinformed COVID-19 vaccine prioritization strategies by age and serostatus. *Science*, 371(6532):916–921, 2021.
- [9] L. F. Buss, C. A. Prete, C. M. Abrahim, A. Mendrone, T. Salomon, C. de Almeida-Neto, R. F. França, M. C. Belotti, M. P. Carvalho, A. G. Costa, et al. Three-quarters attack rate of SARS-CoV-2 in the brazilian amazon during a largely unmitigated epidemic. *Science*, 371(6526):288–292, 2021.
- [10] F. M. Castonguay, J. C. Blackwood, E. Howerton, K. M. Shea, C. Sims, and J. N. Sanchirico. Spatial allocation of scarce vaccine and antivirals for COVID-19. *medRxiv*, 2020.
- [11] Centers for Disease Control and Prevention. COVID-19 pandemic planning scenarios. https://www.cdc.gov/coronavirus/2019-ncov/hcp/planning-scenarios-h.pdf [Online, accessed 02 May 2021.], 10 July 2020.
- [12] C. Cervia, J. Nilsson, Y. Zurbuchen, A. Valaperti, J. Schreiner, A. Wolfensberger, M. E. Raeber, S. Adamo, S. Weigang, M. Emmenegger, et al. Systemic and mucosal antibody responses specific to SARS-CoV-2 during mild versus severe COVID-19. *Journal of Allergy and Clinical Immunology*, 147(2):545–557, 2021.
- [13] J. Chen, S. Hoops, A. Marathe, H. Mortveit, B. Lewis, S. Venkatramanan, A. Haddadan, P. Bhattacharya, A. Adiga, A. Vullikanti, et al. Prioritizing allocation of COVID-19 vaccines based on social contacts increases vaccination effectiveness. *medRxiv*, 2021.
- [14] L. Dandona, R. Dandona, G. A. Kumar, D. Shukla, V. K. Paul, K. Balakrishnan, D. Prabhakaran, N. Tandon, S. Salvi, A. Dash, et al. Nations within a nation: variations in epidemiological transition across the states of India, 1990–2016 in the global burden of disease study. *The Lancet*, 390(10111):2437– 2460, 2017.
- [15] N. G. Davies, S. Abbott, R. C. Barnard, C. I. Jarvis, A. J. Kucharski, J. D. Munday, C. A. Pearson, T. W. Russell, D. C. Tully, A. D. Washburne, et al. Estimated transmissibility and impact of SARS-CoV-2 lineage B.1.1.7 in England. *Science*, 372(6538), 2021.

- [16] Department of Health and Family Welfare Services. COVID-19 Media Bulletin, Government of Karnataka. https://covid19.karnataka.gov.in/govt_bulletin/en.
- [17] Department of Health and Family Welfare Services. Novel Coronavirus (COVID-19) Media Bulletin. https://covid19.karnataka.gov.in/storage/pdf-files/EMB-APR21/22-04-2021HMBEnglish. pdf, 22 April 2021.
- [18] Department of Health and Family Welfare Services. Novel Coronavirus (COVID-19) Media Bulletin, Department of Health and Family Welfare Services. https://covid19.karnataka.gov.in/pdfs/mar21/ en/02-03-2021HMBEnglish.pdf, 02 March 2021.
- [19] Department of Health and Family Welfare Services. Novel Coronavirus (COVID-19) Media Bulletin, Department of Health and Family Welfare Services. https://covid19.karnataka.gov.in/storage/ pdf-files/EMB-APR21/02-04-2021HMBEnglish.pdf, 02 April 2021.
- [20] Department of Health and Family Welfare Services. Order To re-start COVID-19 Apthamitra-TeleMedicine the Services for the control of (in Kanhttps://covid19.karnataka.gov.in/storage/pdf-files/GovernmentOrders/ nada). Order-Tore-starttheApthamitra-TeleMedicineServicesforthecontrolofCOVID-19.pdf, 08 April 2021.
- [21] Directorate of Economics and Statistics, Bangalore. Projected Population of Karnataka 2012-2021 (Provisional), DES 22 of 2013. https://des.kar.nic.in/docs/ProjectedPopulation2012-2021. pdf, 18 February 2013.
- [22] M. Dogan, L. Kozhaya, L. Placek, C. Gunter, M. Yigit, R. Hardy, M. Plassmeyer, P. Coatney, K. Lillard, Z. Bukhari, et al. SARS-CoV-2 specific antibody and neutralization assays reveal the wide range of the humoral immune response to virus. *Communications Biology*, 4(1):1–13, 2021.
- [23] N. Ferguson, D. Laydon, G. Nedjati Gilani, N. Imai, K. Ainslie, M. Baguelin, S. Bhatia, A. Boonyasiri, Z. Cucunuba Perez, G. Cuomo-Dannenburg, et al. Report 9: Impact of non-pharmaceutical interventions (NPIs) to reduce COVID19 mortality and healthcare demand. *Tech. Report*, 2020.
- [24] B. H. Foy, B. Wahl, K. Mehta, A. Shet, G. I. Menon, and C. Britto. Comparing COVID-19 vaccine allocation strategies in India: A mathematical modelling study. *International Journal of Infectious Diseases*, 103:431–438, 2021.
- [25] S. Ganesan and D. Subramani. Spatio-temporal predictive modeling framework for infectious disease spread. *Scientific Reports*, 11(1):1–8, 2021.
- [26] Government of Karnataka. Guidelines to break the chain of COVID-19 transmission in the state. https://covid19.karnataka.gov.in/storage/pdf-files/Government%200rders/Order-Revised% 20Guidelines%20to%20break%20the%20Chain%20of%20C0VID-19%20Transmission%20(extension) %20in%20the%20State.pdf, 26 April 2021.
- [27] S. L. Klein, A. Pekosz, H.-S. Park, R. L. Ursin, J. R. Shapiro, S. E. Benner, K. Littlefield, S. Kumar, H. M. Naik, M. J. Betenbaugh, et al. Sex, age, and hospitalization drive antibody responses in a COVID-19 convalescent plasma donor population. *The Journal of Clinical Investigation*, 130(11):6141– 6150, 2020.
- [28] N. Kumar, S. K. S. Hameed, G. R. Babu, M. M. Venkataswamy, P. Dinesh, P. K. Bg, D. A. John, A. Desai, and V. Ravi. Descriptive epidemiology of SARS-CoV-2 infection in Karnataka state, South India: Transmission dynamics of symptomatic vs. asymptomatic infections. *EClinicalMedicine*, 32:100717, 2021.
- [29] V. M. Kumar, S. R. Pandi-Perumal, I. Trakht, and S. P. Thyagarajan. Strategy for COVID-19 vaccination in India: the country with the second highest population and number of cases. *npj Vaccines*, 6(1):1–7, 2021.
- [30] E. H. Lau, O. T. Tsang, D. S. Hui, M. Y. Kwan, W.-h. Chan, S. S. Chiu, R. L. Ko, K. H. Chan, S. M. Cheng, R. A. Perera, et al. Neutralizing antibody titres in SARS-CoV-2 infections. *Nature Communications*, 12(1):1–7, 2021.

- [31] J. V. Lazarus, S. C. Ratzan, A. Palayew, L. O. Gostin, H. J. Larson, K. Rabin, S. Kimball, and A. El-Mohandes. A global survey of potential acceptance of a COVID-19 vaccine. *Nature Medicine*, 27(2):225–228, 2021.
- [32] J. C. Lemaitre, D. Pasetto, M. Zanon, E. Bertuzzo, L. Mari, S. Miccoli, R. Casagrandi, M. Gatto, and A. Rinaldo. Optimizing the spatio-temporal allocation of COVID-19 vaccines: Italy as a case study. *medRxiv*, 2021.
- [33] Q.-X. Long, B.-Z. Liu, H.-J. Deng, G.-C. Wu, K. Deng, Y.-K. Chen, P. Liao, J.-F. Qiu, Y. Lin, X.-F. Cai, et al. Antibody responses to SARS-CoV-2 in patients with COVID-19. *Nature Medicine*, 26(6):845–848, 2020.
- [34] C. McAloon, A. Collins, K. Hunt, A. Barber, A. W. Byrne, F. Butler, M. Casey, J. Griffin, E. Lane, D. McEvoy, et al. Incubation period of COVID-19: a rapid systematic review and meta-analysis of observational research. *BMJ Open*, 10(8):e039652, 2020.
- [35] Ministry of Health and Family Welfare Services, Government of India. Liberalised pricing and accelerated national COVID-19 vaccination strategy. https://www.mohfw.gov.in/pdf/ LiberalisedPricingandAcceleratedNationalCovid19VaccinationStrategy2042021.pdf, 21 April 2021.
- [36] F. Muecksch, H. Wise, B. Batchelor, M. Squires, E. Semple, C. Richardson, J. McGuire, S. Clearly, E. Furrie, N. Greig, et al. Longitudinal serological analysis and neutralizing antibody levels in coronavirus disease 2019 convalescent patients. *The Journal of Infectious Diseases*, 223(3):389–398, 2021.
- [37] Office of the Registrar General and Census Commissioner, India. Age distribution of states in India. https://censusIndia.gov.in/vital_statistics/SRS_Report_2017/12. SRSStatisticalReport-Detailedtables-2017.pdf, 2017.
- [38] K. Prem, A. R. Cook, and M. Jit. Projecting social contact matrices in 152 countries using contact surveys and demographic data. *PLoS Computational Biology*, 13(9):e1005697, 2017.
- [39] E. C. Sabino, L. F. Buss, M. P. Carvalho, C. A. Prete, M. A. Crispim, N. A. Fraiji, R. H. Pereira, K. V. Parag, P. da Silva Peixoto, M. U. Kraemer, et al. Resurgence of COVID-19 in manaus, brazil, despite high seroprevalence. *The Lancet*, 397(10273):452–455, 2021.
- [40] S. Shekatkar, B. Pujari, M. Arjunwadkar, D. K. Hazra, P. Chaudhuri, S. Sinha, G. I. Menon, A. Sharma, and V. Guttal. INDSCI-SIM a state-level epidemiological model for India, 2020. Ongoing Study at https://indscicov.in/indscisim.
- [41] L. Singh and T. Barnagarwala. Mumbai third sero survey: 36% have Covid antibodies. https://Indianexpress.com/article/cities/mumbai/ mumbai-third-sero-survey-36-have-covid-antibodies-7287791/, 25 April 2021.
- [42] Siva Athreya, Nitya Gadhiwala, Abhiti Mishra, Srigyan Nandi, and Srivatsa B. COVID-19 India Timeline: An Understanding across States and Union Territories. https://www.isibang.ac.in/~athreya/ incovid19/data.php, 22 April 2021.
- [43] Siva Athreya, Nitya Gadhiwala, Abhiti Mishra, Srigyan Nandi, and Srivatsa B. . One week predictions for districts of Karnataka. https://www.isibang.ac.in/~athreya/incovid19/wp, 23 April 2021.
- [44] R. Thiruvengadam, S. Chattopadhyay, F. Mehdi, B. K. Desiraju, S. Chaudhuri, S. Singh, V. Bhartia, P. Kshetrapal, U. C. M. Natchu, N. Wadhwa, et al. Longitudinal serology in SARS-CoV-2 infected individuals in India–a prospective cohort study. *medRxiv*, 2021.
- [45] S. Venkatramanan, P. Bhattacharya, P. Porebski, and B. Klahn. NSSAC/PatchSim: First official release. https://doi.org/10.5281/zenodo.4313095, December 2020.
- [46] S. Venkatramanan, J. Chen, A. Fadikar, S. Gupta, D. Higdon, B. Lewis, M. Marathe, H. Mortveit, and A. Vullikanti. Optimizing spatial allocation of seasonal influenza vaccine under temporal constraints. *PLoS Computational Biology*, 15(9):e1007111, 2019.

- [47] M. Voysey, S. A. C. Clemens, S. A. Madhi, L. Y. Weckx, P. M. Folegatti, P. K. Aley, B. Angus, V. L. Baillie, S. L. Barnabas, Q. E. Bhorat, et al. Single-dose administration and the influence of the timing of the booster dose on immunogenicity and efficacy of ChAdOx1 nCoV-19 (AZD1222) vaccine: a pooled analysis of four randomised trials. *The Lancet*, 397(10277):881–891, 2021.
- [48] A. Wajnberg, M. Mansour, E. Leven, N. M. Bouvier, G. Patel, A. Firpo-Betancourt, R. Mendu, J. Jhang, S. Arinsburg, M. Gitman, et al. Humoral response and PCR positivity in patients with COVID-19 in the New York City region, USA: An observational study. *The Lancet Microbe*, 1(7):e283–e289, 2020.

Appendix

More on the calibration and the piece-wise constant contact rate periods

Our entire model is a spatio-temporal age-stratified SEIRV compartmental model with mobility and antibody waning. To determine the periods when the contact rates can be held constant, let us first turn to a simplified model to make some insightful observations.

Decay and growth phases: Let us consider a simplified setting where we ignore mobility, age-stratified interaction, vaccination and antibody waning. Focus on fractions of the population in each compartment and write $s_j(t) = S_j(t)/N_j$ for the susceptible fraction in unit j at time t; similarly define $e_j(t)$, $i_j(t)$, $r_j(t)$ in unit j at time t. Then the difference equations in (1)-(4) separate for each unit j. Let us suppress the unit index j in the notation for this section. The difference equations constitute the Euler discretisation for the ordinary differential equation system:

$$\begin{aligned} \frac{de}{dt} &= \beta(t)s(t)i(t) - \alpha e(t) \\ \frac{ds}{dt} &= -\beta(t)s(t)i(t) \\ \frac{di}{dt} &= \alpha e(t) - \gamma i(t) \\ \frac{dr}{dt} &= \gamma i(t). \end{aligned}$$

Assuming $\beta(t)$ and s(t) are constant⁷, setting $\beta(t)s(t) \equiv \overline{\beta}$, we have the simpler linear system

$$\dot{x}(t) = Ax(t)$$

where

$$x(t) = \begin{bmatrix} e \\ i \end{bmatrix}$$
 and $A = \begin{bmatrix} -lpha & \overline{eta} \\ lpha & -\gamma \end{bmatrix}$.

It is easy to verify that A is diagonalisable for generic values of $\alpha, \overline{\beta}, \gamma$, and that its two eigenvalues are

$$\lambda_1 = \frac{-(\alpha + \gamma) + \sqrt{(\alpha + \gamma)^2 - 4\alpha(\gamma - \overline{\beta})}}{2} \quad \text{and} \quad \lambda_2 = \frac{-(\alpha + \gamma) - \sqrt{(\alpha + \gamma)^2 - 4\alpha(\gamma - \overline{\beta})}}{2}$$

Then $x(t) = e^{At}x(0)$ and hence i(t) (and also e(t)) will grow or will decay exponentially as $i(t) \sim Ce^{\lambda_1 t}$, where λ_1 is the larger of the two eigenvalues. This suggests that λ_1 may be estimated as the slope using linear regression on the trajectory $\log(i(t))$ during the phase when $\overline{\beta} \equiv \beta(t)s(t)$ is constant. Using standard methods in least squares regression, a 95%- confidence interval for the estimate of λ_1 can be obtained using Student's t-distribution, in this simplified model.

From the plot in Figure 10, we see that the $\overline{\beta}$ is roughly constant during November 2020 to February 2021, and then from mid-March 2021. These motivate the calibration periods used in this work.

The quantity $\overline{\beta}$ can be recovered from the eigenvalue as:

$$\overline{\beta} = \gamma \left(1 + \lambda_1 / \gamma \right) \left(1 + \lambda_1 / \alpha \right).$$

Observe that the reproduction number for this oversimplified model is $R_t = \overline{\beta}/\gamma$, which is given by $R_t = (1 + \lambda_1/\gamma) (1 + \lambda_1/\alpha)$. Clearly $R_t > 1$ if $\lambda_1 > 0$ and $R_t < 1$ if $\lambda_1 < 0$, as expected.

These observations suggest a least-squares regression to estimate λ_1 and then an inversion using the R_t formula to obtain $\overline{\beta}$. Further, a linear approximation can be used to obtain confidence intervals for $\overline{\beta}$ from that of λ_1 , in this simplified model.

The actual implementation is a nonlinear least-squares regression that directly searches for the β . We take the loss function to be minimised as the squared error objective $\sum_t (\log i(t) - \log \hat{i}_{\beta}(t))^2$ where $\hat{i}_{\beta}(t)$ is the model-predicted infections at time t for contact rate β , and the sum is taken over the period for which $\beta(\cdot)$ is assumed constant (called a phase). Note that $\hat{i}_{\beta}(\cdot)$ is a β -dependent solution to the difference equations (1)-(4) and respects the actual number of susceptible individuals at each time instant. We implement an iterative gradient descent algorithm to arrive at a good β for each unit. Further, since (1)-(4) is a coupled

⁷During the early exponential growth phase, the contact rate $\beta(t)$ being static seems like a fair assumption. Further, during this early exponential growth phase, the susceptible fraction is likely quite large and the s(t) changes at a slower time-scale.

system of patches (units with age-stratification), in each phase, for each iteration, we update the β_j in a round-robin fashion across the units.

Calibration before resurgence: Between 01–31 March 2021, we noted in our simulations that while there were residual infections in the Bengaluru units, there was a need to seed infections in some other units to match the timing of the resurgence in these units. The resurgences in these units were likely due to seeding by mobility from other places where the infection was still prevalent. So we model a cross-unit mobility $\Theta(t) = (1 - \varepsilon)I + \epsilon J$ for the period of March 2021 with $\varepsilon = 1/100$. Once seeded, we conservatively model only local transmission, and so we remove the cross-unit mobility from 01 April 2021 onward.

There is also significant stochasticity during 01–31 March 2021. Mean-field models like ours need to be enhanced to handle this stochasticity during such phases. To compensate for the stochasticity, we divide the period into two phases, where we estimate the contact rates. However, the estimation/calibration is only to put the system in a realistic initial state (how many in each of the SEIRV compartments) to learn the best-fit contact rates during the exponential growth phase starting 01 April 2021. Some units see this exponential phase from 15 March 2021 itself, so splitting the month of March into two phases allows some units to fit better the early upsurge (e.g., BBMP units). In summary, the two calibrations during this period is largely to seed infections through mobility, handle stochasticity, and put the system in a more realistic initial state prior to the exponential growth phase starting 01 April 2021.

Contact rates

Table 10: Calibrated contact rates for each unit. The time periods T_1, \ldots, T_5 are 11–31 October 2020, 01 November 2020 – 28 February 2021, 01–15 March 2021, 16 March – 07 April 2021, 08 April onward, respectively.

Unit	T_1	T_2	T_3	T_4	T ₅
BBMP-Bommanahalli	0.018	0.025	0.074	0.04	0.047
BBMP-Dasarahalli	0.016	0.022	0.056	0.038	0.042
BBMP-East	0.041	0.022	0.06	0.043	0.047
BBMP-Mahadevpura	0.013	0.021	0.062	0.042	0.043
BBMP-RR-Nagar	0.019	0.023	0.092	0.043	0.062
BBMP-South	0.033	0.026	0.084	0.048	0.058
BBMP-West	0.032	0.029	0.08	0.047	0.062
BBMP-Yelahanka	0.019	0.022	0.057	0.039	0.043
Bagalkot	0.006	0.017	0.018	0.008	0.043
Ballari	0.01	0.025	0.057	0.015	0.056
Belgaum	0.013	0.025	0.032	0.012	0.046
Bengaluru-Rural	0.019	0.025	0.061	0.046	0.049
Rest of Bengaluru-Urban	0.018	0.024	0.045	0.03	0.042
Bidar	0.019	0.025	0.045	0.045	0.029
Chamarajanagar	0.02	0.022	0.033	0.018	0.064
Chikkaballapur	0.014	0.021	0.022	0.011	0.056
Chikmagalur	0.014	0.024	0.031	0.031	0.048
Chitradurga	0.021	0.024	0.039	0.017	0.04
Dakshina-Kannada	0.014	0.025	0.054	0.029	0.039
Davanagere	0.014	0.028	0.019	0.013	0.048
Dharwad	0.01	0.02	0.021	0.012	0.035
Gadag	0.007	0.016	0.011	0.005	0.031
Hassan	0.017	0.023	0.047	0.035	0.045
Haveri	0.01	0.019	0.01	0.009	0.041
Kalaburagi	0.008	0.026	0.06	0.042	0.037
Kodagu	0.021	0.024	0.045	0.025	0.078
Kolar	0.017	0.021	0.045	0.026	0.055
Koppal	0.015	0.016	0.007	0.01	0.052
Mandya	0.023	0.021	0.036	0.036	0.055
Mysuru	0.017	0.024	0.047	0.026	0.05
Raichur	0.01	0.022	0.017	0.017	0.063
Ramanagar	0.011	0.02	0.031	0.016	0.058
Shivamogga	0.009	0.021	0.021	0.019	0.041
Tumakuru	0.017	0.024	0.08	0.034	0.059
Udupi	0.011	0.025	0.07	0.034	0.039
Uttara-Kannada	0.012	0.021	0.027	0.013	0.047
Vijayapura	0.025	0.026	0.051	0.033	0.046
Yadgir	0.007	0.022	0.032	0.011	0.057

Seroprevalence from the model

Table 11: Seroprevalence from the model and the data. Once the second round serosurvey data is available (both active infection and antibody prevalence estimates), the last two columns could be used for comparison and calibration. Number are in percentages.

Unit	Active Infection %	Antibody(IgG)	Antibody (IgG) Preva-
	on 18 February 2021	Prevalence % on	lence % (Round-1)
	(Model)	18 February 2021	
		(Model)	
BBMP-Bommanahalli	0.039	12.359	12.5
BBMP-Dasarahalli	0.045	7.987	13.8
BBMP-East	0.068	9.906	15.5
BBMP-Mahadevpura	0.044	7.176	7
BBMP-RR-Nagar	0.048	12.143	9.1
BBMP-South	0.059	14.208	20.9
BBMP-West	0.072	15.53	25.1
BBMP-Yelahanka	0.05	8.404	13.9
Bagalkot	0.004	3.987	4.1
Ballari	0.01	10.858	22.1
Belgaum	0.04	7.84	23.7
Bengaluru-Rural	0.029	13.002	15.2
Rest of Bengaluru-	0.046	10.825	15
Urban			
Bidar	0.07	4.672	18
Chamarajanagar	0.029	8.953	15.8
Chikkaballapur	0.045	6.708	6.4
Chikmagalur	0.02	12.011	12
Chitradurga	0.044	12.621	10.2
Dakshina-Kannada	0.046	10.419	14.5
Davanagere	0.019	13.527	16.4
Dharwad	0.017	2.673	7.1
Gadag	0.001	2.101	6.3
Hassan	0.025	12.012	8.2
Haveri	0.005	8.062	14.8
Kalaburagi	0.053	8.975	17.1
Kodagu	0.072	12.311	12
Kolar	0.034	8.17	10.1
Koppal	0.002	4.655	19.6
Mandya	0.014	9.631	18.5
Mysuru	0.05	9.665	18.8
Raichur	0.007	8.097	22.8
Ramanagar	0.006	8.735	13.9
Shivamogga	0.023	7.514	7.7
Tumakuru	0.045	12.424	6.8
Udupi	0.018	11.31	16.2
Uttara-Kannada	0.031	7.077	8.1
Vijayapura	0.019	13.731	23.9
Yadgir	0.01	8.551	15.4



Figure 10: Log plot of the daily infections (averaged over seven days to avoid weekly periodicity issues). The different phases are colour coded. During October 2020, we initialise the simulator. During 01 November 2020 – 28 February 2021, we capture the receding trend. As seen above, both are well-captured by straight lines. During 01 March 2021 – 31 March 2021, we calibrate to re-seed the infections across the units and handle stochasticity. From 01 April 2021 - 01 May 2021, we capture the resurgence and the λ_1 associated with it. All projections are from 02 May 2021 onward.

Time series of daily and cumulative new infections





(h) BBMP-Mahadevpura

31 Dec 2021

31 Dec 2021

31 Dec 2021

31 Dec 2021









Rest of Bengaluru-Urban -- Cumulative













1600

1400

1200 ں۔ 1000 sess م

400

200 D

Daily 600

Figure 11: Daily predicted number of cases in each unit until 31 December 2021 for various NPI intensities. Vaccination capacity is 167000 per day allocated in proportion to population in the units.

No NPI

1/3 NPI 1/2 NPI

2/3 NPI

2021

No NPI

1/3 NPI

1/2 NPI 2/3 NPI

31 Dec 2021

No NPI

1/3 NPI

1/2 NPI

2/3 NPI

31 Dec 2021

No NPI

1/3 NPI

1/2 NPI

2/3 NPI

01 Nov 2021

01 Nov 2021 01 Dec 2021 31 Dec 2021

01 Dec 2021

01 Nov 2021

.

01 Nov 2021

01 Dec 202:

Susceptible fraction

No NPI

1/3 NPI

1/2 NPI

2/3 NPI

31 Dec 2021

No NPI

1/3 NPI

1/2 NPI

2/3 NPI

No NPI

1/3 NPI

1/2 NPI

2/3 NPI

No NPI 1/3 NPI

1/2 NPI

2/3 NPI

31 Dec 2021

01 Dec 2021

.

01 Dec 2021

No NPI

1/3 NPI

1/2 NPI

2/3 NPI

31 Dec 2021

No NPI

1/3 NPI

1/2 NPI

2/3 NPI

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01 Nov 2021

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01 Dec 2021

01 Oct 2021

01 Oct 2021 01 Nov 2021 01 Dec 2021 31 Dec 2021

> . No NPI

1/3 NPI

1/2 NPI

2/3 NPI

31 Dec 2021

No NPI 1/3 NPI

1/2 NPI

2/3 NPI

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01 Aug 202 01 Sep 202:

01 Aug 2021

01 Aug 2021 01 Sep 2021 01 Oct 2021 01 Nov 2021 01 Dec 2021 31 Dec 2021

01 Sep 2021

2021

01 Aug 2

Figure 12: Susceptible fractions in each unit for various NPI intensities. Vaccination capacity is 167000 per day allocated in proportion to population in the units.

2021

22 Apr

2021

22 Apr.

2021

22 Apr 3

2021

22 Apr 2

01 May 2021

01 May 2021

01 May 2021

01 May 2021

11 May 2021

11 May 2021

11 May 2021

11 May 2021

Carton State

Validation Plots

Figure 13: Validation plots for each unit. Calibration has been performed from 08–22 April 2021 and validated during 23 April – 11 May 2021. The vertical line separates the calibration and the validation phases.