



Sub-national Level Analysis of Measles Prevalence in India

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Motivation

- India is committed to eliminate measles by December 2023
- Measles outbreak in India (November, December 2022)
 - Mumbai and other parts of India
 - Highly concentrated
- Analysis of measles prevalence at subnational level

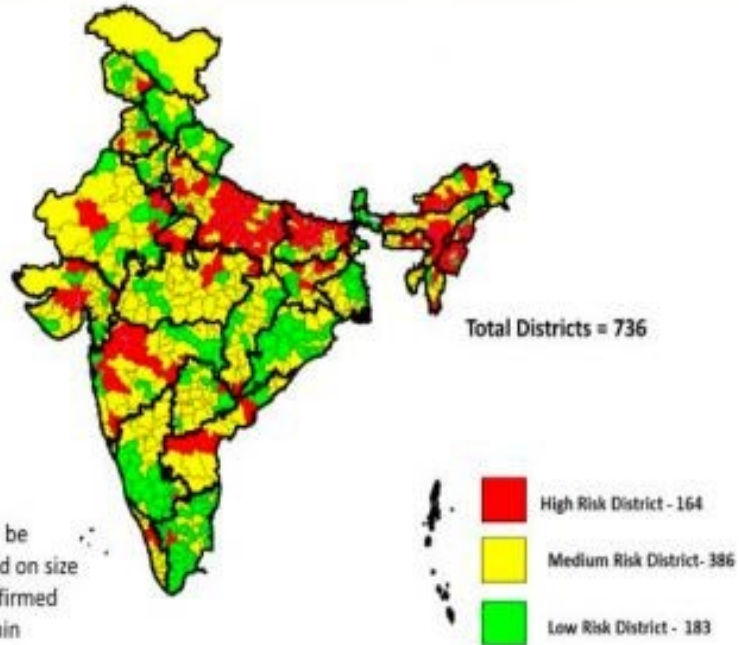


Objectives

- Modelling the measles burden at subnational level.
- Analysis of supplemental immunization activities (SIA)



MR Risk Assessment, India



From the figure we observe that the most **high-risk districts** were located in **Uttar Pradesh (41)**, followed by **Bihar (19)**, **Arunachal Pradesh (14)**, **Maharashtra (12)** & **Manipur (12)**

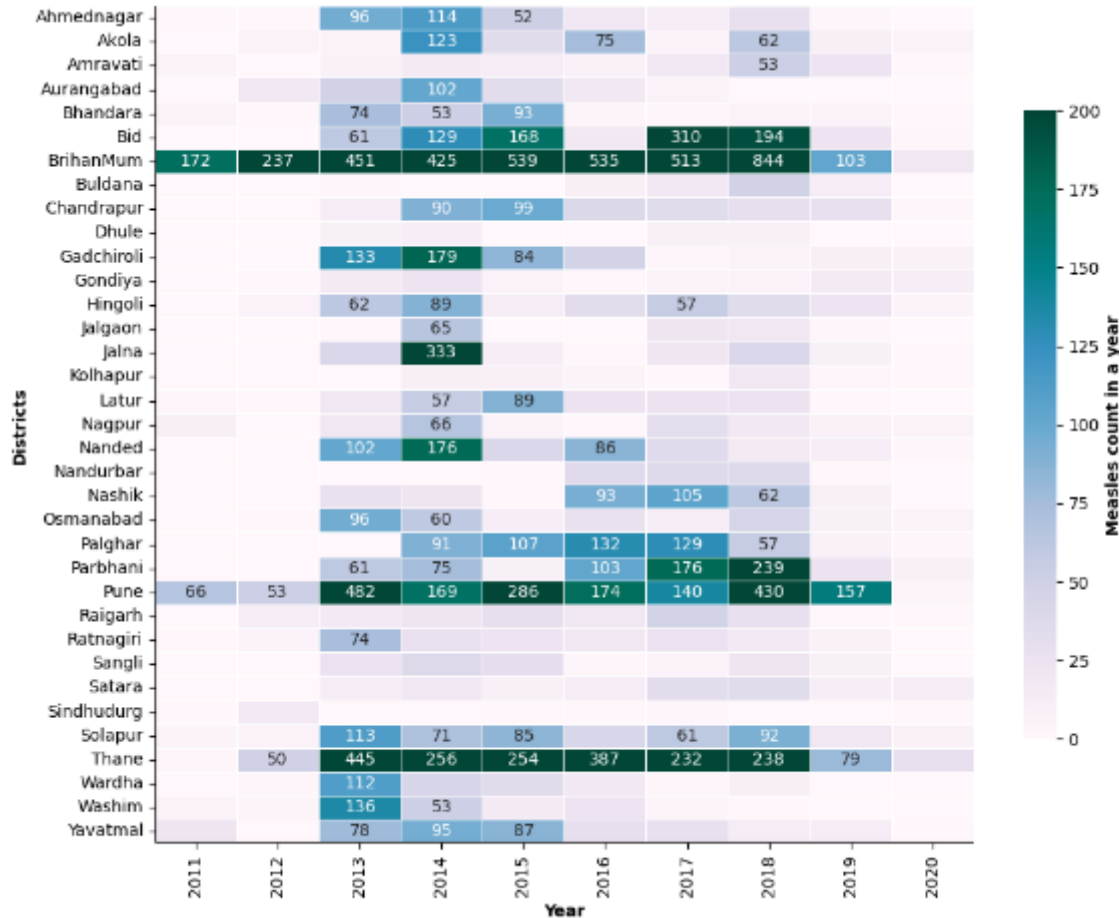
Source: Roadmap to Measles and Rubella Elimination in India by 2023



Data Sources

- **HMIS - Monthly data (April 2013 - March 2020)**
 - Data on child immunization 1st and 2nd dose
 - Live births
 - Measles disease count in children below 5
 - Child deaths with probable cause being measles (1-11 months and 1-5yrs)
- **NFHS-5 - Districts Fact Sheets**
 - Data on MCV-1 for the years 2019-20, 2015-16
 - Data on MCV-2 for the year 2019-20
- **Census of India, 2011**
 - District level population age-wise

Identifying the High Risk Districts of Maharashtra



According to the heat map our districts of focus should be **Mumbai**, **Pune** and **Thane**



Methods

- Time series SIR model: Grenfell *et al.* (2002) and Thakkar *et al.* (2019)
 - Fit the model for three high risk districts of Maharashtra: Mumbai, Pune and Thane
 - Based on the fitted models predict the effect of future SIAs on estimated susceptible population and measles infections
- State space model: Chen, Fricks and Ferrari (2012)
 - Fit the model for Mumbai



TSIR Model

- Following Grenfell *et al.* (2002) and Thakkar *et al.* (2019), we model S_t and I_t as a discrete stochastic dynamical system
 - $S_t = (1 - \mu_{t-1})(B_t + S_{t-1} + I_t)$
 - $I_t = \beta_t I_{t-1}^\alpha S_{t-1} \varepsilon_t$
 - $C_t \sim \text{Binom}\{I_t, p\}$
- β_t - unknown transmission rate



TSIR Modelling

- $B_t = \bar{B}_t [1 - 0.9V_{1,t}(1 - V_{2,t}) - 0.99V_{1,t}V_{2,t}]$
- NFHS data - MCV1 & MCV2 coverages
- Monthly fluctuations in $V_{1,t}$ and $V_{2,t}$ from HMIS
- TSIR Model has been fitted using monthly data from April'13-March'20 (84 time points for each district).
- SIA at 2018 December with 60% efficacy is incorporated.



TSIR Model Calibration

- In this method we have utilized the live birth counts for the next 20 years (Apr'20 to Mar'40) to calculate the adjusted birth counts from the National Population Projection Report published by the Government of India
- We used Crude Birth Rate (CBR), Population growth rate and total Population of respective districts reported in the Government report to calculate the live births.
- To Calculate the adjusted live births we have used MCV1 coverage as 75% and MCV2 coverage as 20% (constant throughout the time period) and the above obtained live birth counts.

State Space Model



- Chen, Fricks and Ferrari (2012)
- Observation equation: $C_t = \theta_2 \{1 - \exp(-\theta_1 S_{t-1}/N_{t-1})\} S_{t-1} + \varepsilon_t$
- System equation: $S_t = S_{t-1} - \left[\left\{ 1 - \exp\left(-\frac{\theta_1 S_{t-1}}{N_{t-1}}\right) \right\} S_{t-1} + X_t \right] (1 - Y_{t-1}) + \eta_t$
- Where $\varepsilon_t \sim iid N(0, \theta_3^2)$ and $\eta_t \sim iid N(0, \theta_4^2)$
- X_t : adjusted birth rate accounting for MCV1 and MCV2 coverage
- Y_t : Coverage of SIA at time t
- N_t : Total population at time t
- $\theta_1 - \theta_4$: unknown parameters



State Space Model Calibration

- The number of susceptible individuals (S_t) is treated as unobservable Markovian process while the case counts (C_t) are treated as observed time series.
- Both C_t and S_t are modelled simultaneously via non-linear time series regression models with covariates as population, at risk birth cohort and outreach of vaccination campaigns.
- We use extended Kalman filter to estimate C_t and S_t .



Findings for Mumbai: RMSE and MAPE

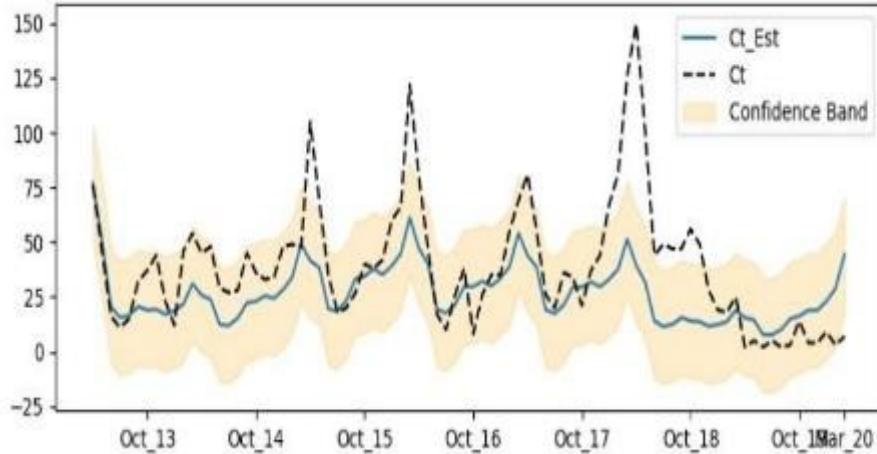
The root-mean-squared error (RMSE) and mean absolute prediction error (MAPE) in the estimation of C_t are given below.

Method	RMSE	MAPE
TSIR	25.9300	0.8583
State Space	21.1627	0.8318

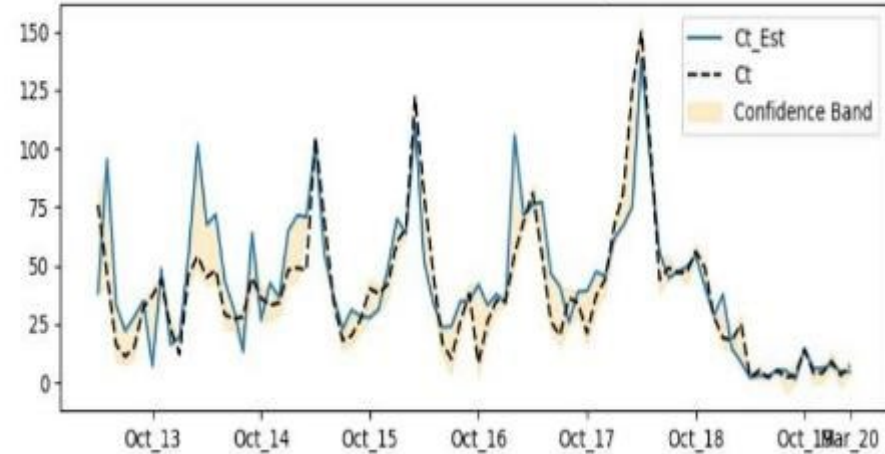


Findings for Mumbai : Estimated Case Counts

Actual and Estimated Ct: TSIR



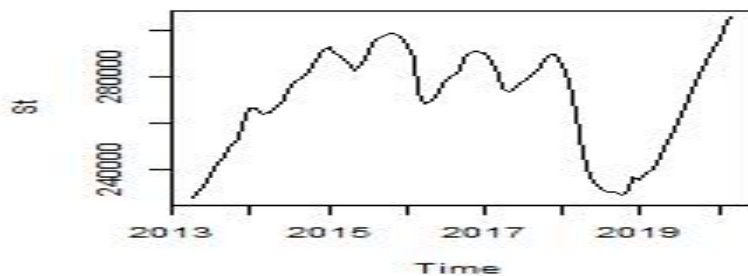
Actual and Estimated Ct: State Space



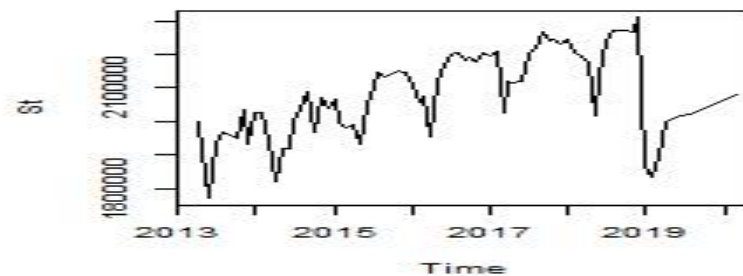


Findings for Mumbai : Estimated Susceptibility

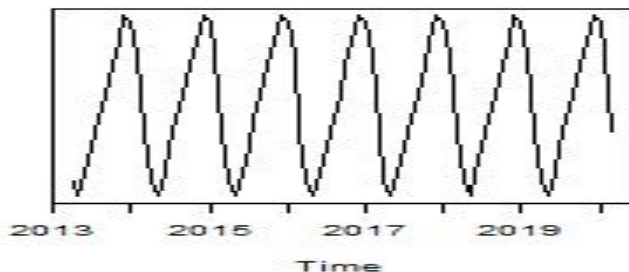
Estimated Susceptibility: TSIR



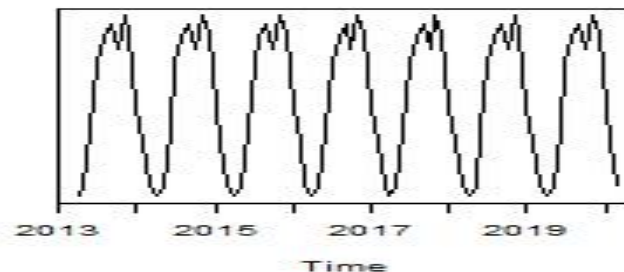
Estimated Susceptibility: State Space



Estimated seasonality in St: TSIR



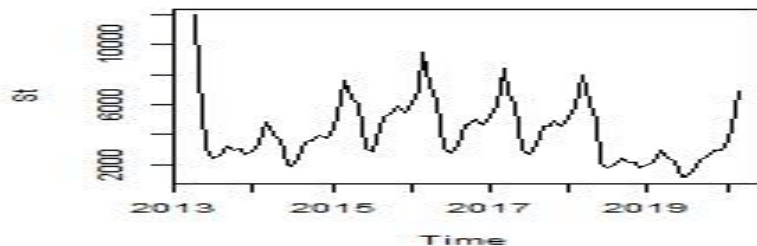
Estimated seasonality in St: State Space



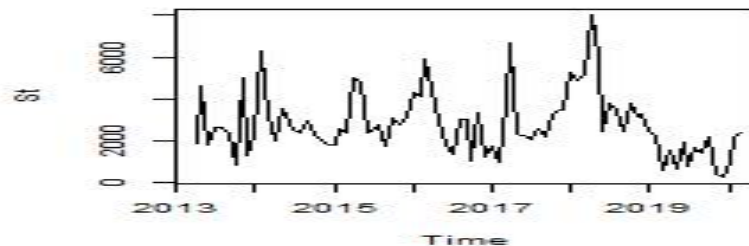


Findings for Mumbai : Estimated Infectious Population

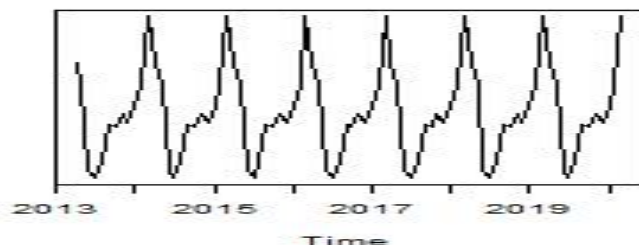
Estimated Infectious: TSIR



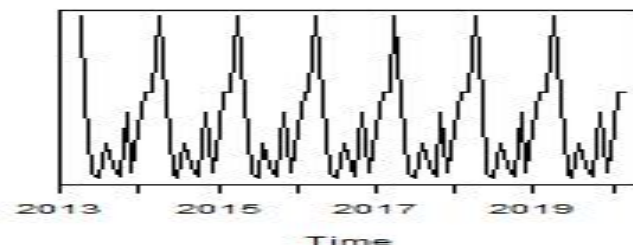
Estimated Infectious: State Space



Estimated seasonality in I_t: TSIR



Estimated seasonality in I_t: State Space

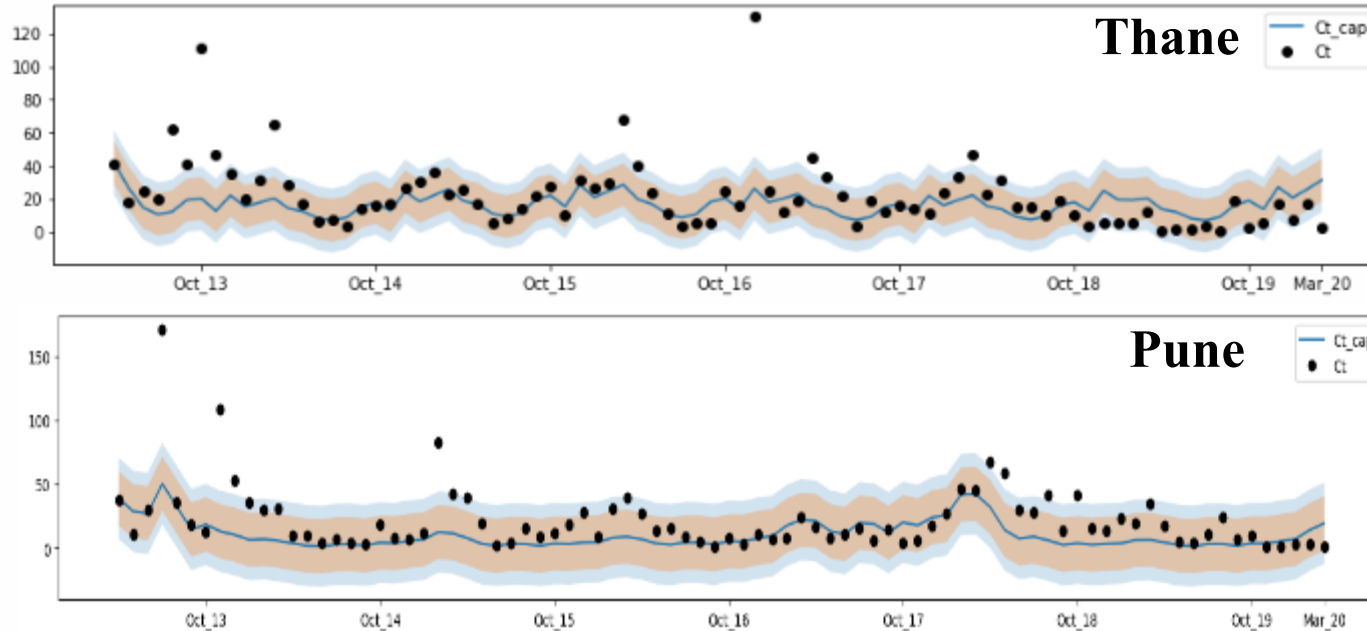




Findings for Mumbai

- Both the approaches show similar trend and annual seasonality in S_t and I_t .
- Effect of 2018 SIA is visible – temporary reduction in estimated susceptibility and infectious population
- This calls for the periodic SIA analysis (TSIR).

Findings for Thane and Pune based on TSIR



Actual case counts Vs. predicted case counts using TSIR model with SIA at Dec'18
(Blue shaded region indicates 3σ confidence interval, red shaded region indicates 2σ confidence interval)

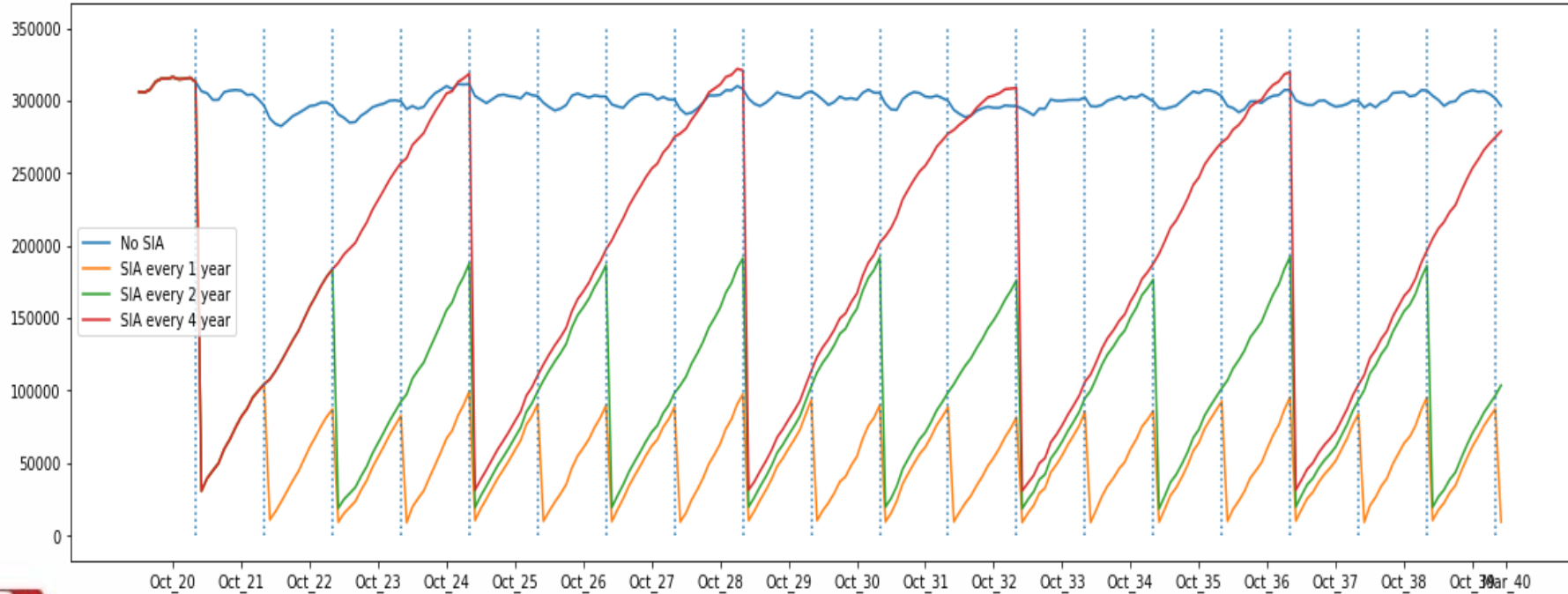


SIA Scenarios

- Coverage scenarios considered
 - Covering children upto 5 years
 - Covering children upto 14 years
 - Covering population upto 40 years



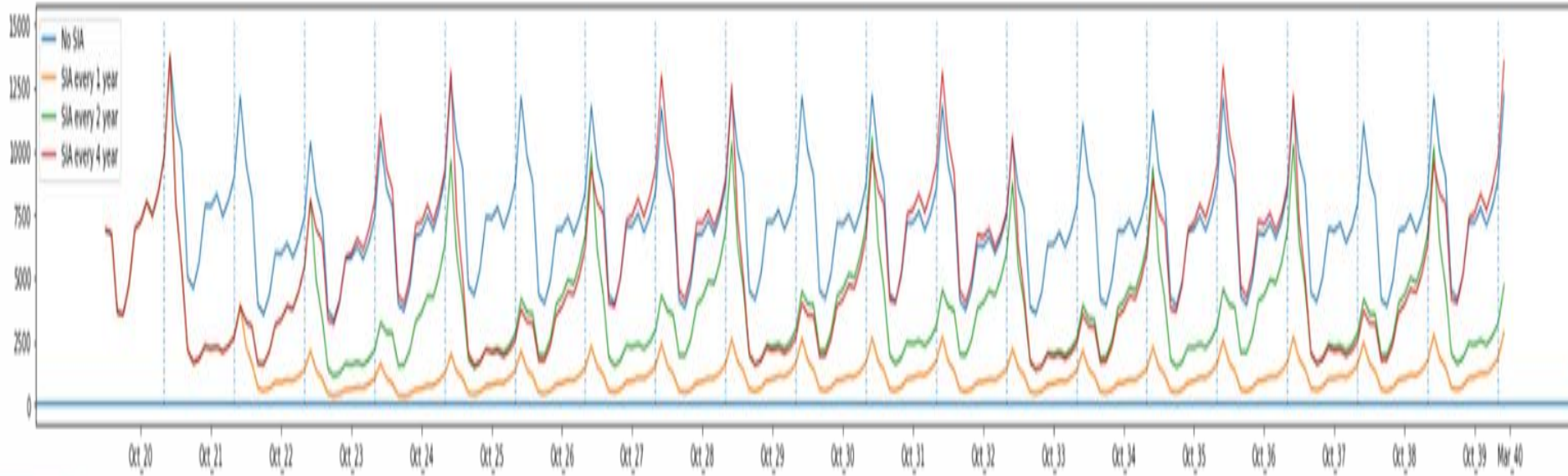
SIA plot for Mumbai: Susceptibility



Estimation based on HMIS, NFHS data for India



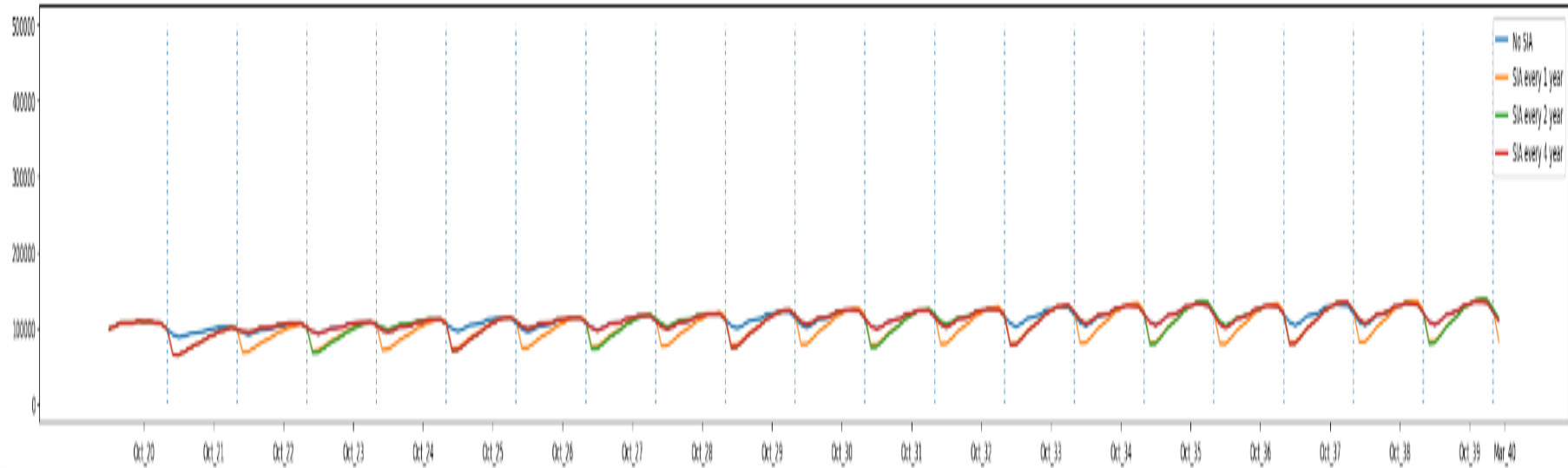
SIA plot for Mumbai: Infectious population



Estimation based on HMIS, NFHS data for India



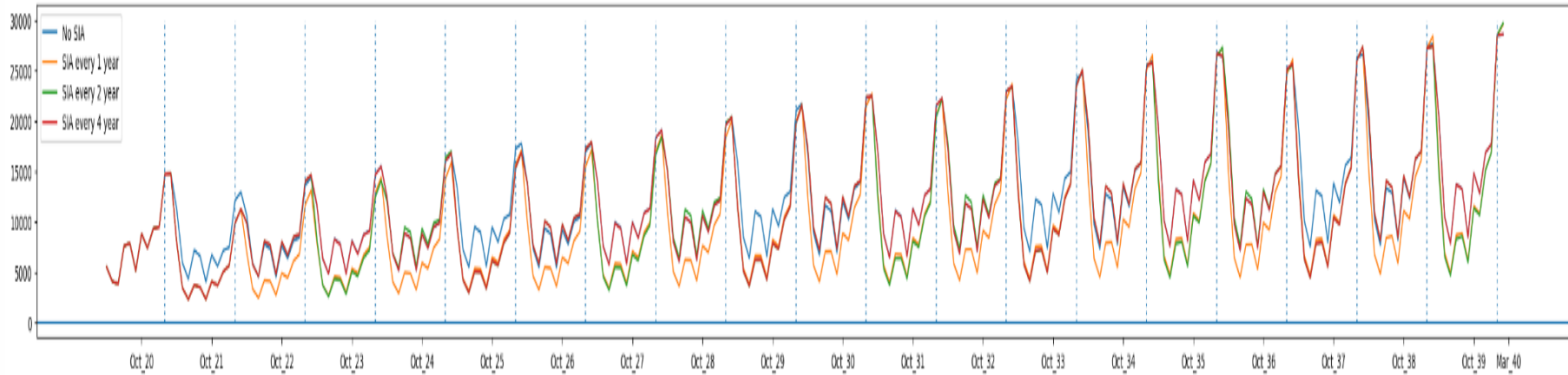
SIA Covering Children upto 14 years in Pune: Susceptibility



Estimation based on HMIS, NFHS data for India

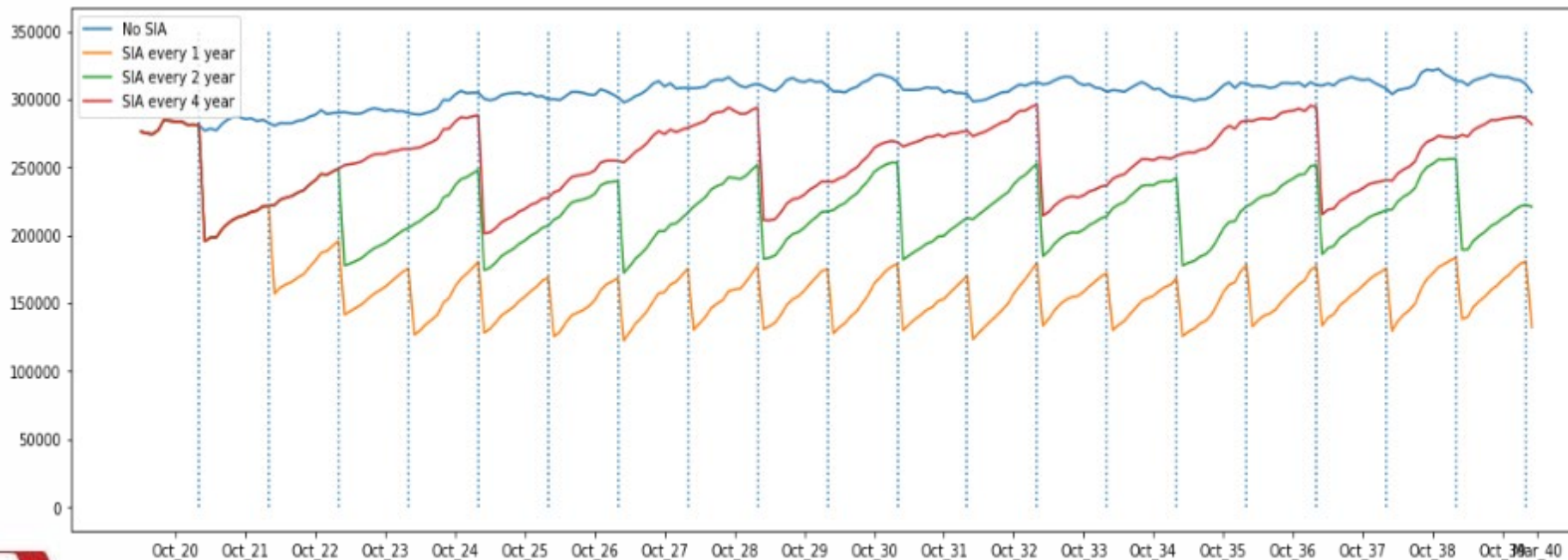


SIA Covering Children upto 14 years in Pune: Infectious population



Estimation based on HMIS, NFHS data for India

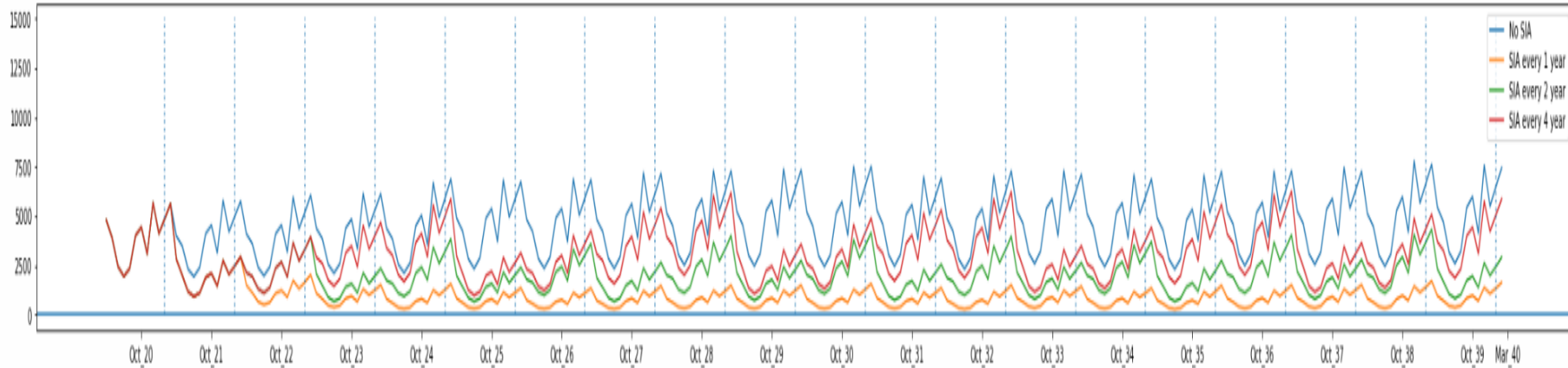
SIA Covering Children upto 14 years in Thane: Susceptibility



Estimation based on HMIS, NFHS data for India



SIA Covering Children upto 14 years in Thane: Infectious population



Estimation based on HMIS, NFHS data for India

Concluding Remarks



- The trend in the estimates of C_t and S_t are similar for both the approaches.
- State space modeling provides smaller RMSE, MAPE and confidence intervals.
- SIA analysis shows the reduction in measles incidences if regular SIAs are implemented.
- The susceptibility is seen to be increasing which calls for more aggressive vaccination campaigns and SIAs.



Future Work

- Incorporate spatio-temporal dependence amongst the measles burden across various districts.
- Analyze the effect of migrations.
- Include the age-wise stratification in the models.
- Include additional covariates which may act as potential predictors of measles burden.



Thank You