



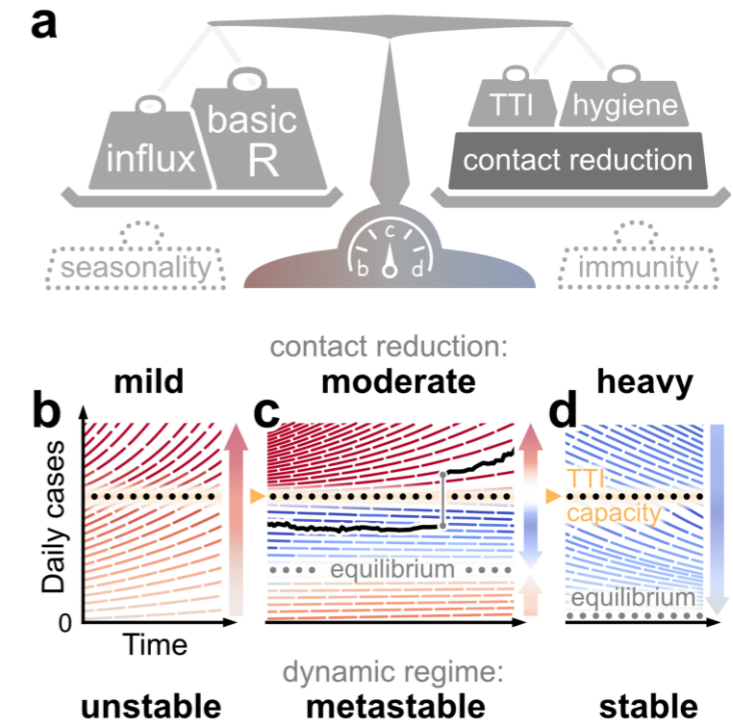
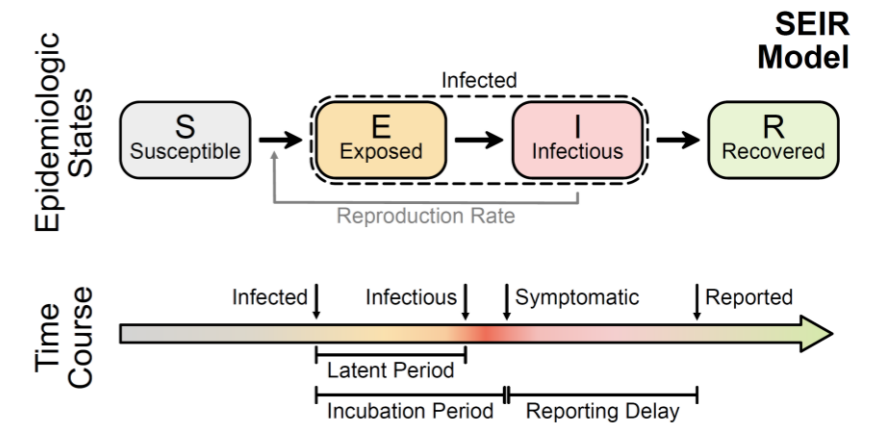
Spreading dynamics in neural networks - and of COVID-19

Viola Priesemann

Max-Planck-Institut für
Dynamik und
Selbstorganisation
Göttingen

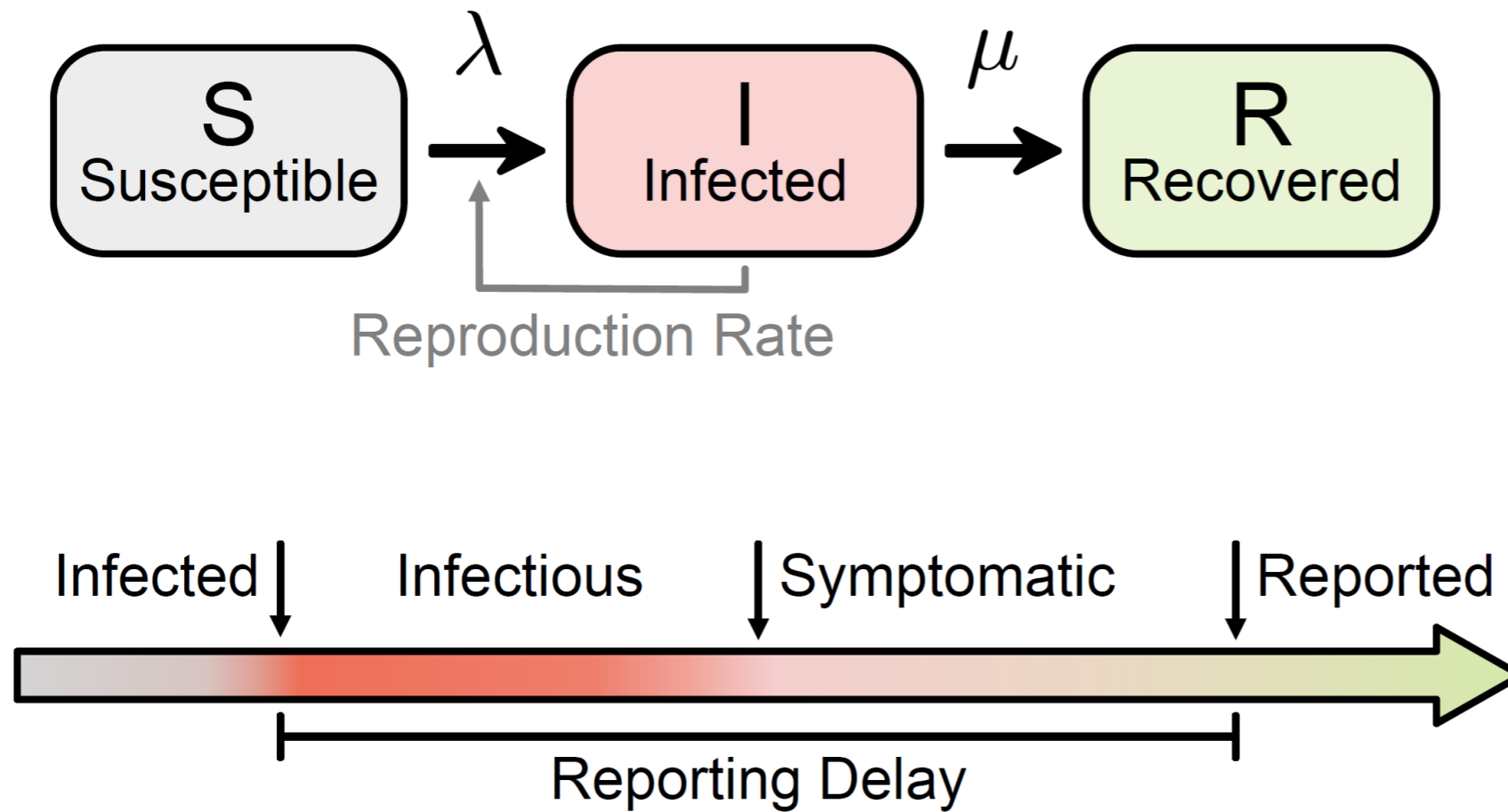
Content

- **Bayesian Inference and short-term forecast:**
SIR-Model, Hamiltonian MCMC, weekly change points
- **Progress of vaccination:** How fast can we lift „non-pharmaceutical interventions“ (NPIs) given the planned vaccination progress?
- **Derive effectiveness of measures:**
Contribution of test-trace-isolate (TTI) to containment of COVID-19



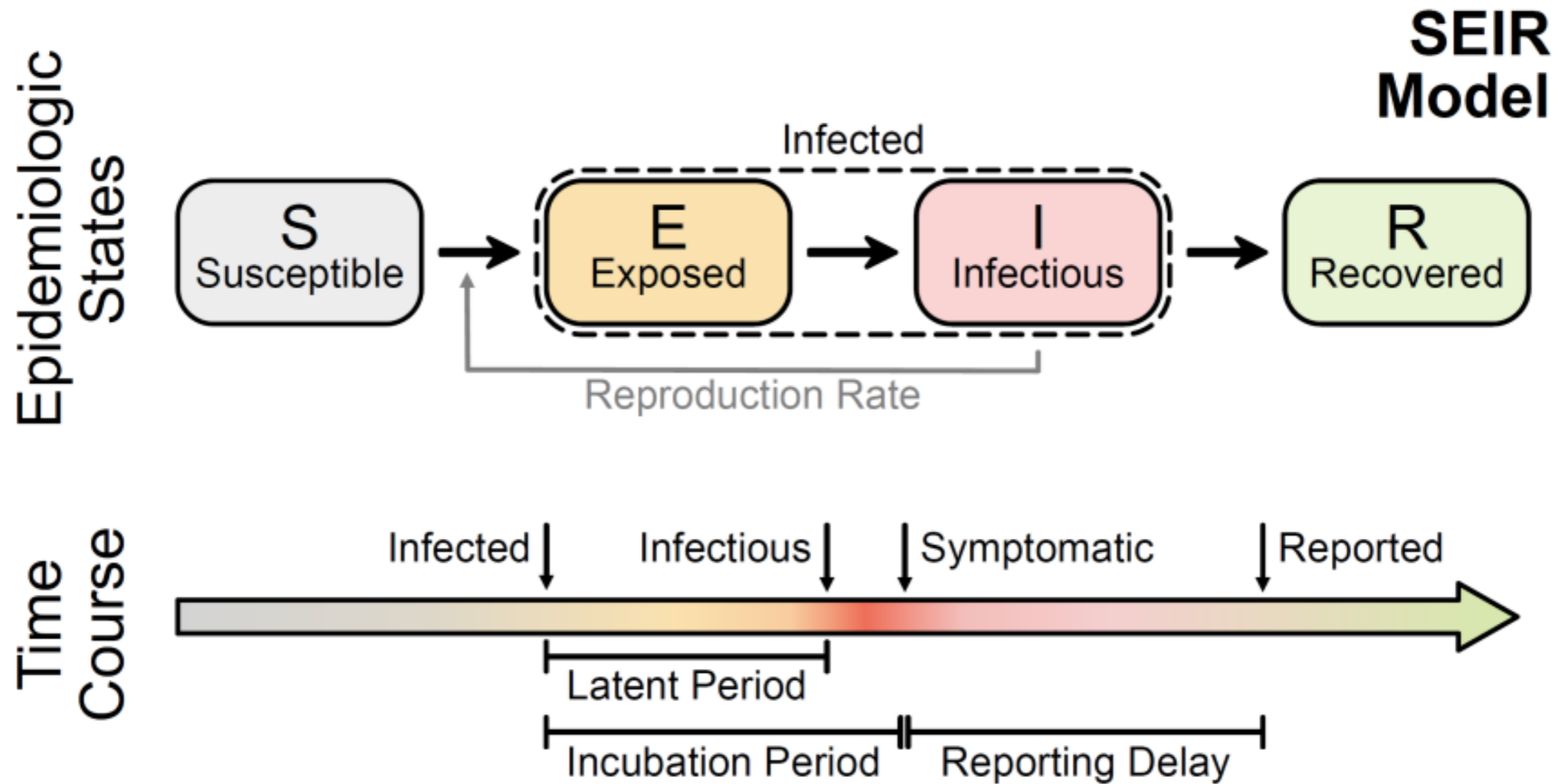
SIR: Susceptible-Infected-Recovered

SIR Model

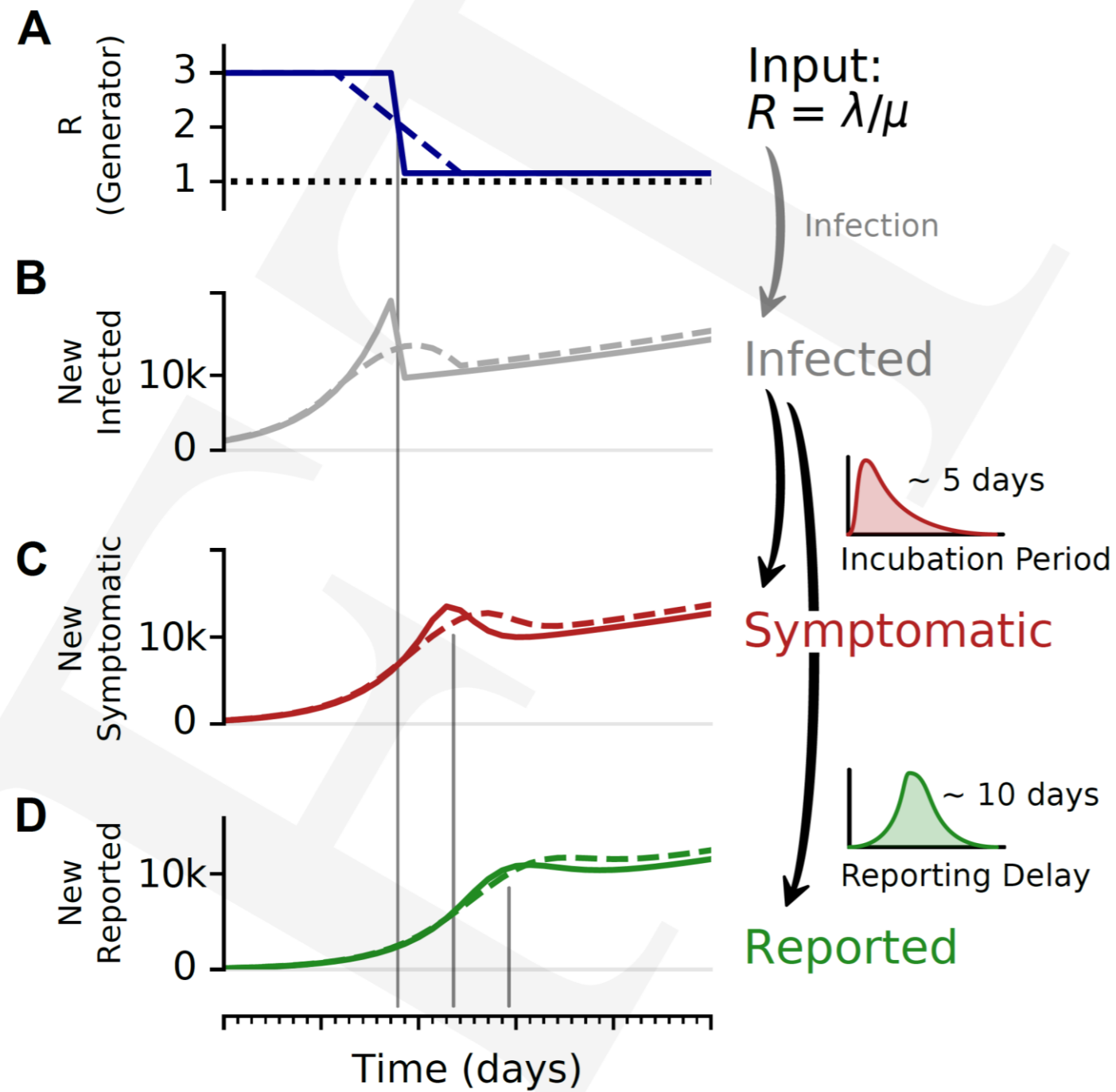


$$\begin{aligned}\frac{dS}{dt} &= -\lambda \frac{SI}{N} \\ \frac{dI}{dt} &= \lambda \frac{SI}{N} - \mu I \\ \frac{dR}{dt} &= \mu I\end{aligned}$$

SEIR: Susceptible-Exposed-Infected-Recovered



Inference of R under fast changes



Bayesian Inference of R

- SIR-model:

- Set begin: $I_0 = I_0, S_0 = N - I_0$
- recursion: $I_{t+1} - I_t = \lambda I_t \frac{S}{N} - \mu I_t$
 $S_{t+1} - S_t = -\lambda I_t \frac{S}{N}$

I_t Infected persons at time t

S_t Susceptible persons at time t

λ Infection rate

μ Recovery rate

N Total number of persons in the population

Bayesian Inference of R

- SIR-model:

- Set begin: $I_0 = I_0, S_0 = N - I_0$
- recursion: $I_{t+1} - I_t = \lambda I_t \frac{S}{N} - \mu I_t$
 $S_{t+1} - S_t = -\lambda I_t \frac{S}{N}$

I_t Infected persons at time t

S_t Susceptible persons at time t

λ Infection rate

μ Recovery rate

N Total number of persons in the population

- The cases are reported D days later: $C_{t+D} = I_t$

- change points of lambda with times t_i , and length Δt_i

- Weekly modulation with amplitude f_w and offset Φ_w .

$$f(t) = (1 - f_w) \cdot 1 - |\sin(\frac{\pi}{7}t - \frac{1}{2}\Phi_w)|$$

-> Likelihood $P(C_{t,meas} | \theta) \sim \text{StudentT}_{\nu=4} \left(\text{mean} = C_t(\theta), \text{width} = \sigma \sqrt{C_t(\theta)} \right)$

- We use Hamiltonian Monte Carlo which adds a momentum, a modified proposal density and rejection criterion

Bayesian Inference of R

- SIR-model:

- Set begin: $I_0 = I_0, S_0 = N - I_0$
- recursion: $I_{t+1} - I_t = \lambda I_t \frac{S}{N} - \mu I_t$
 $S_{t+1} - S_t = -\lambda I_t \frac{S}{N}$

I_t Infected persons at time t

S_t Susceptible persons at time t

λ Infection rate

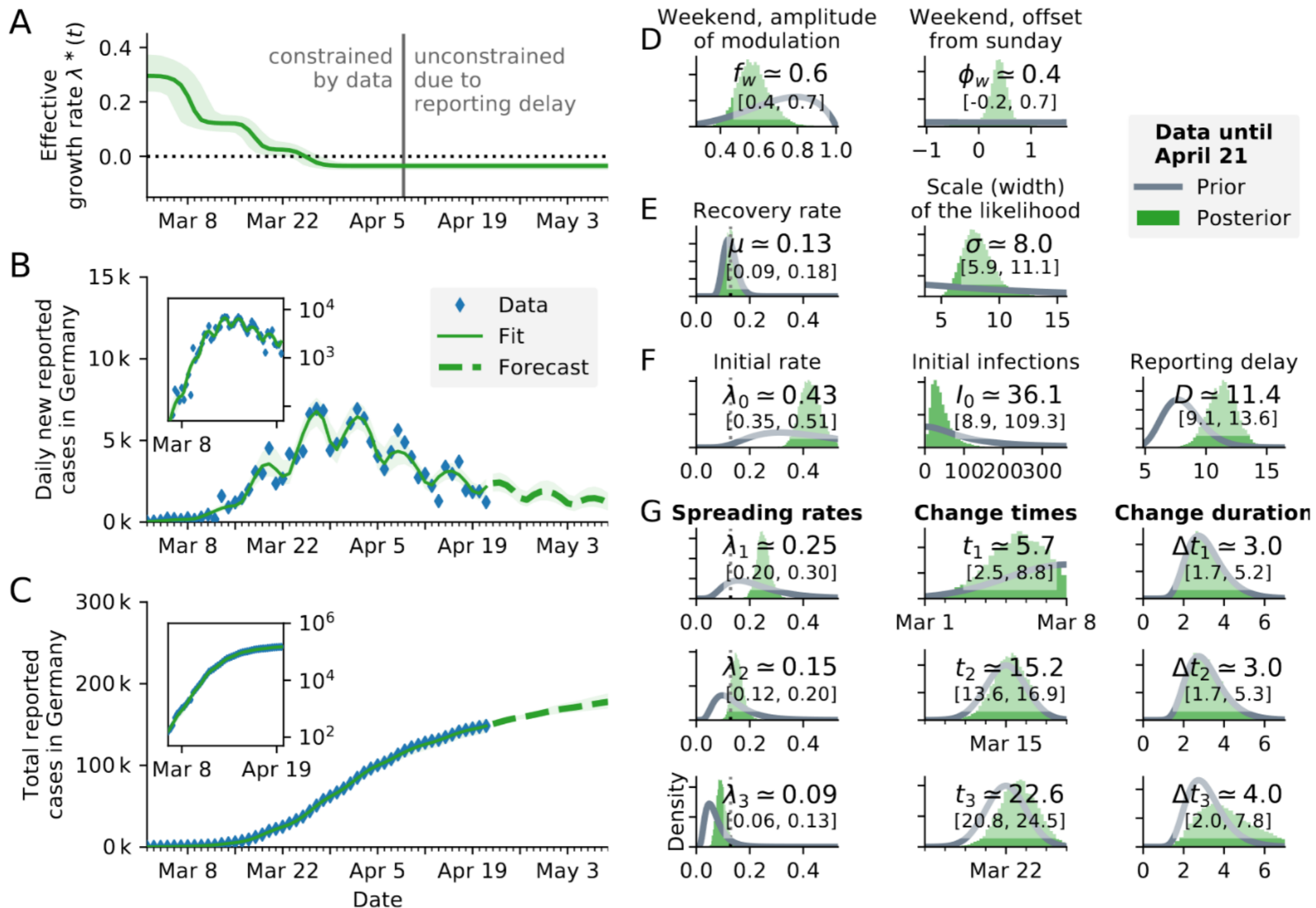
μ Recovery rate

N Total number of persons in the population

Priors:

Parameter	Variable	Prior distribution
Change points	t_i	Normal(Sunday of week i , 7)
Change duration	Δt_i	LogNormal($\log(4)$, 0.5)
Spreading rates	λ_0	LogNormal($\log(1/8)$, 0.5)
	λ_i	Relative to $i - 1$ with factor (country & week dependent)
Recovery rate	μ	LogNormal($\log(1/8)$, 0.2)
Reporting delay	D	LogNormal($\log(8)$, 0.2)
Weekly modulation amplitude	f_w	Beta(0.7, 0.17)
Weekly modulation phase	Φ_w	vonMises(0, 0.01) (nearly flat)
Initially infected	$I(0)$	HalfCauchy(depending on country)

Results March/April 2020

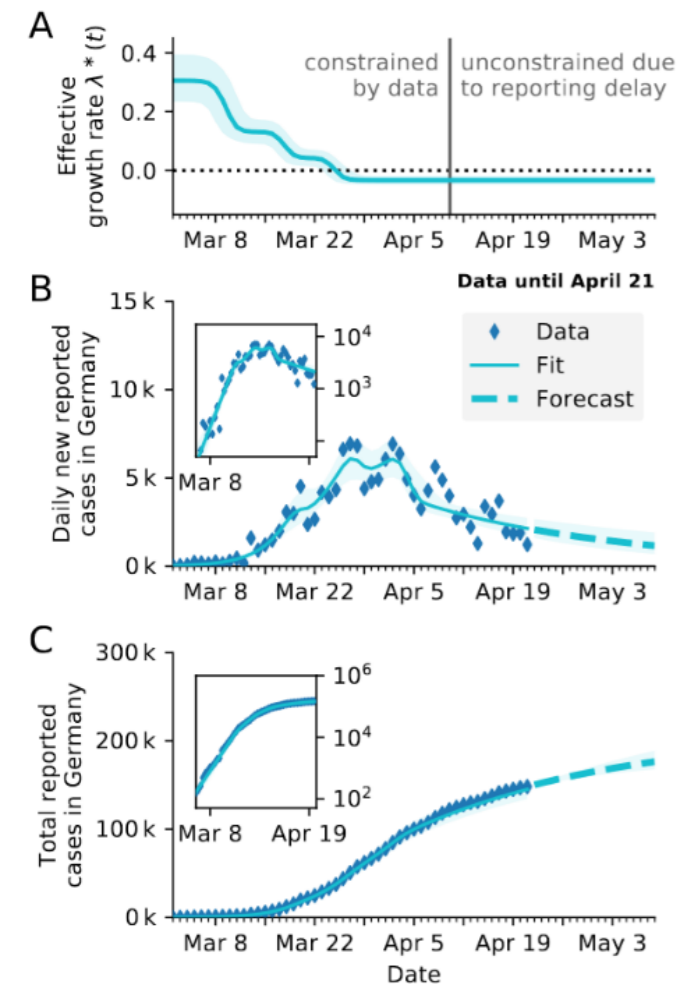


Adapted from [Dehning et al., Priesemann, Science, 2020]

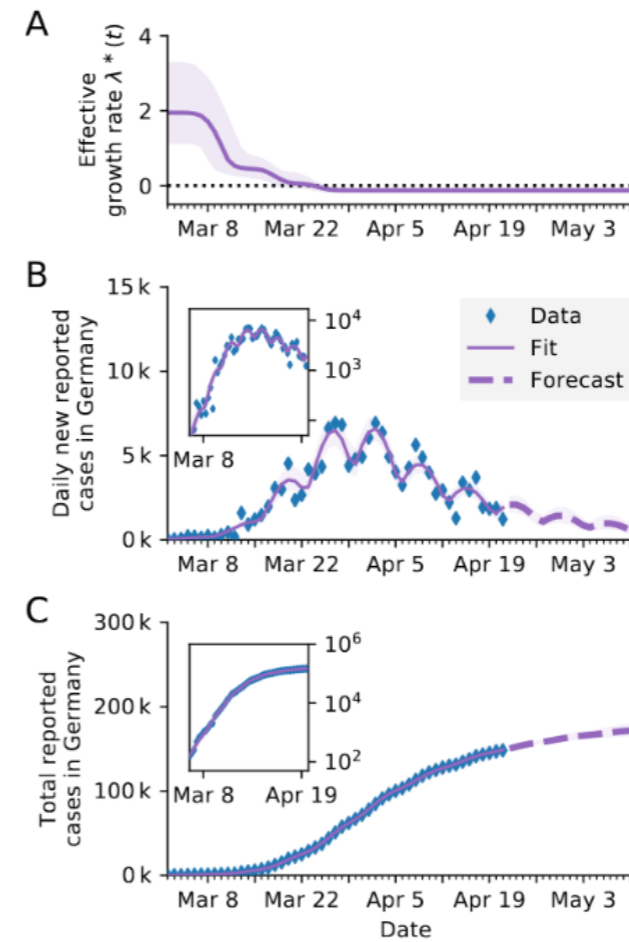
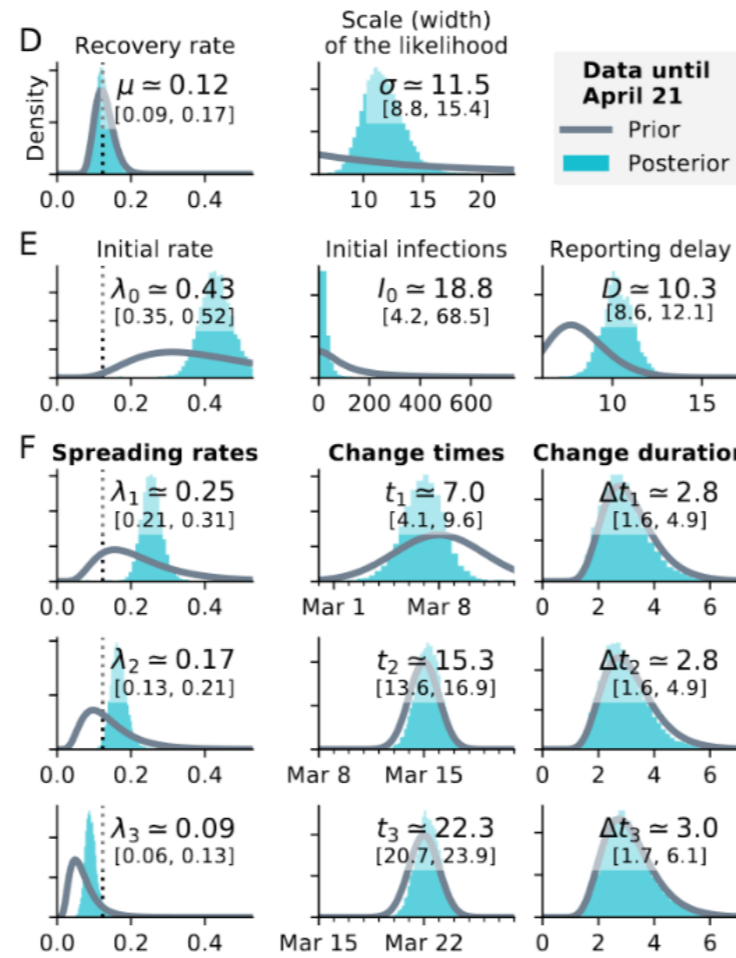
[Dehning et al., Priesemann, medRxiv]

<https://www.medrxiv.org/content/medrxiv/early/2020/09/18/2020.09.16.20187484.full.pdf>

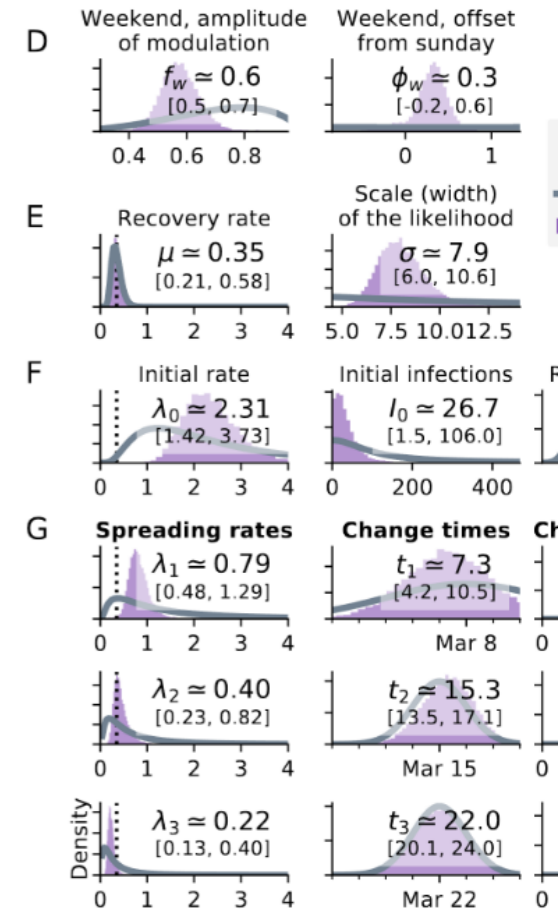
Different Models



Without weekend modulation



With incubation period

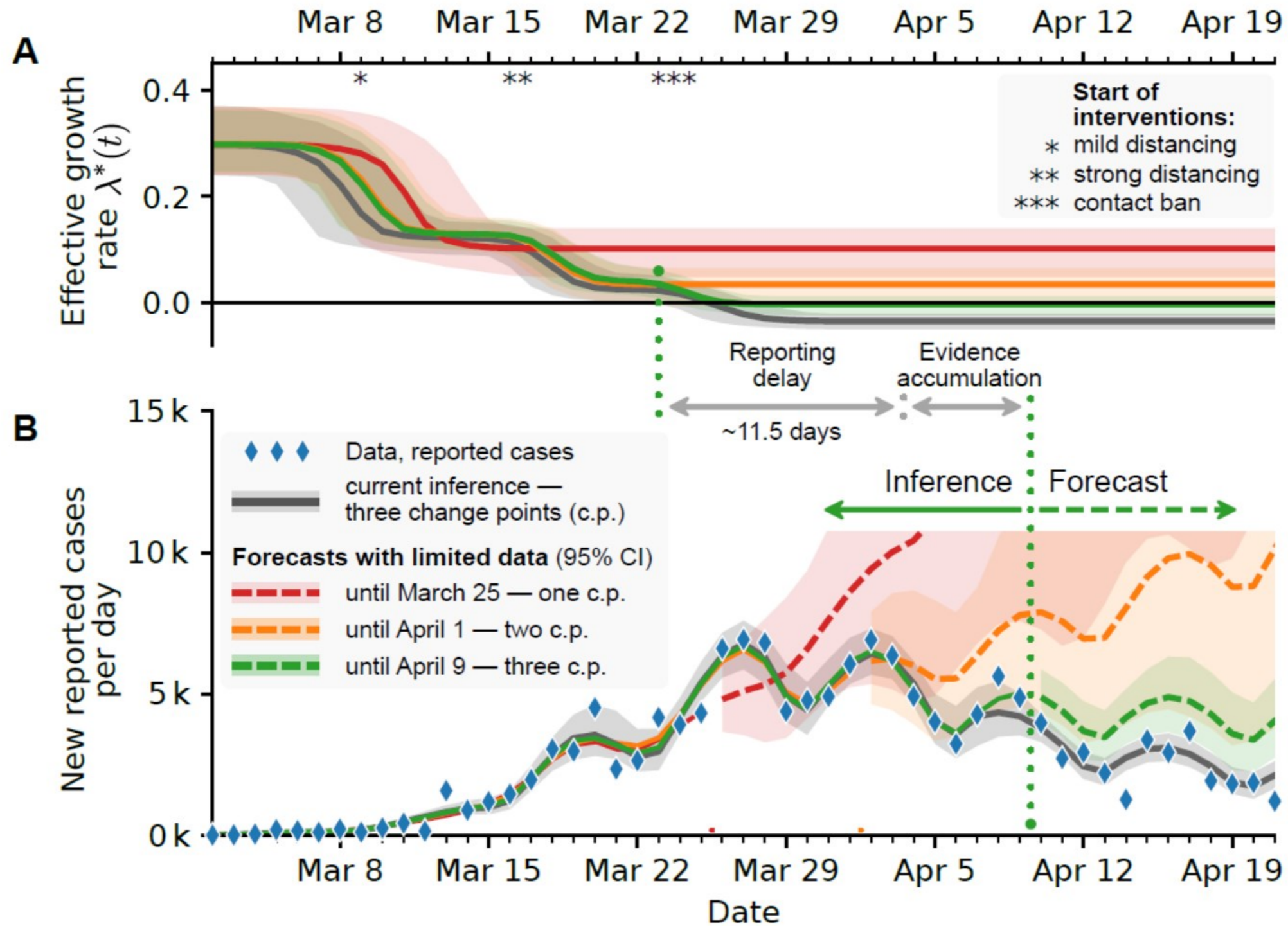


Adapted from [Dehning et al., Priesemann, Science, 2020]
[Dehning et al., Priesemann, medRxiv]

<https://www.medrxiv.org/content/medrxiv/early/2020/09/18/2020.09.16.20187484.full.pdf>

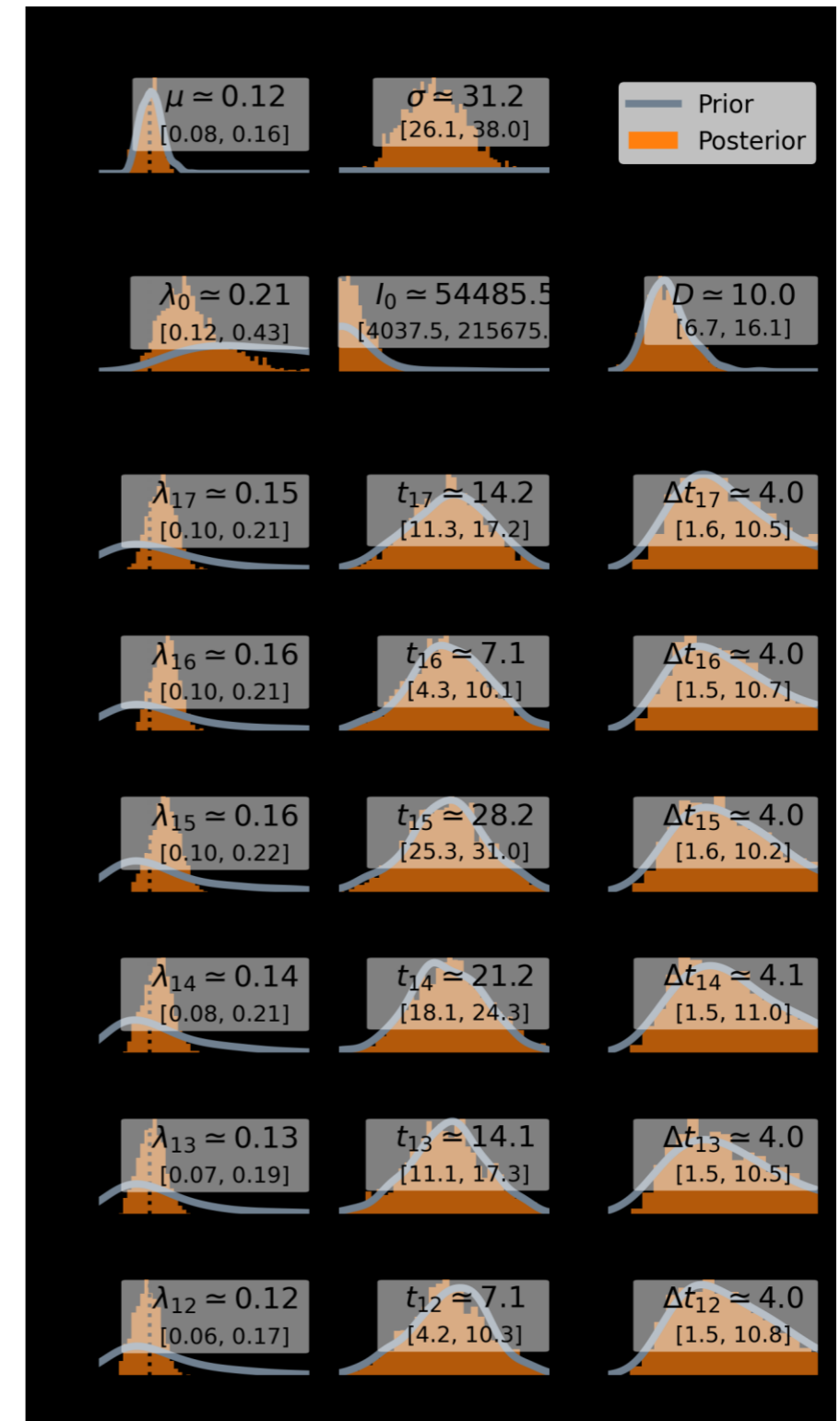
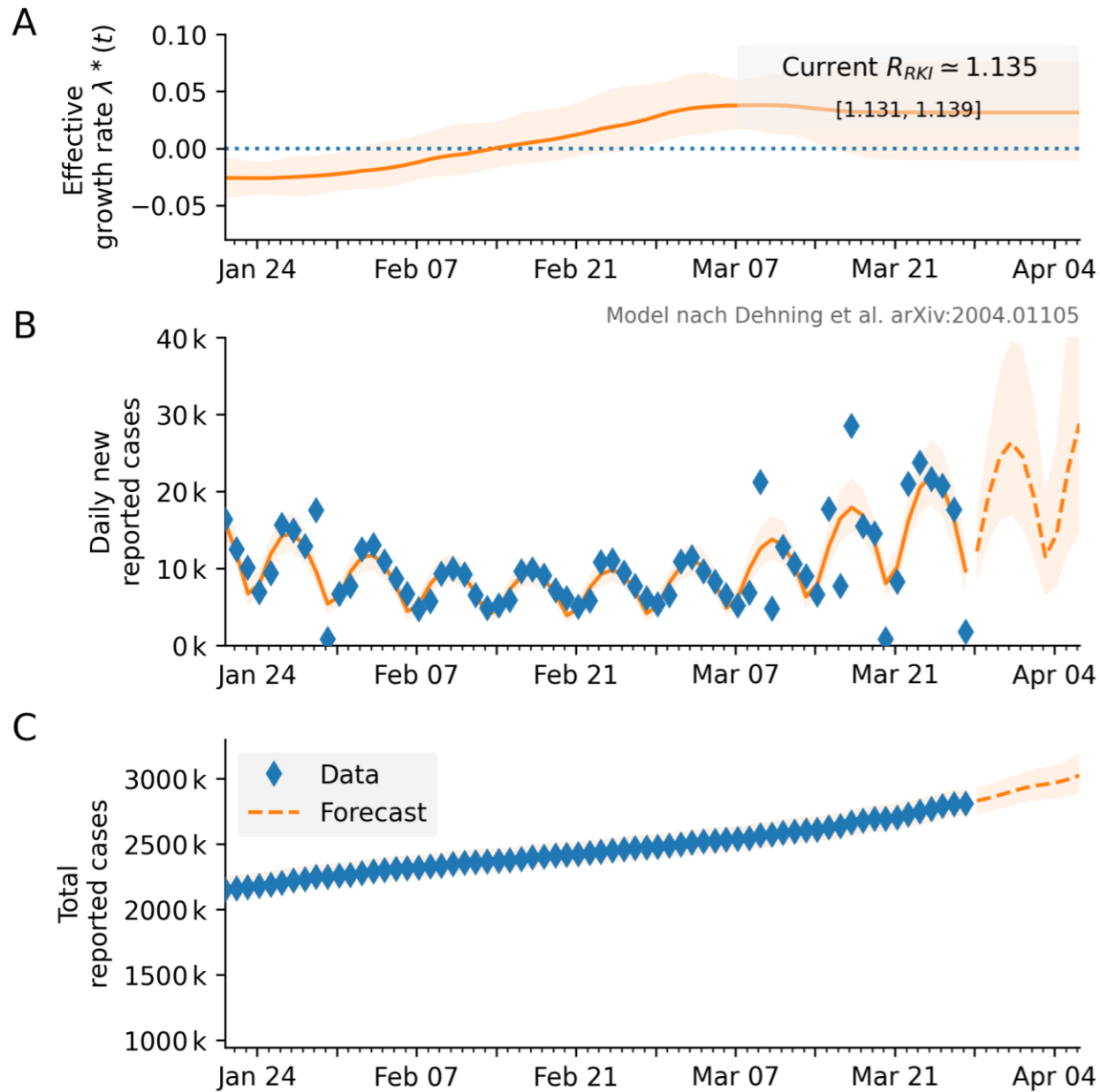
Analysis of the first wave

COVID-19 analysis Germany (as of April 21, 2020)



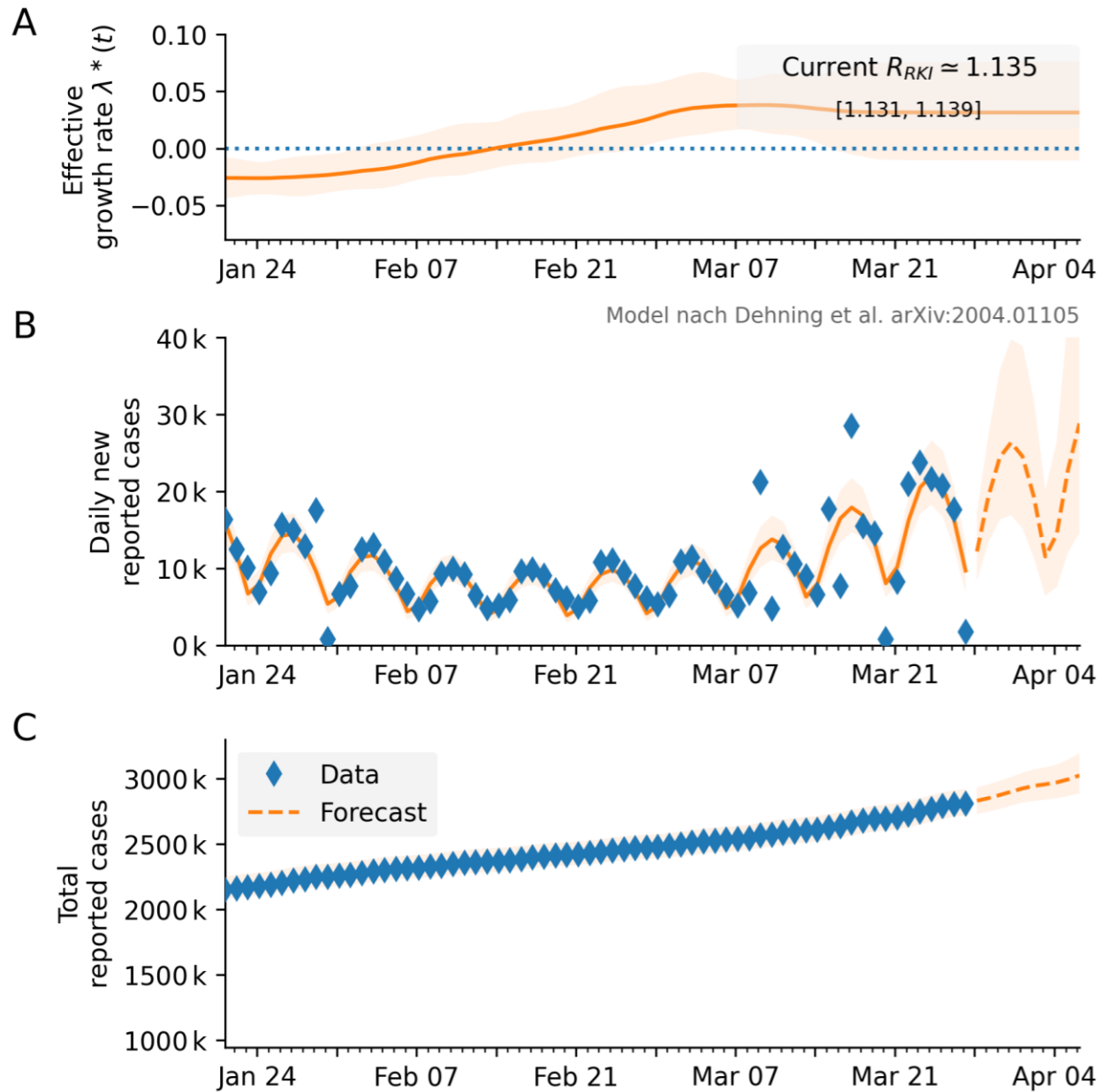
Daily new estimates for Germany

Updated on
2021/03/30



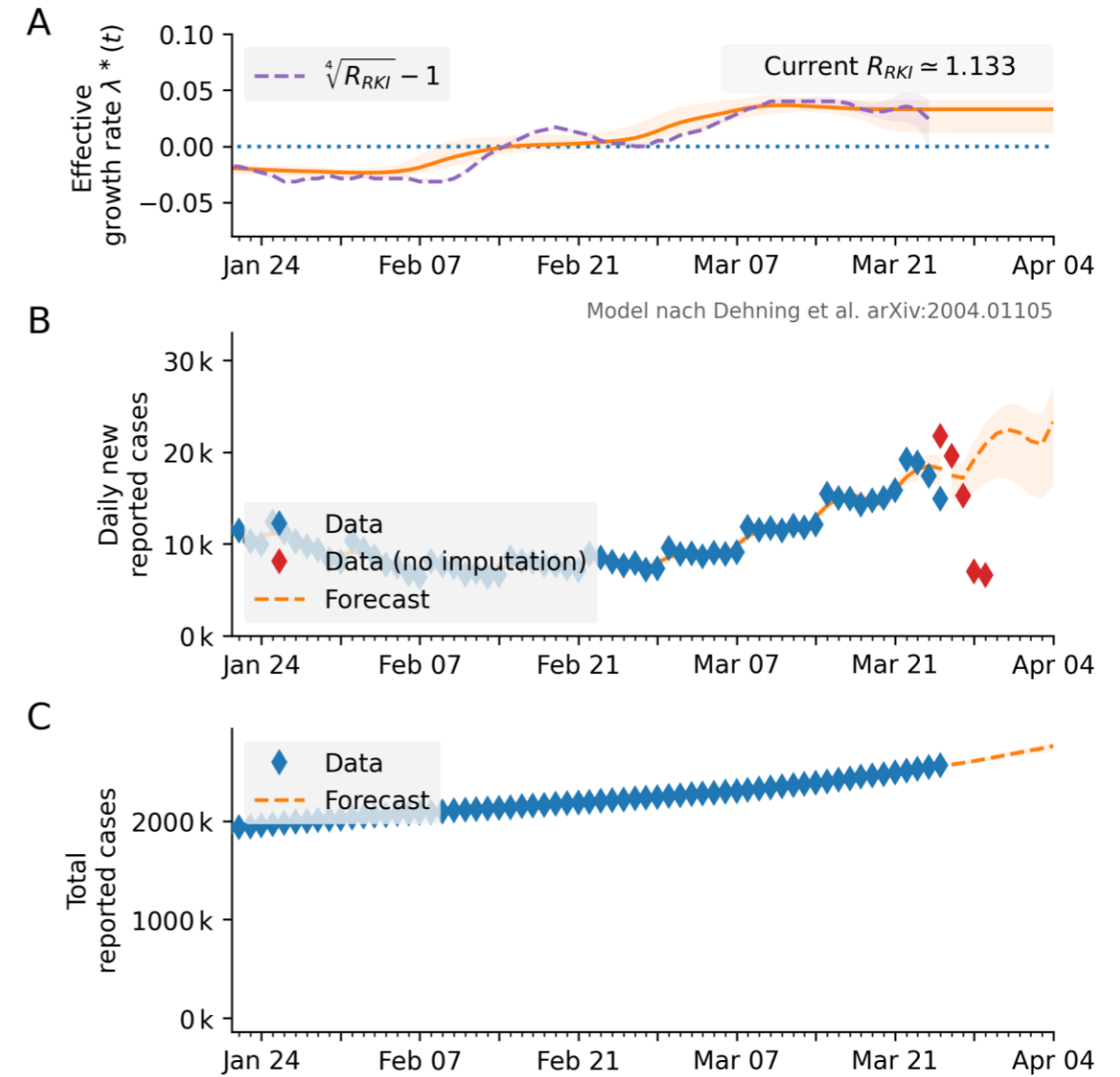
Daily new estimates for Germany

Updated on
2021/03/30



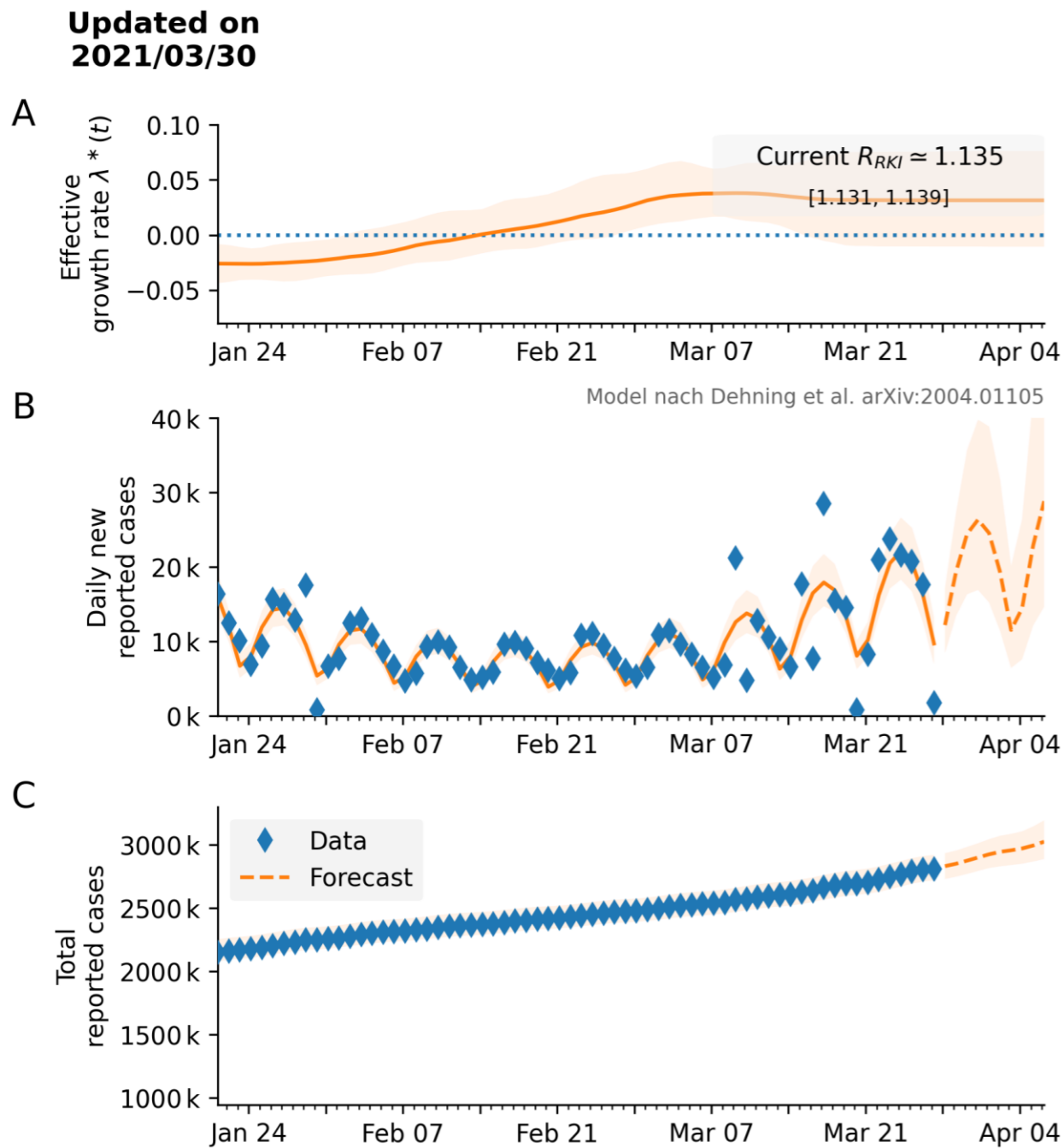
Data Source: OWD

Updated on
2021/03/30

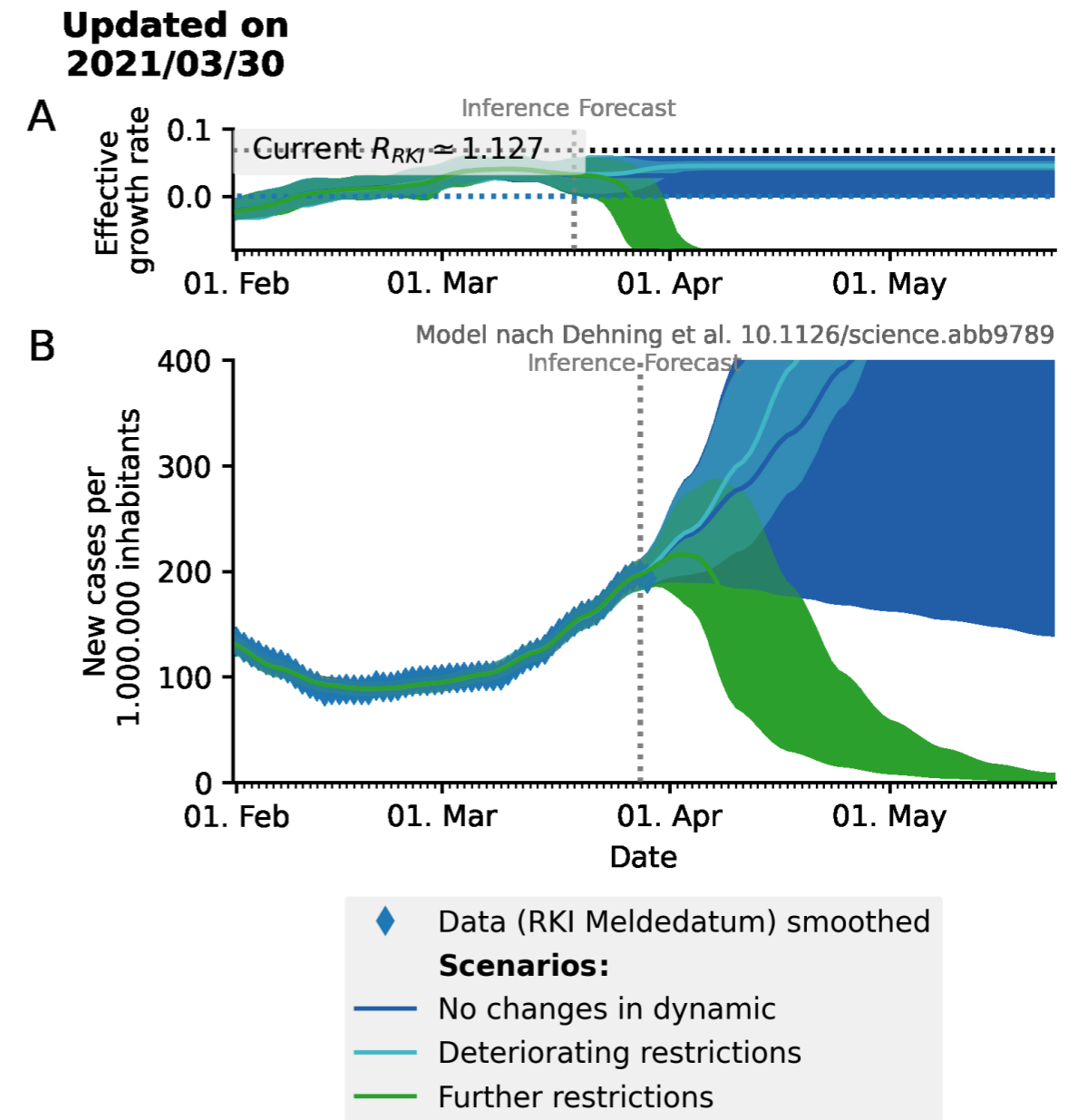


Data Source: RKI Nowcast

Daily new estimates for Germany



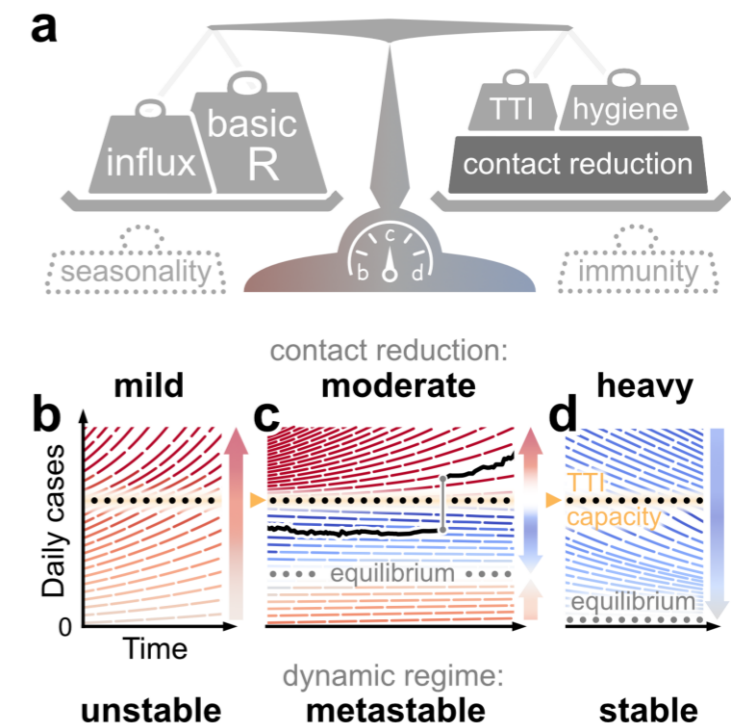
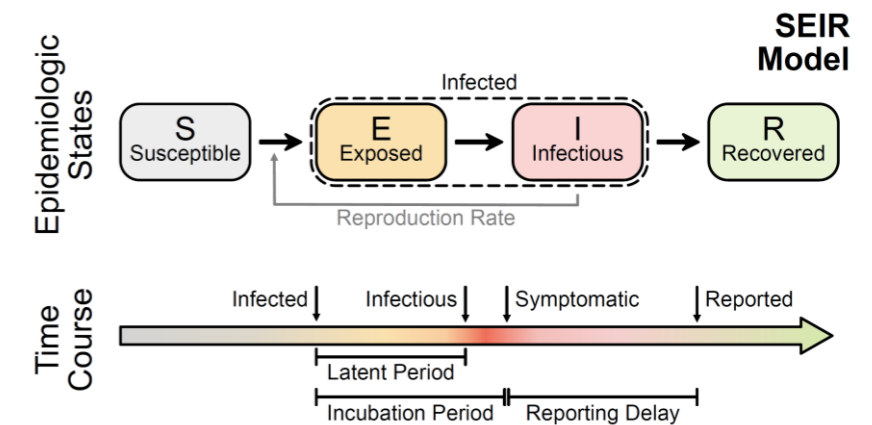
Data Source: OWD



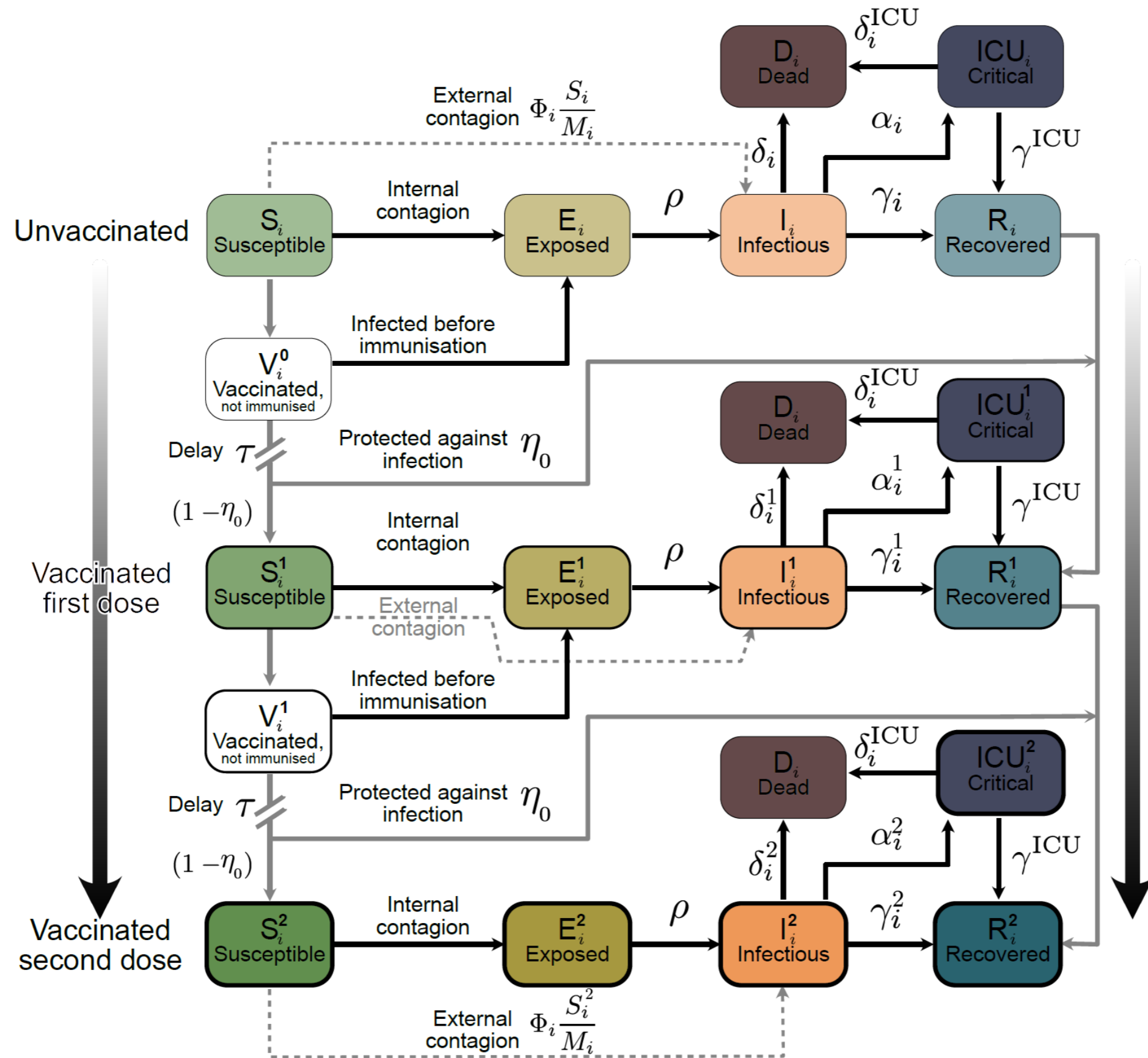
Data Source: RKI Reporting Date

Content

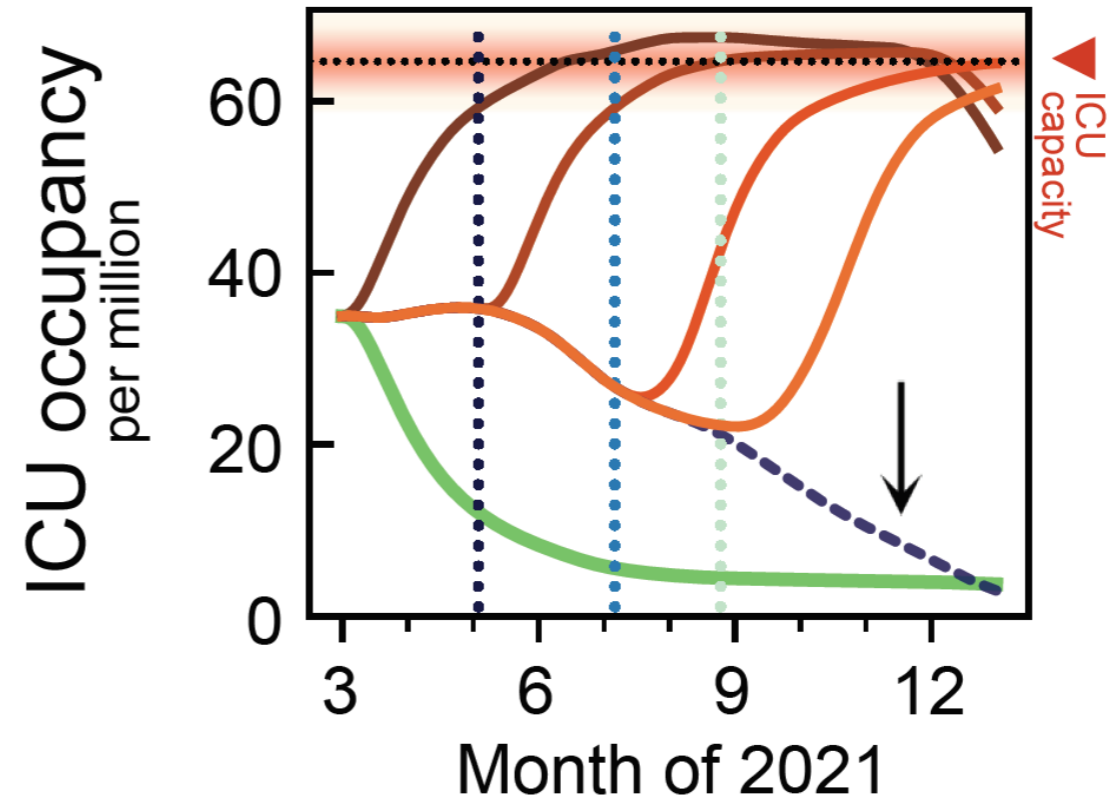
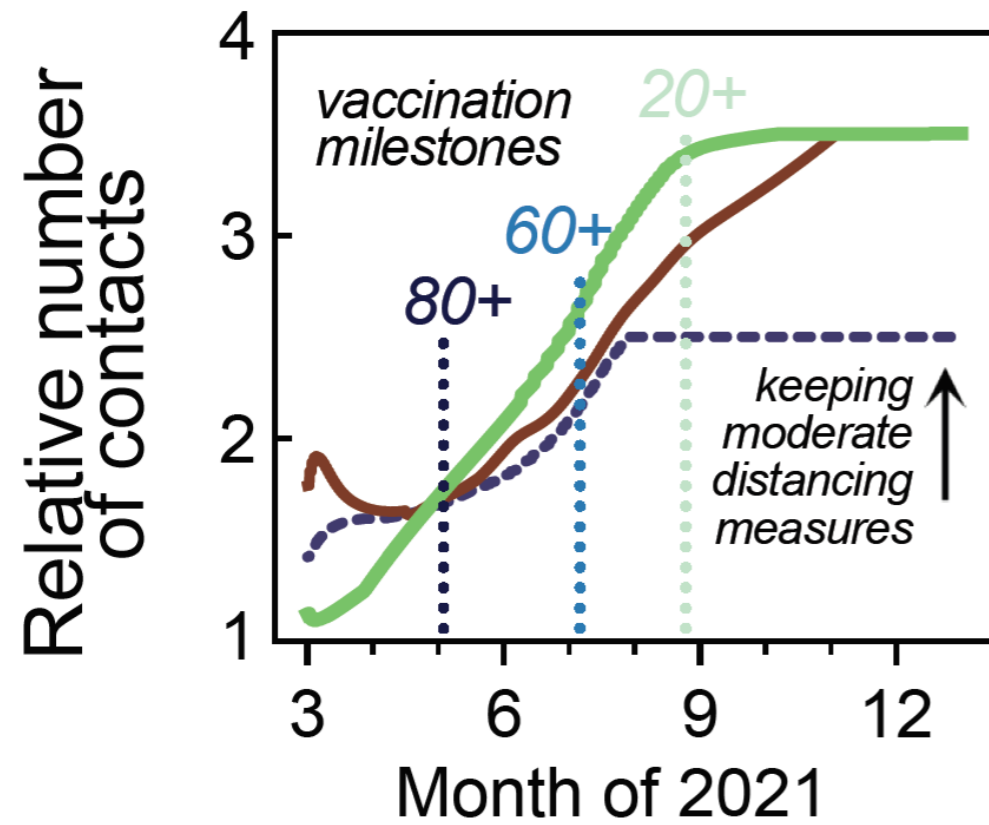
- **Bayesian Inference and short-term forecast:**
SIR-Model, Hamiltonian MCMC, weekly change points
- **Progress of vaccination determines opening steps.**
How fast can we lift „non-pharmaceutical interventions“ (NPIs) given the planned vaccination progress?
- **Derive effectiveness of measures:**
Contribution of test-trace-isolate (TTI) to containment of COVID-19



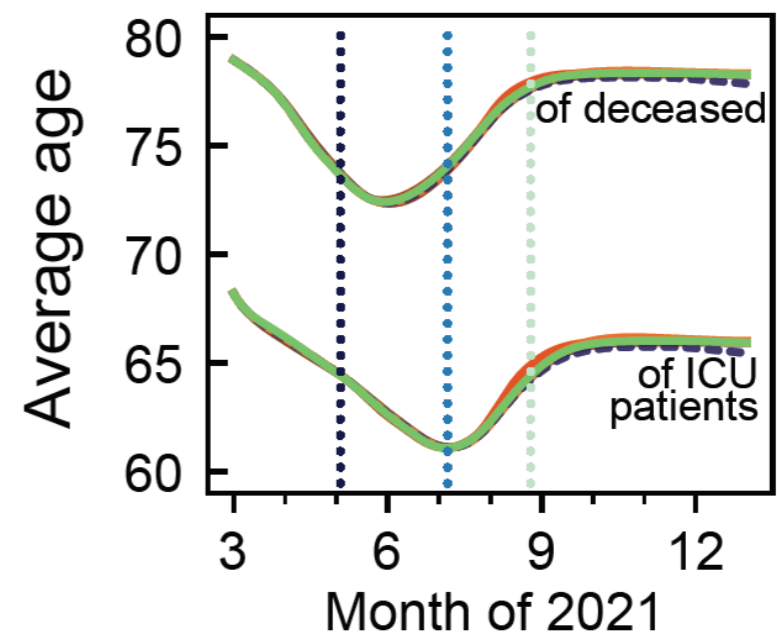
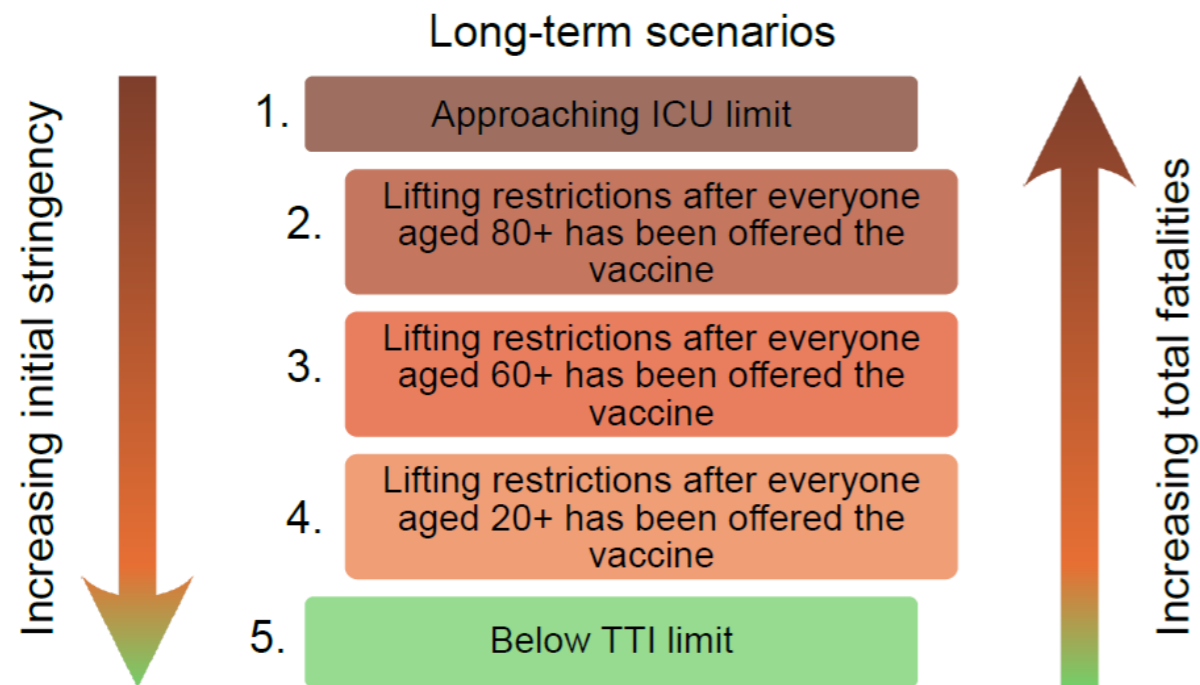
Compartmental Vaccination Model – Age Dependent



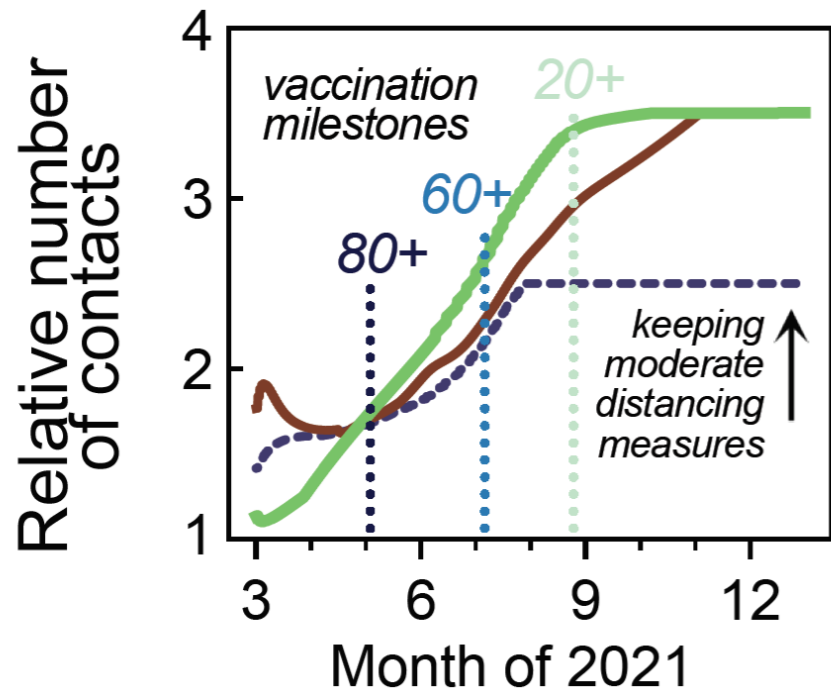
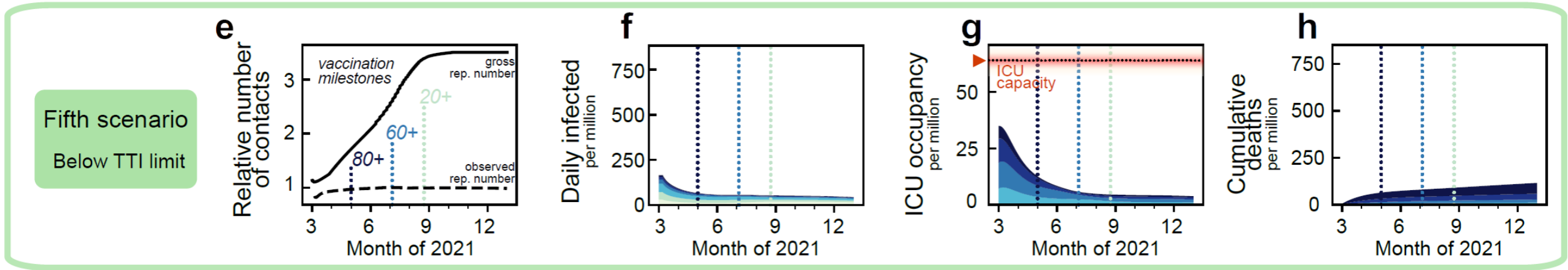
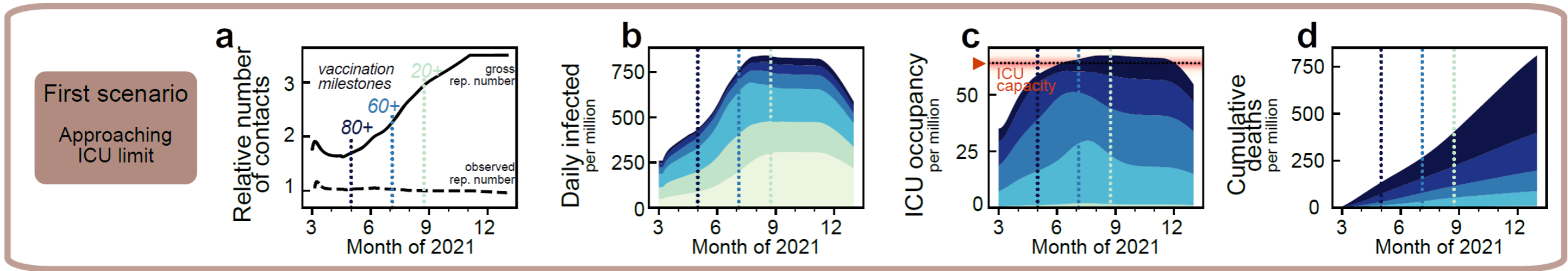
Different Strategies of Opening



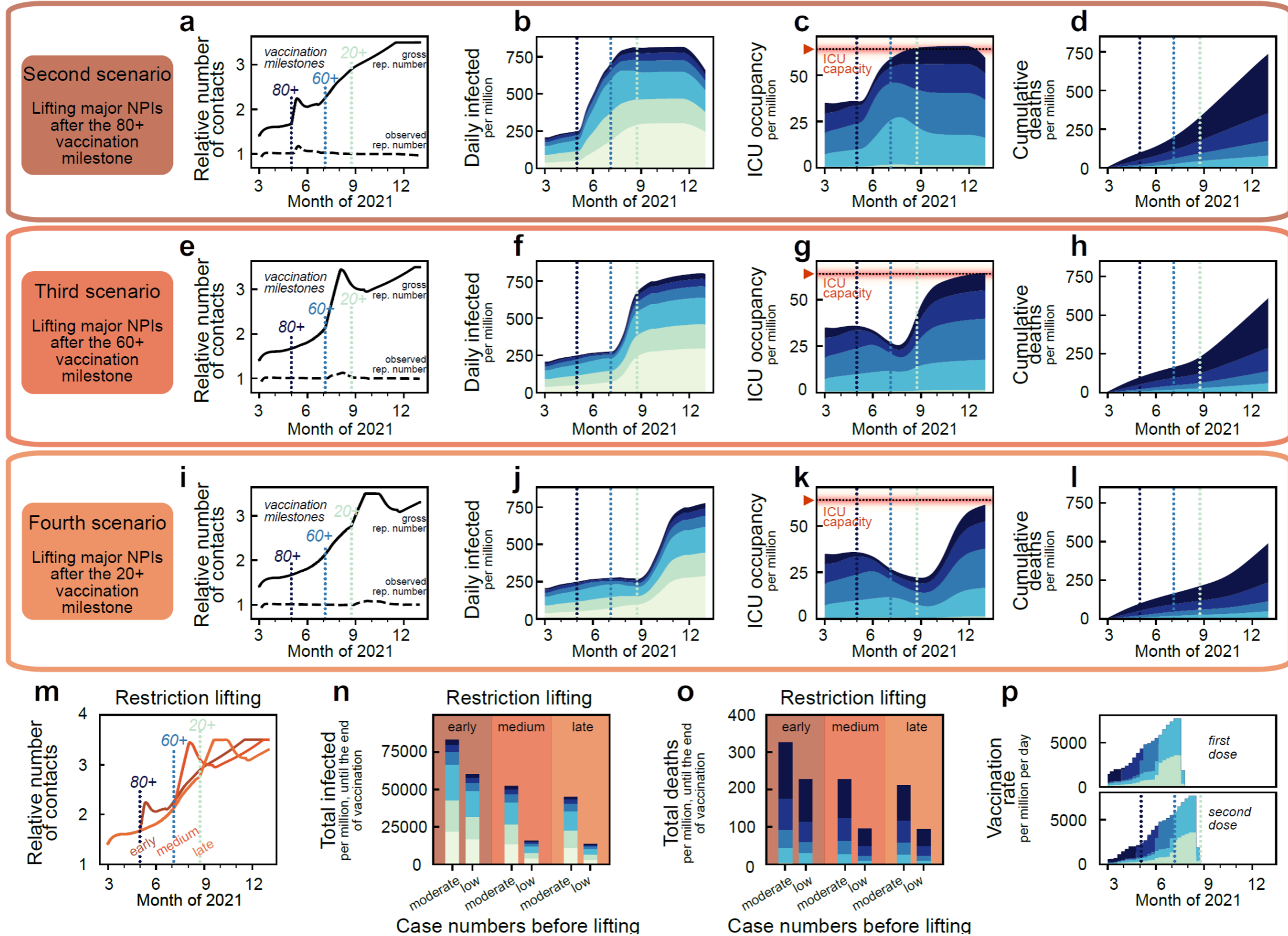
b



Different Strategies of Opening

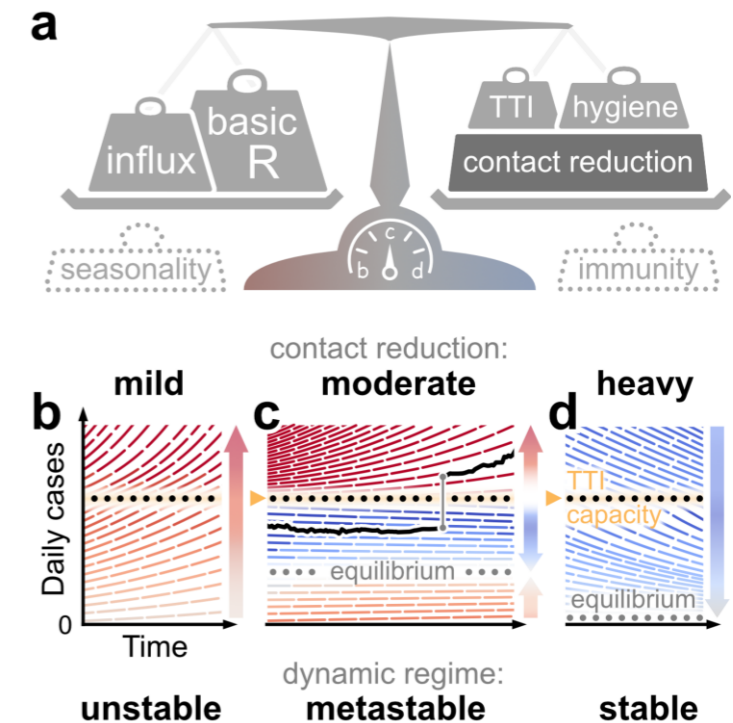
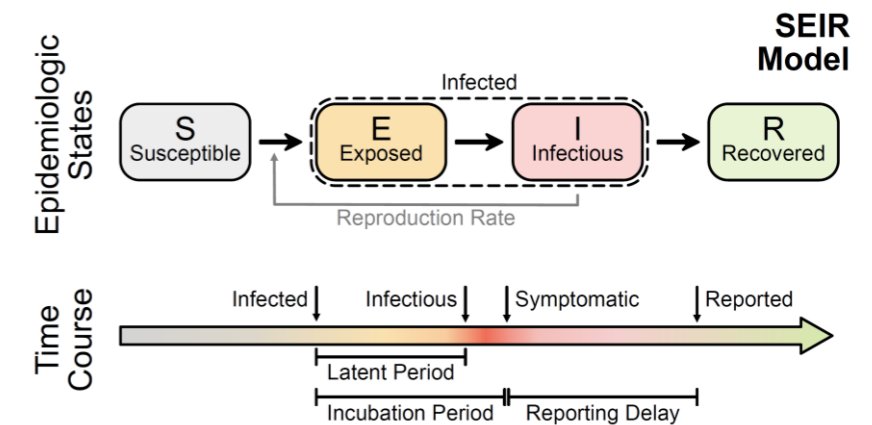


Different Strategies of Opening

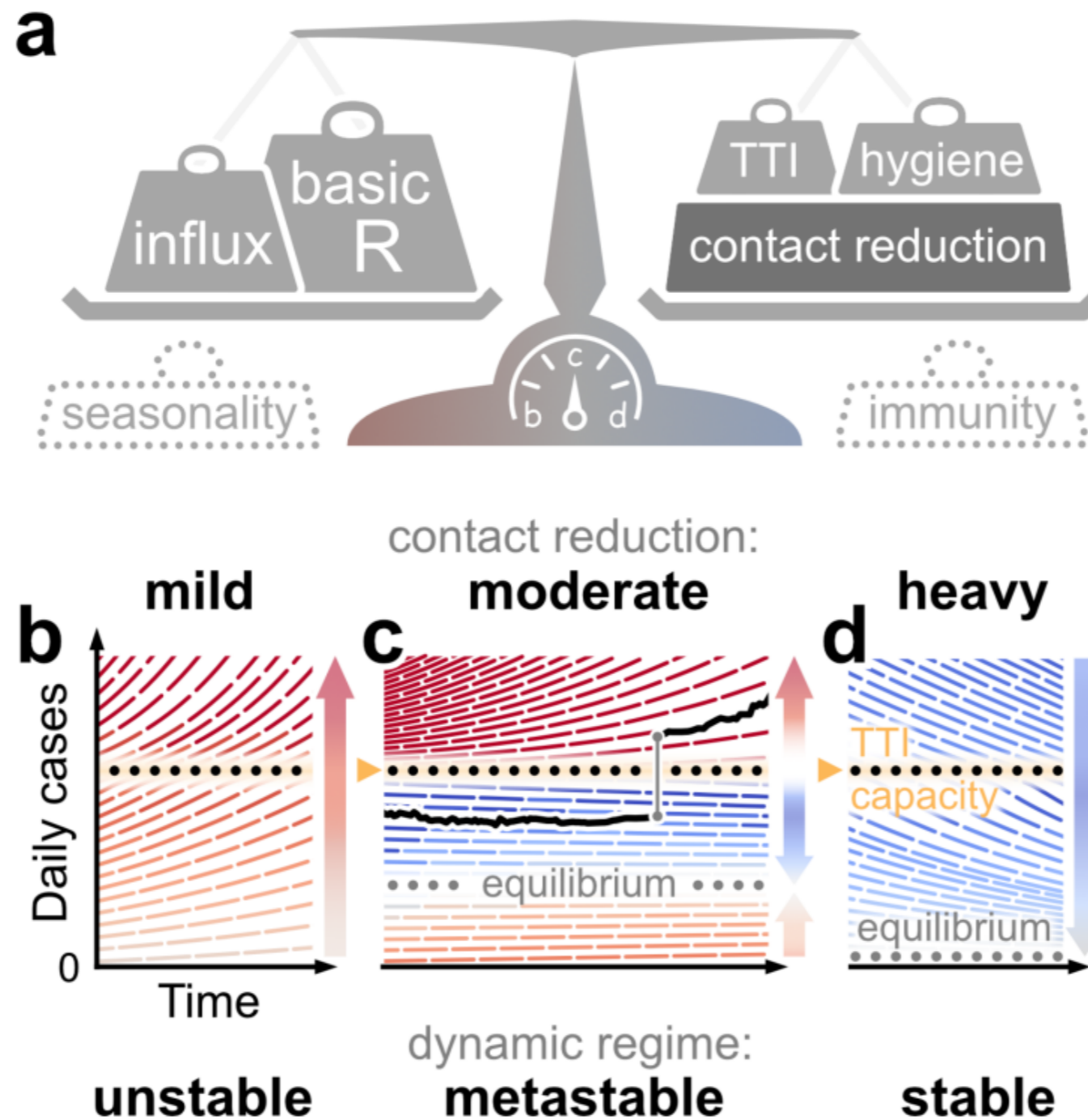


Content







- **Bayesian Inference and short-term forecast:**
SIR-Model, Hamiltonian MCMC, weekly change points
- **Progress of vaccination determines opening steps.**
How fast can we lift „non-pharmaceutical interventions“ (NPIs) given the planned vaccination progress?
- **Derive effectiveness of measures:**
Contribution of test-trace-isolate (TTI) to containment of COVID-19



Combined measures to contain COVID-19

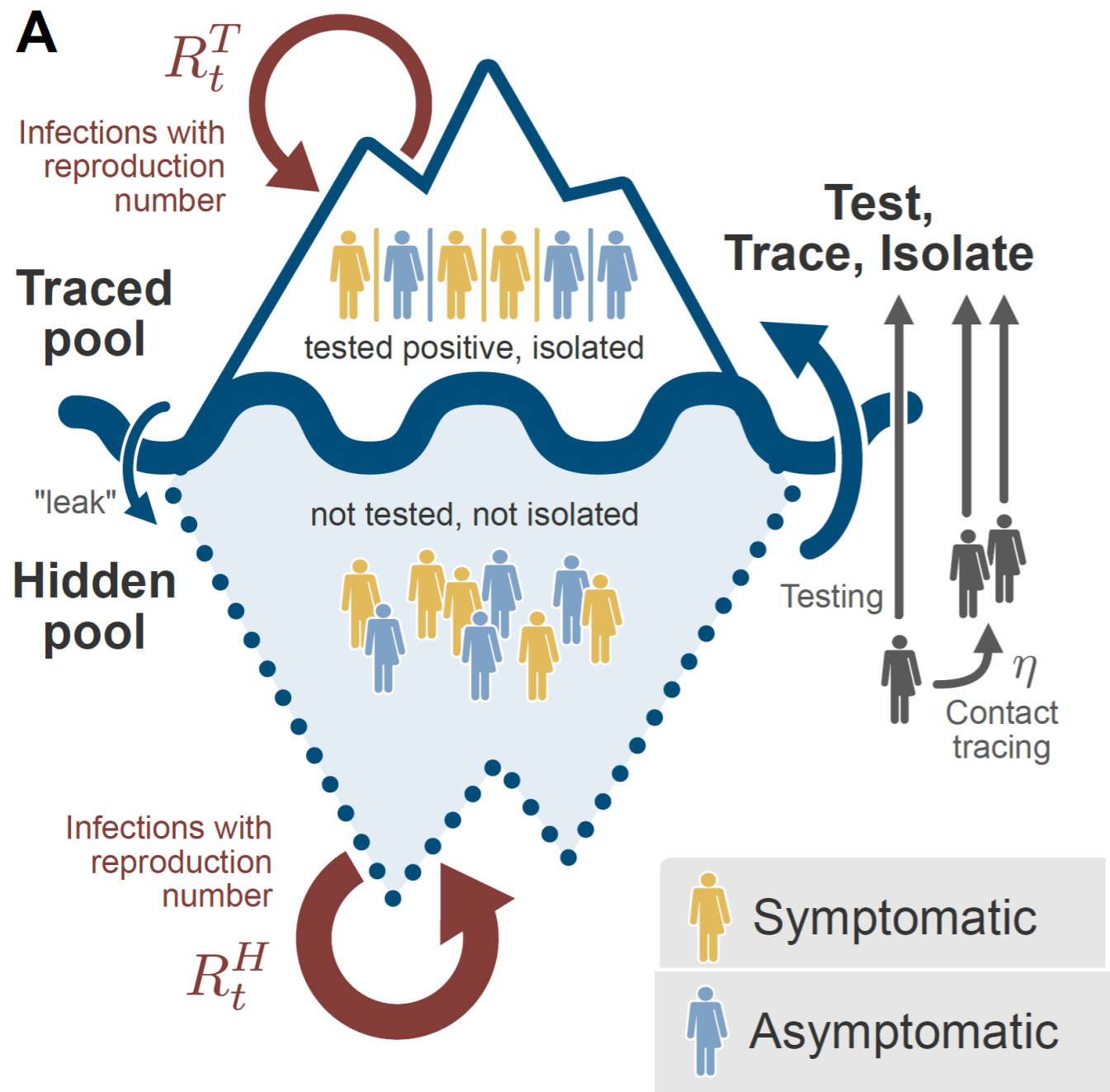


The challenges of containing SARS-CoV-2 via test-trace-and-isolate

Sebastian Contreras ^{1,2,5}, Jonas Dehning^{1,5}, Matthias Loidolt ^{1,5}, Johannes Zierenberg ¹, F. Paul Spitzner¹, Jorge H. Urrea-Quintero¹, Sebastian B. Mohr ¹, Michael Wilczek ^{1,3}, Michael Wibral⁴ & Viola Priesemann ^{1,3}✉

Without a cure, vaccine, or proven long-term immunity against SARS-CoV-2, test-trace-and-isolate (TTI) strategies present a promising tool to contain its spread. For any TTI strategy, however, mitigation is challenged by pre- and asymptomatic transmission, TTI-avoiders, and undetected spreaders, which strongly contribute to “hidden” infection chains. Here, we study a semi-analytical model and identify two tipping points between controlled and uncontrolled spread: (1) the behavior-driven reproduction number R_t^H of the hidden chains becomes too large to be compensated by the TTI capabilities, and (2) the number of new infections exceeds the tracing capacity. Both trigger a self-accelerating spread. We investigate how these tipping points depend on challenges like limited cooperation, missing contacts, and

Test-Trace-and-Isolate (TTI) contributes to containment



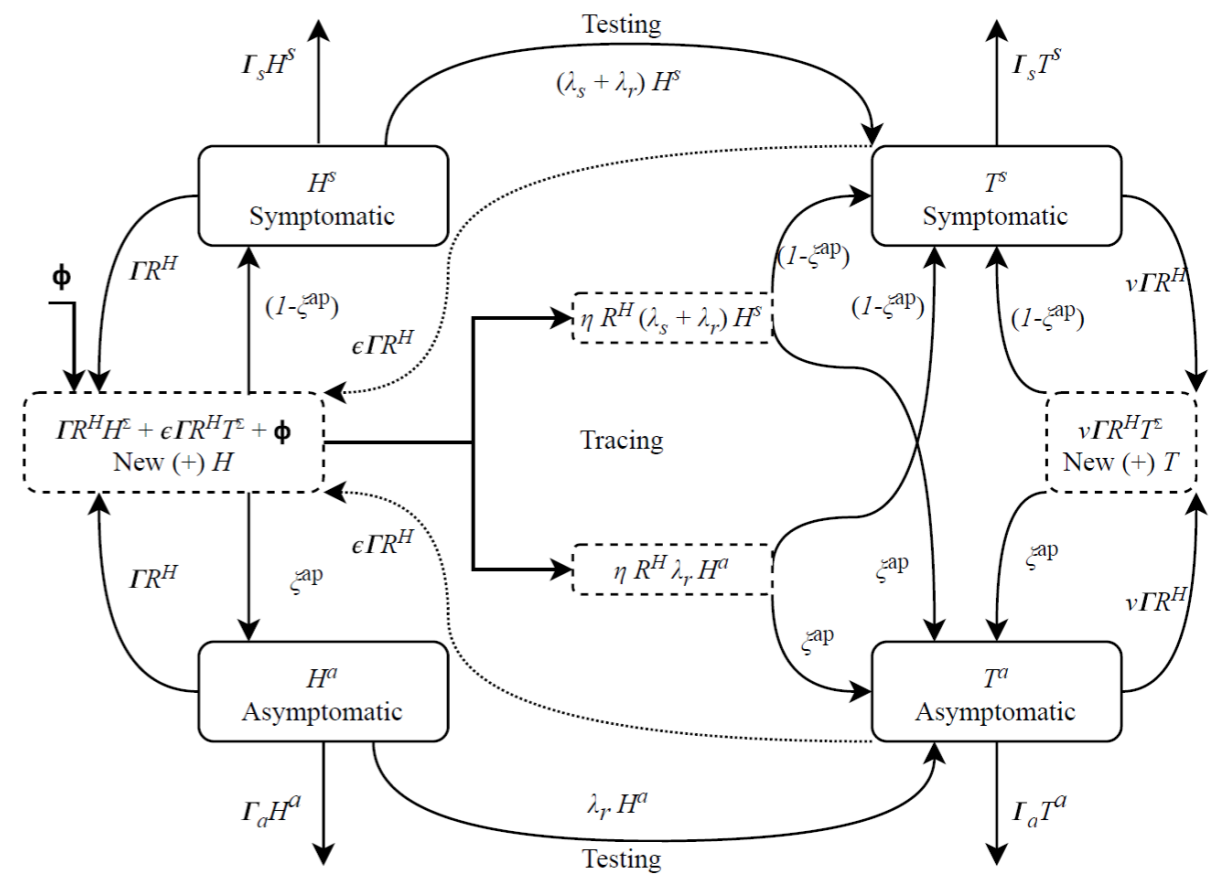
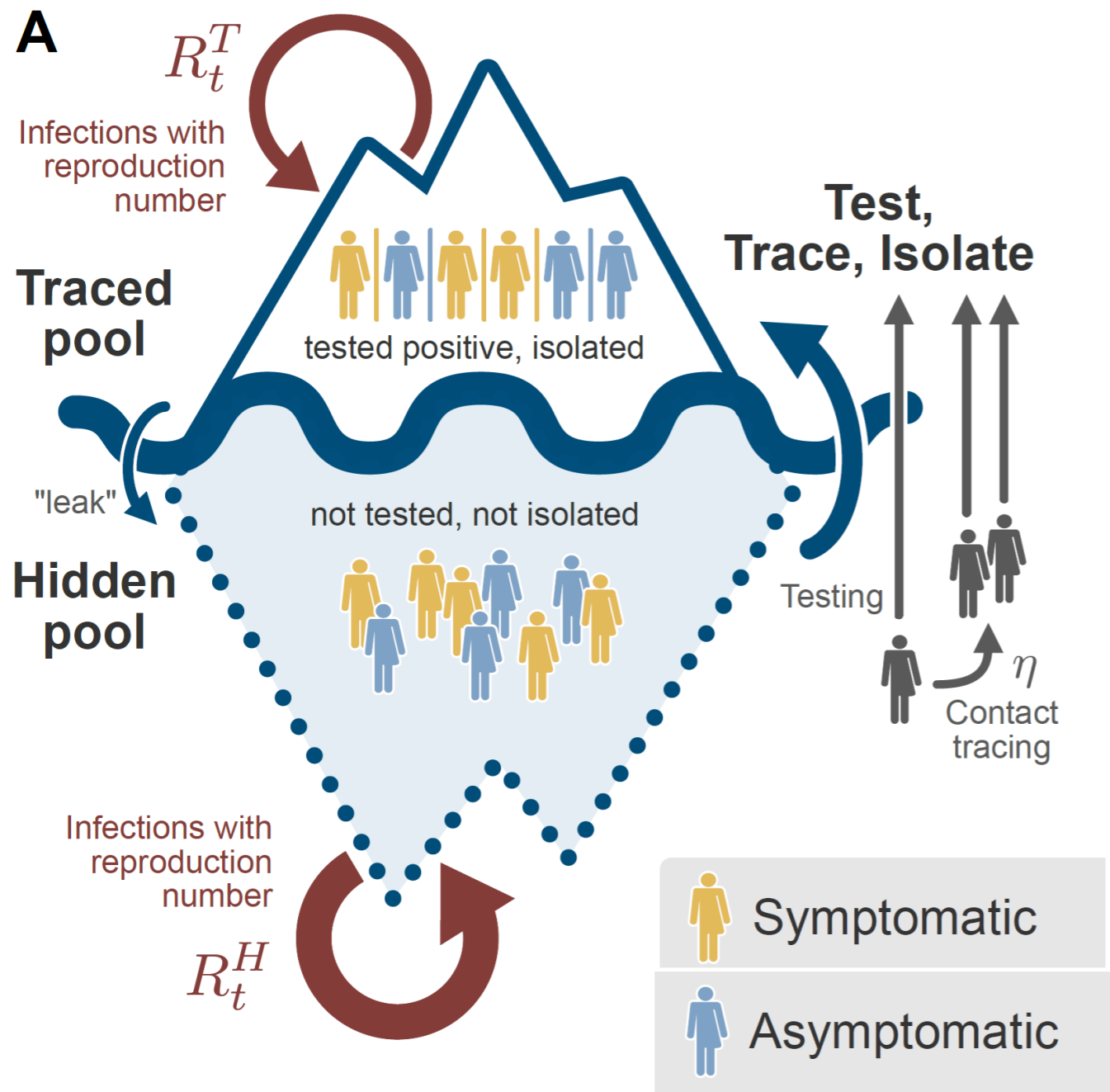
Test & isolated

- Random (0)
- Symptoms (50 % of sympt., on average after 5 days)
- Test contact persons

Contact tracing is difficult:

- Pre- und asymptomatic infection
- 1/3 of contacts are overlooked
- Quarantine is not perfect
- People who do not get tested (20%)
- Introduction of new infectious from abroad
- Limited capacities of health offices for testing and tracing

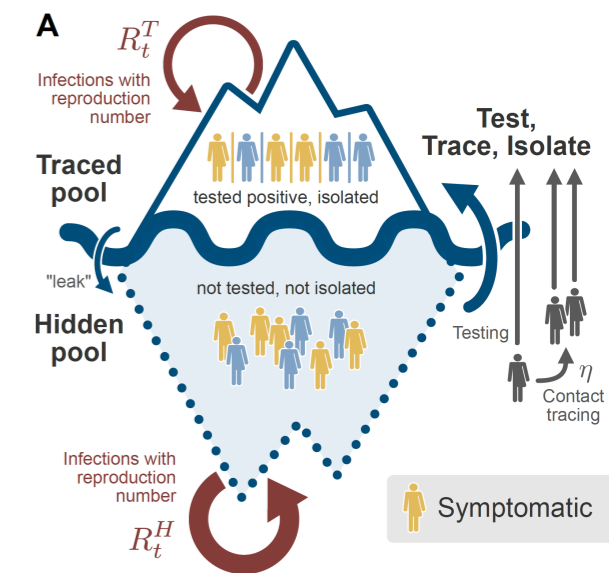
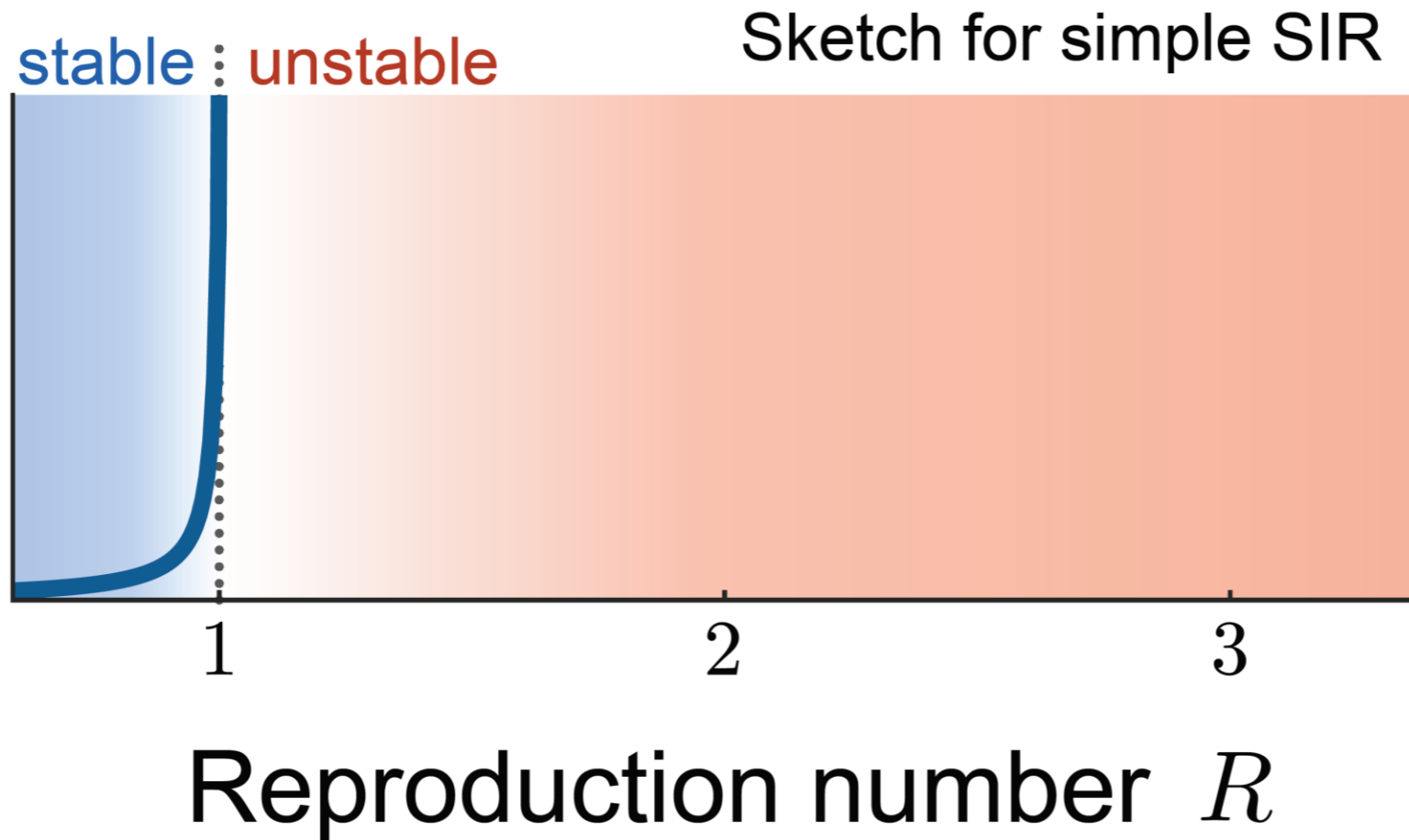
Test-Trace-and-Isolate (TTI) contributes to containment



The reproduction number R and the external influx of new cases Φ determine the level of new infections N

$$N \approx \frac{\Phi}{R_c - R} = \frac{\Phi}{1 - R}, \quad \text{für } R < 1$$

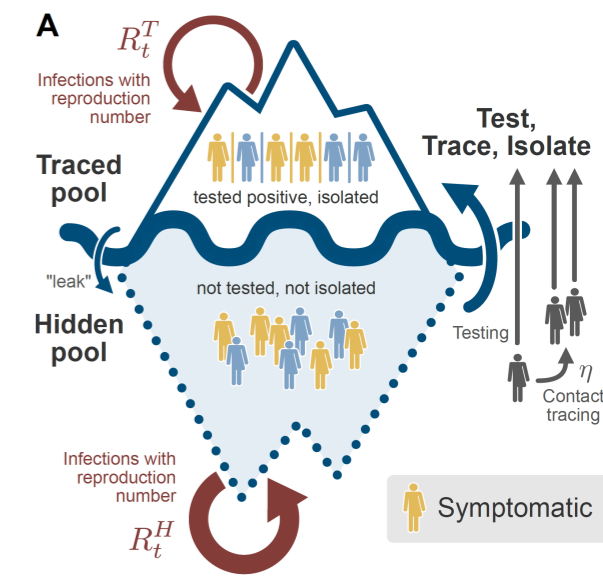
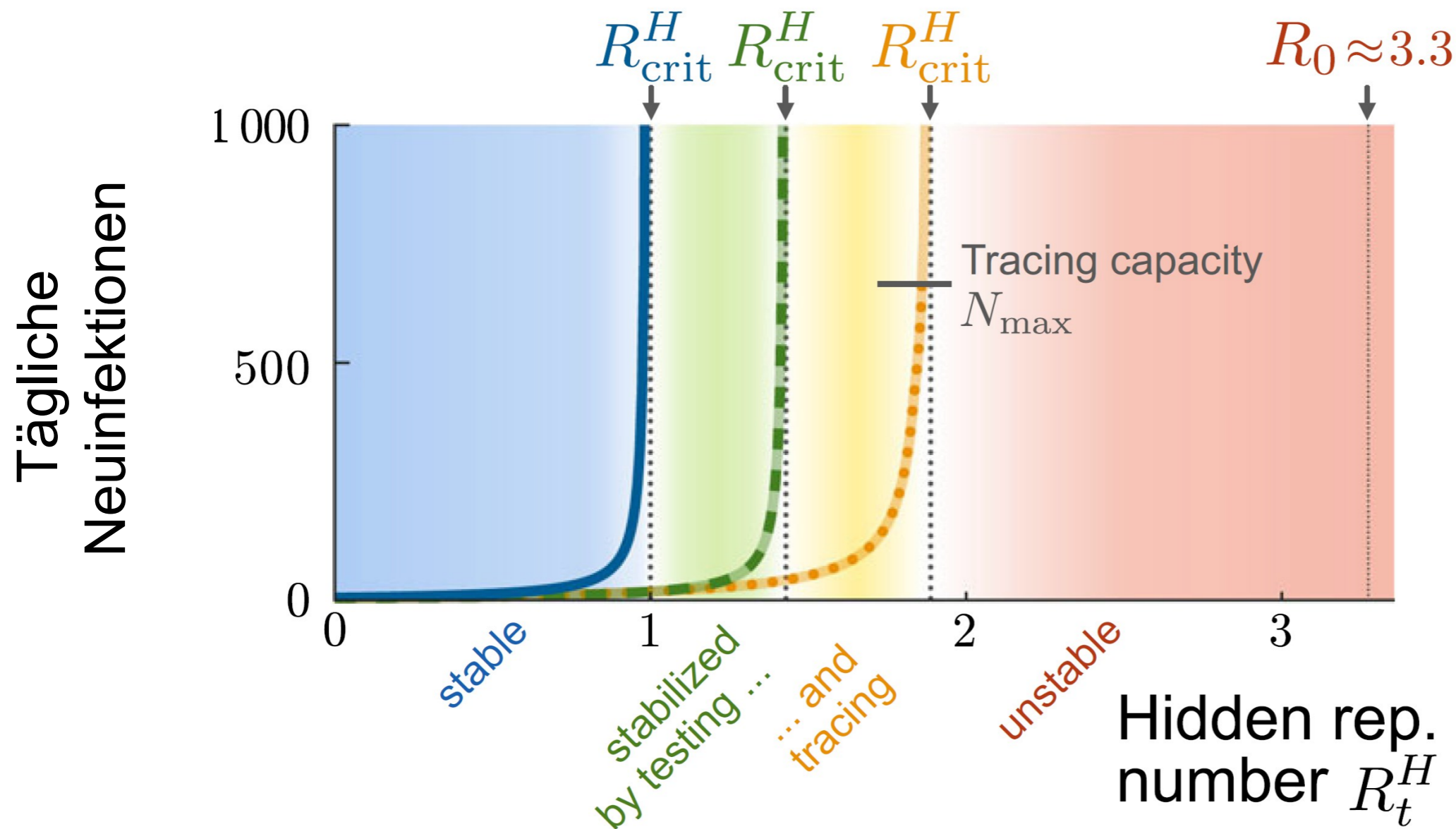
Observed new cases $\hat{N}_\infty^{\text{obs}}$



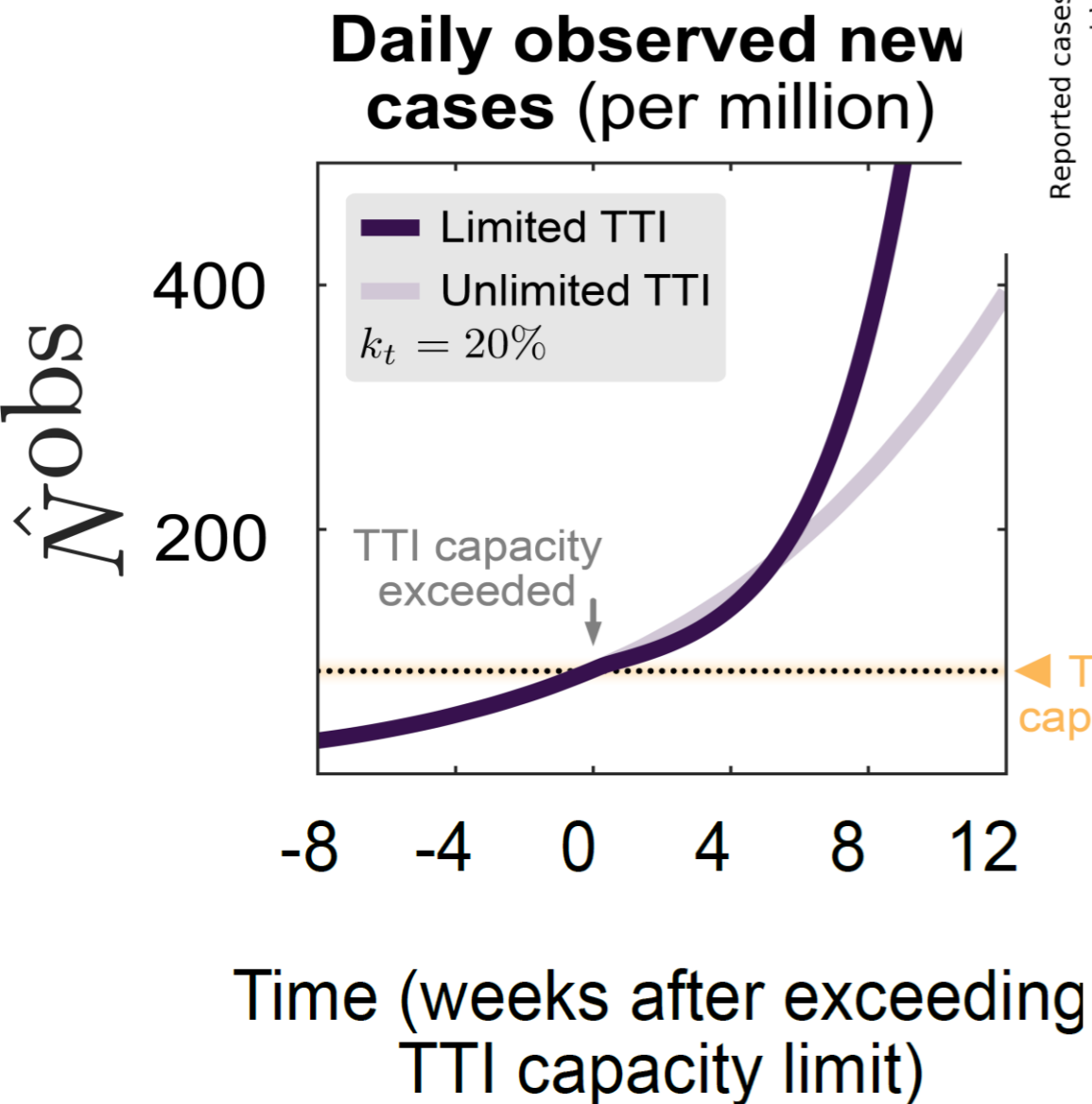
Test-Trace-Isolate (TTI) moves the stability limit:

Without TTI, R must be below 1

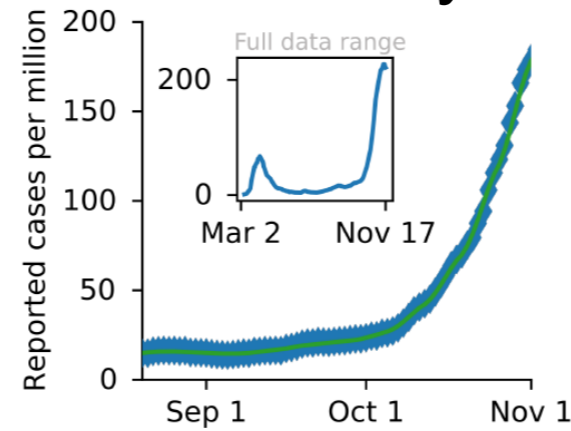
With TTI, R in the day to day life can be up to two



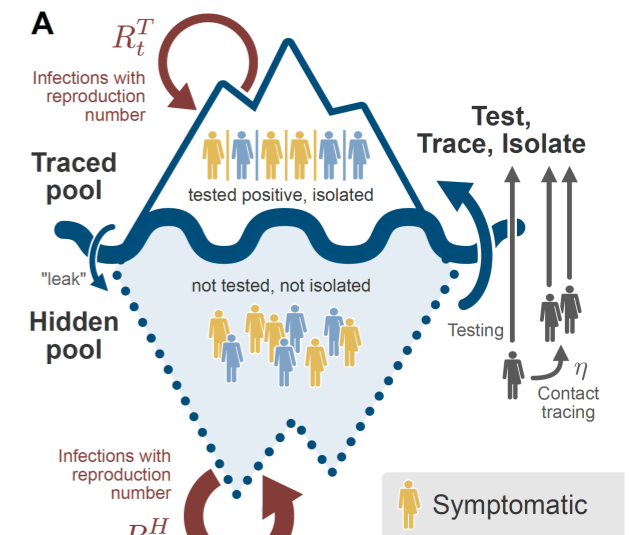
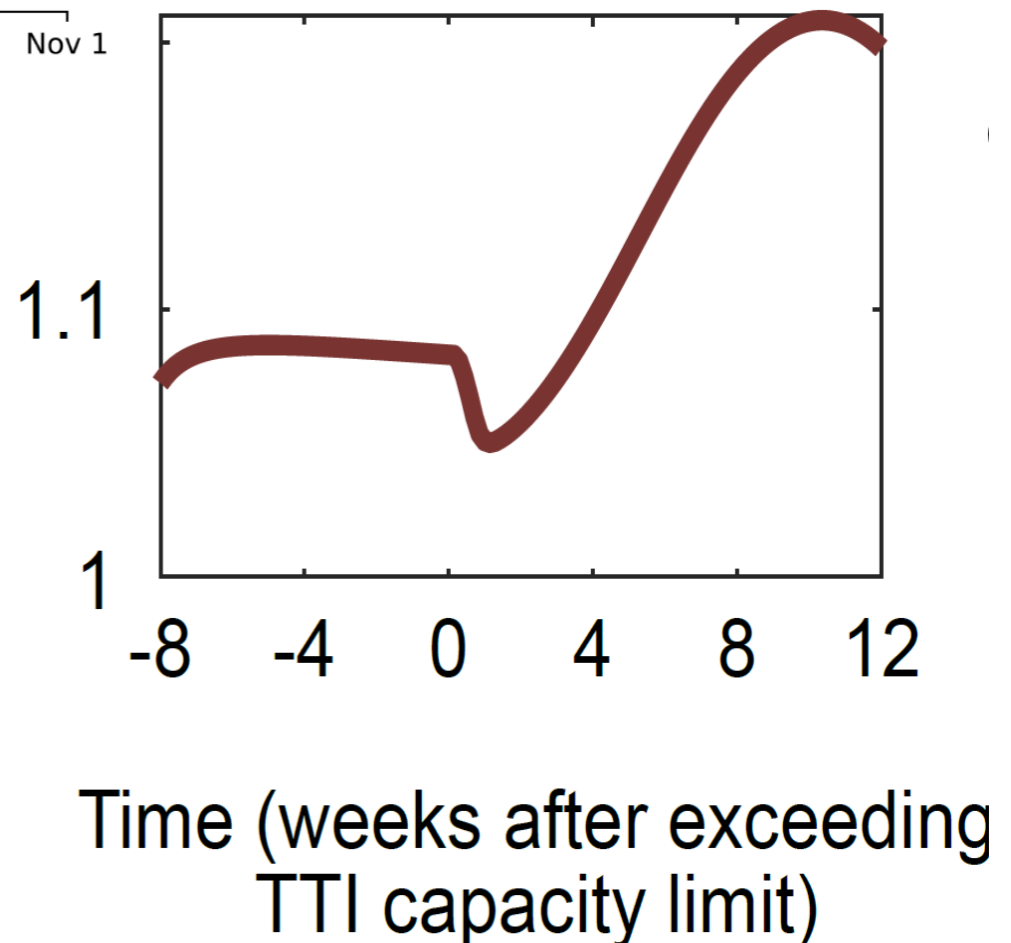
Crossing the TTI Limit: Case numbers grow faster than exponential



New infections Germany

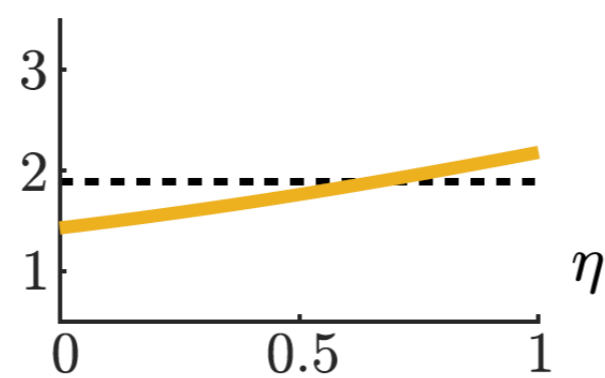
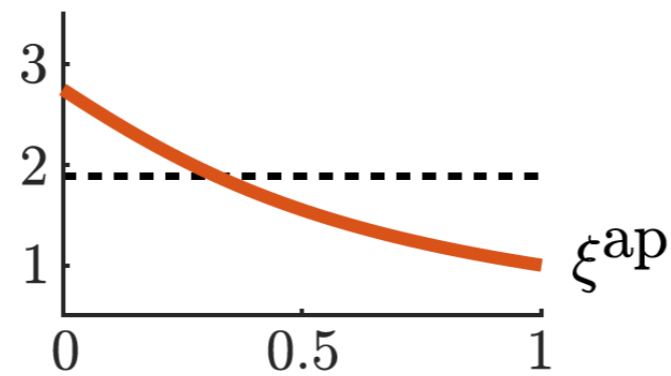
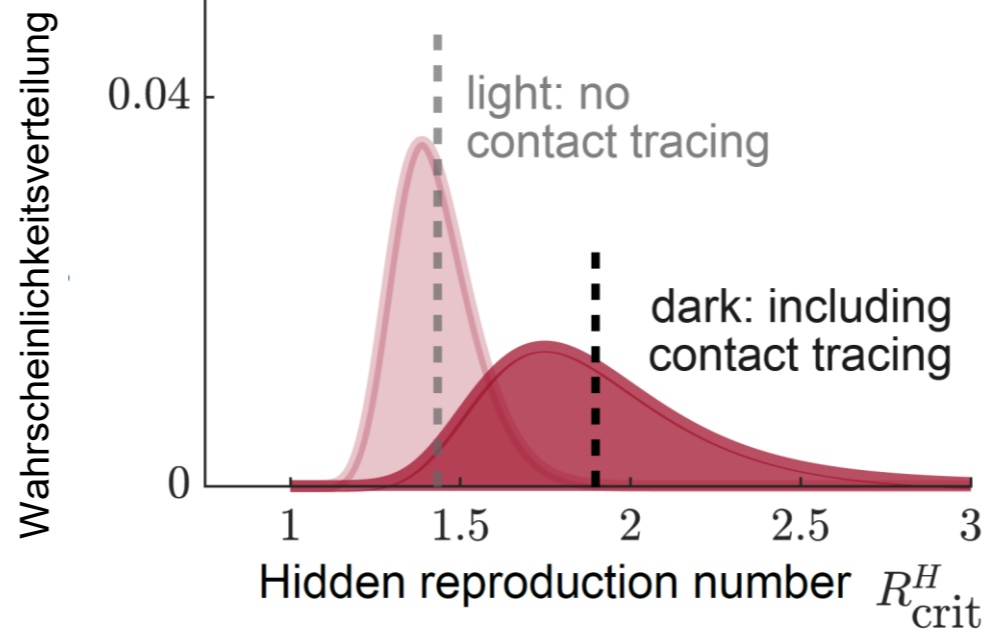
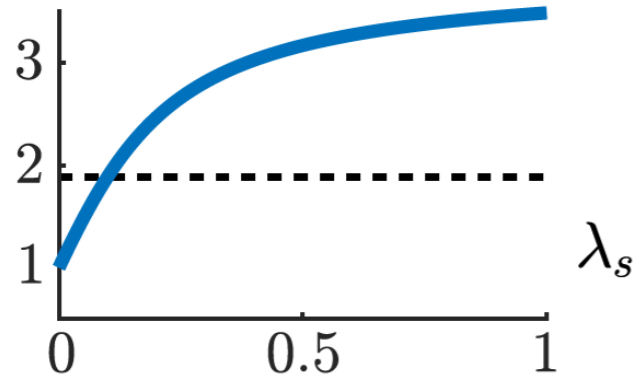
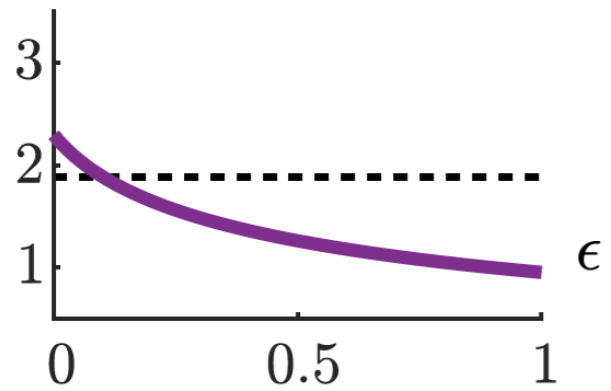
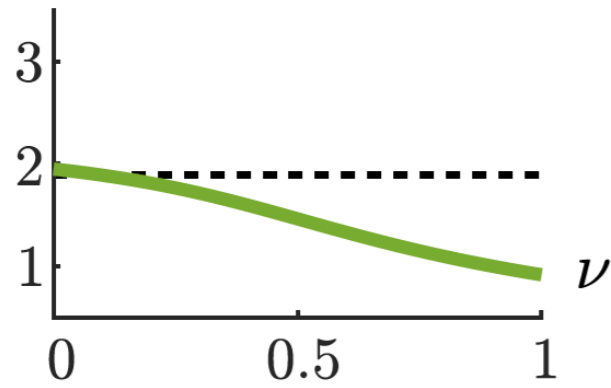


\hat{R}_t^{obs}

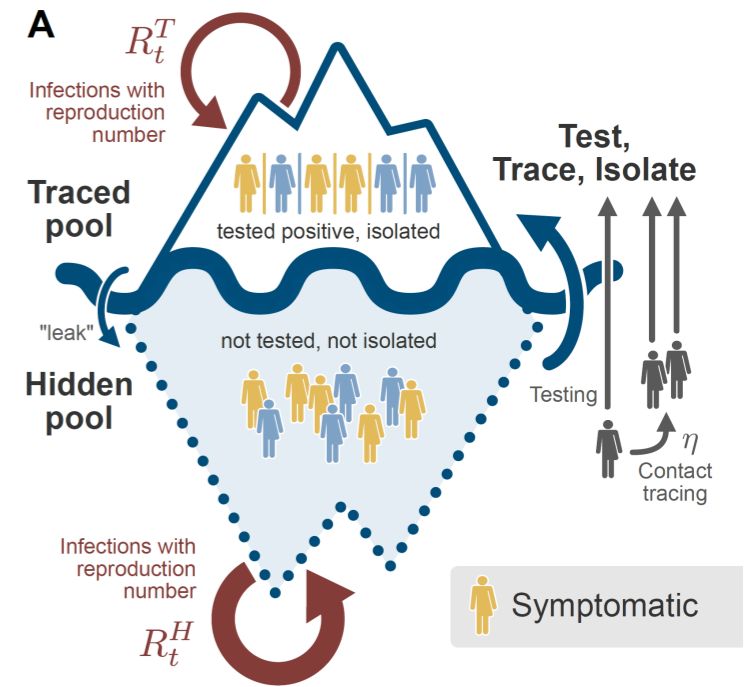


Sensitivity Analysis

Stabilitätsgrenze R_H

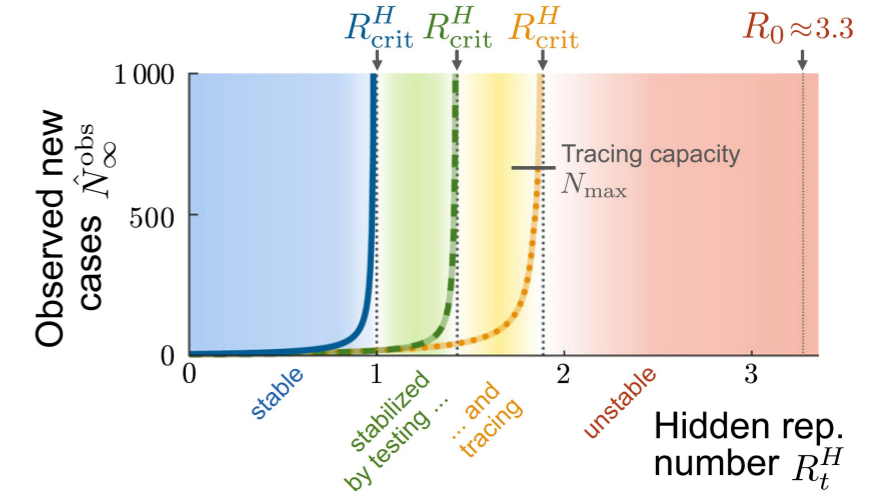
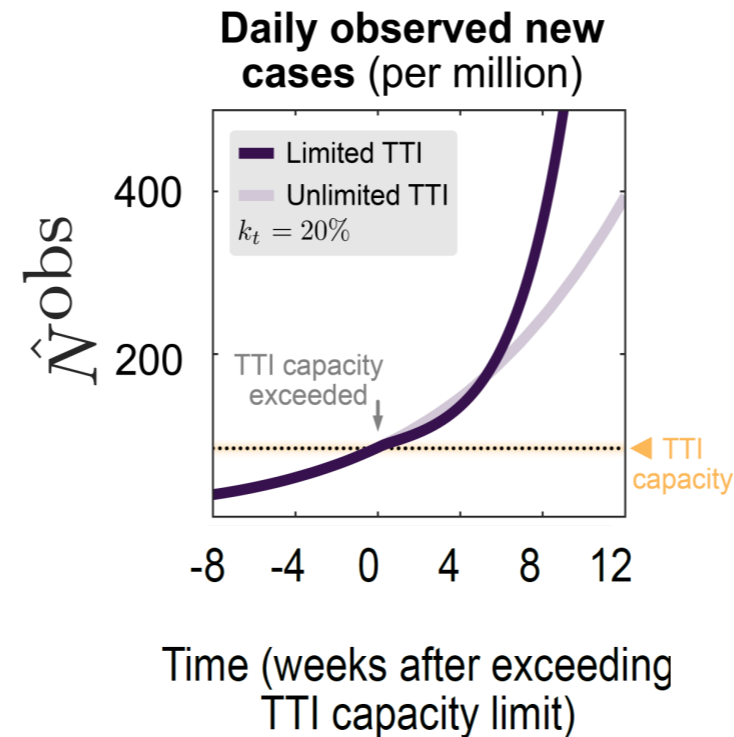
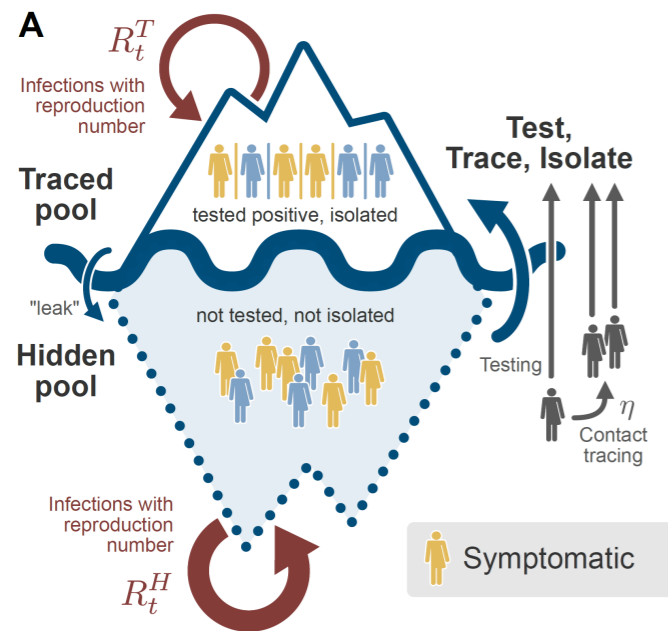


- ν Isolation factor
- ϵ "leak" factor
- λ_s Symptom-driven testing
- ξ^{ap} Apparent asymptomatic fraction
- η Tracing efficiency



Summary of the TTI strategy

Test-Trace-Isolate (TTI) contributes to containing COVID-19:



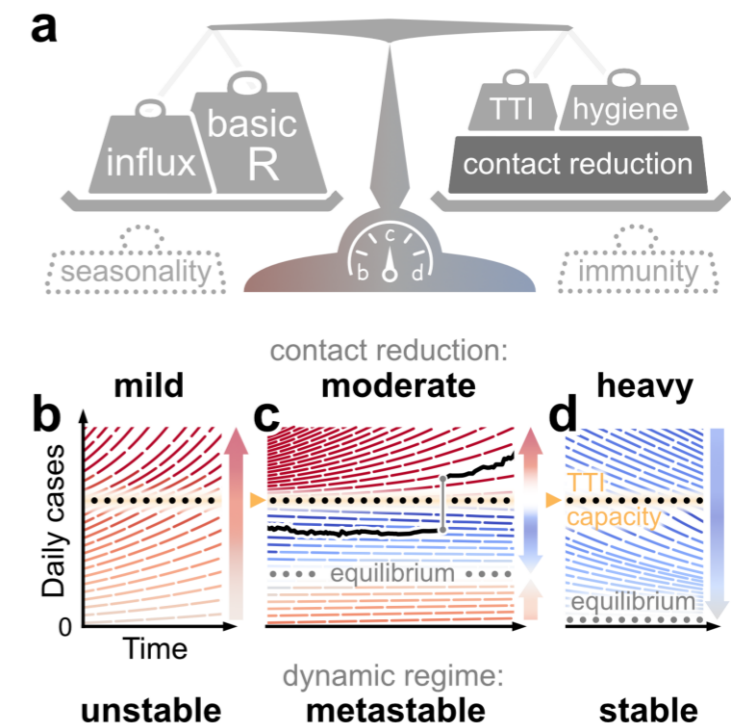
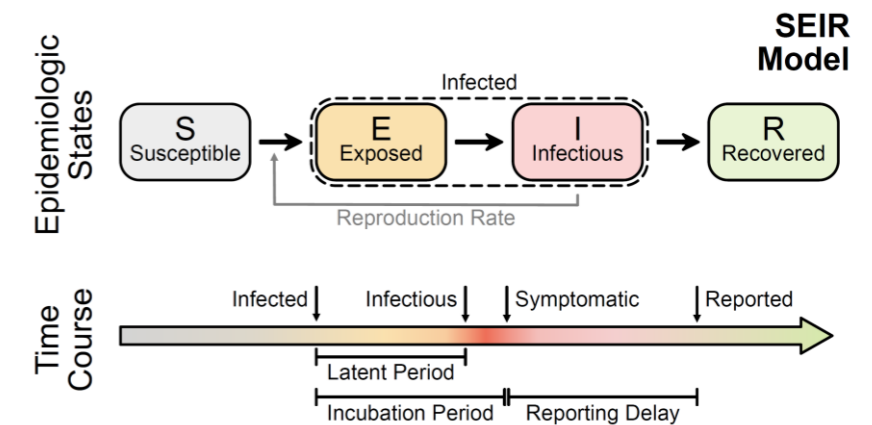
The undetected cases contribute most strongly to the spread

If the TTI capacity is surpassed, a tipping point is crossed, and growth self-accelerates.

TTI enables every single person to have more contacts: Instead of one, about two persons can be infected →
Compensaiton by TTI.

Content

- **Bayesian Inference and short-term forecast:**
SIR-Model, Hamiltonian MCMC, weekly change points
- **Derive effectiveness of measures:**
Contribution of test-trace-isolate (TTI) to containment of COVID-19
- **Progress of vaccination determines opening steps.**
How fast can we lift „non-pharmaceutical interventions“ (NPIs) given the planned vaccination progress?



Thank you!

Priesemann Group

Sebastian Contreras
Jonas Dehning
David Ehrlich
Daniel Gonzalez Marx
Benedikt Grüger
Kira Herff
Emil Iftekhar
Matthias Linden
Matthias Loidolt
Fabian Mikulasch
Sebastian Mohr
Valentin Neuhaus
Lucas Rudelt
Alexander Schmidt
Andreas Schneider
Julian Schulz
Paul Spitzner
Patrick Vogt
Johannes Zierenberg
+ you?



MAX-PLANCK-GESELLSCHAFT



SPP 2205
Evolutionary optimization
of neuronal processing



**Discussions on COVID within
the Göttingen Campus and beyond:**
Heike Bickeböller, Philip Bittihn, Eberhard
Bodenschatz, Wolfgang Brück, Alexander Ecker,
Andreas Leha, Theo Geisel, Ramin Golestanian,
Helmut Grubmüller, Stephan Herminghaus, Gerald
Haug, Reinhard Jahn, Jürgen Jost, Norbert Lossau,
Vladimir Zykov, Michael Meyer-Hermann, Iris Pigeot,
Simone Scheithauer, Anita Schöbel, Ferdi Schüth,
Michael Wibrál & Michael Wilczek

External PhD students (co-supervised)

Benjamin Cramer (U Heidelberg)
Madhura Ketkar (ENI Göttingen)
Corentin Nelias (MPI-DS)

Alumni

Victor Brasch (EPFL)
Henrik von der Emde (Cambridge)
Jan Geisler (Max Planck School)
Jorge de Heuvel (U Mainz)
Annika Hagemann (Bosch)
Helge Heuer (U Göttingen)
Leonhard Leppin (MPI Garching)

Moritz Layer (Cambridge)
Joao Neto (MPI-DS)
Bruno del Papa (MERK)
Bettina Royen (Max Planck School)
Mathias Sogorski (PSI, Berlin)
Jens Wilting (Bosch)

Referenzen:
[Dehning et al., Science, May 2020]
[Dehning et al., medRxiv 2020]
[Contreras et al., Nature Commun (2021) / arXiv:2009.05732]
[Contreras et al., arXiv:2011.11413]
[Alwan et al., The Lancet 2020]
[Linden et al., Dt. Aerzteblatt Int. / arXiv:2010.05850]
[Priesemann et al., The Lancet, 2021]
[Priesemann et al., The Lancet, in press]

www.containcovid-pan.eu