

Online Supplementary Document

Table S1. Parameters and initial values for the compartmental model.

Age group (i)	Age (years)
1	0–19
2	20–39
3	40–59
4	≥60

Vaccination coverage scenario	Vaccination coverage (12–39 years, 40–59 years, ≥60 years)	Description	Source
90% coverage	90%, 90%, 90%	Assuming 90% of eligible people got vaccinated	–
High coverage	75%, 80%, 90%	Corresponding to the sum of people who are willing to be vaccinated and those who cannot decide yet	(1)
Intermediate coverage	60%, 70%, 85%	Intermediate scenario between High and Low coverages	(1)
Low coverage	45%, 60%, 80%	Corresponding to a proportion of people who are willing to be vaccinated	(1)
No vaccination	0%, 0%, 0%	Scenario without vaccination	–

Parameter	Description	Value	Source
Basic reproduction number	Mean number of secondary cases produced by a single infector in a completely susceptible population	2.5, 3.5, 5.0, 7.5	(2–4)
$R_{i,j}$	Reproduction number from age group j to age group i (Maximal eigenvalue of the matrix $R_{i,j}$ corresponds to a basic reproduction number)	Supplementary Figure 2	*1
N_1	Population of age group 1	23,000,000 (0–11 years, 130,000; 12–19 years, 100,000)*2	Census
N_2	Population of age group 2	32,500,000	Census
N_3	Population of age group 3	33,500,000	Census
N_4	Population of age group 4	37,000,000	Census
$1/\gamma$	Mean generation time (days)	5	(5)
$1/\gamma + 1/\epsilon$	Mean hospitalization period of severe cases (days)	19	*1
δ_1	Severity rate for age group 1	0.03%	*3*4
δ_2	Severity rate for age group 2	0.06%	(6)*3
δ_3	Severity rate for age group 3	1.0%	(6)*3
δ_4	Severity rate for age group 4	8.5%	(6)*3
$\delta_1 \times \mu_1$	Fatality rate for age group 1	0.005%	*3*4
$\delta_2 \times \mu_2$	Fatality rate for age group 2	0.01%	(6)*3
$\delta_3 \times \mu_3$	Fatality rate for age group 3	0.2%	(6)*3
$\delta_4 \times \mu_4$	Fatality rate for age group 4	5.7%	(6)*3
E_a	Vaccine efficacy to prevent infection	90% (very effective), 70% (effective)	(7–15)
$1 - (1 - E_a) \times E_b$	Vaccine efficacy to prevent severe illness and death	95% (very effective), 90% (effective)	(7–16)
$1 - E_c$	Vaccine efficacy to reduce infectivity from patients with a breakthrough infection	50% (very effective), 25% (effective)	(17,18)*5

Compartment	Description	Initial value
S_{ni}	Unvaccinated people in age group i	$N_i \times 0.99 \times (1 - \text{vaccination coverage for age group } i)$
S_{vi}	People vaccinated but not protected in age group i	$N_i \times 0.99 \times (\text{vaccination coverage for age group } i) \times (1 - E_a)$
V_i	People protected by vaccination in age group i	$N_i \times 0.99 \times (\text{vaccination coverage for age group } i) \times E_a$
I_{ni}	Infected (infectious) people among those unvaccinated in age group i	100
I_{vi}	Infected (infectious) people among those vaccinated in age group i	0
C_i	Infected people in age group i with severe illness requiring mechanical ventilation, intensive care unit admission, or extracorporeal membrane oxygenation	0
R_i	Recovered people in age group i (assuming 1% of the total population had been already infected in Japan)	$N_i \times 0.01$
D_i	Dead people due to COVID-19 in age group i	0

*¹ Determined using empirical data from Japan.

*² In age group 1, only those aged 12–19 years are eligible for vaccination.

*³ Data are from 2020. The change in pathogenicity by the Delta variant is probable (19,20), but that was not considered in the simulation due to the lack of data in Japan.

*⁴ Due to the small number of COVID-19 cases for age group 1 in Japan, we could not estimate the severity and fatality rates for the group. We assumed the rates to be half of those for age group 2.

*⁵ Although the vaccine efficacy to reduce infectivity from patients with a breakthrough infection for the Delta variant might decline (21,22), the efficacy seems still maintained to some degree (23,24). We assumed the efficacy to be half of that for the very effective scenario.

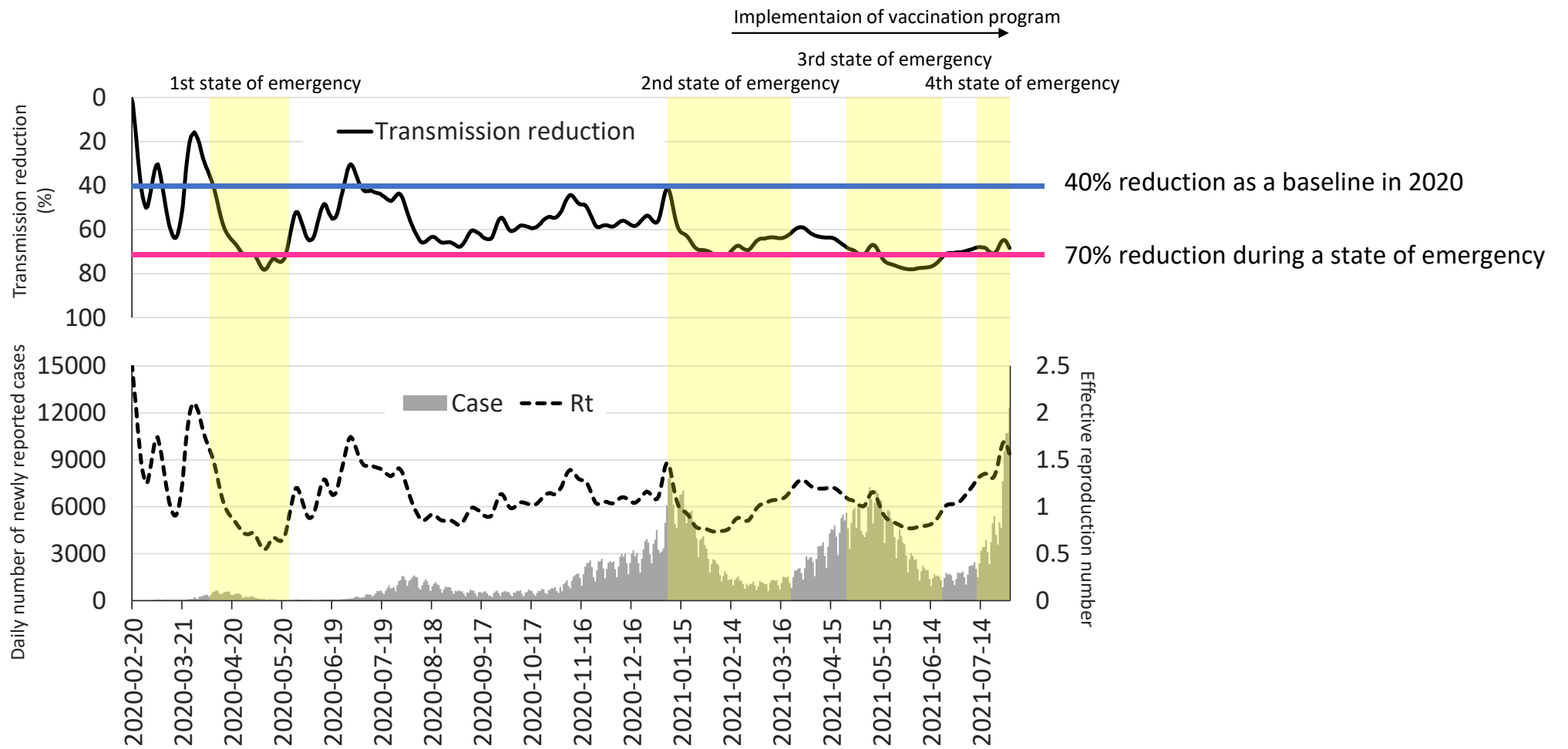


Figure S1. Transmission dynamics of COVID-19 and transmission reduction in Japan from 20 February 2020 to 31 July 2021.

The daily numbers of confirmed COVID-19 cases were retrieved from the website of the Ministry of Health, Labour and Welfare. The effective reproduction number (R_t) was calculated using EpiEstim with the parameter of a serial interval set to 5 days (5,25). Reduction in transmission was calculated as follows: $(1 - \text{Effective reproduction number} / \text{Basic reproduction number})$. The basic reproduction was set to 2.5 (original strain), 3.5 (Alpha variant), or 5.0 (Delta variant) (2–4). The transitions from the original strain to the Alpha variant and from the Alpha variant to the Delta variant were considered to have occurred in February–April 2021 and July 2021, respectively (26). Shaded areas in yellow indicate periods under a state of emergency. Note that transmission reduction was achieved by nonpharmaceutical interventions plus the effect of vaccination after February 2021.

A

Homogeneous transmission

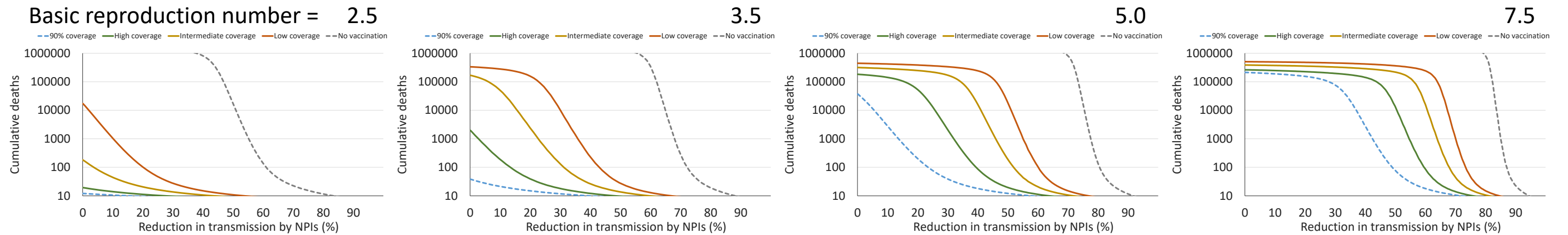
≥60Y (age group 4)	1	1	1	1
40–59Y (age group 2)	1	1	1	1
20–39Y (age group 2)	1	1	1	1
0–19Y (age group 1)	1	1	1	1
To (i) From (j)	0–19Y (age group 1)	20–39Y (age group 2)	40–59Y (age group 3)	≥60Y (age group 4)

Heterogeneous transmission

≥60Y (age group 4)	1	4	4	4
40–59Y (age group 2)	1	4	16	1
20–39Y (age group 2)	1	16	4	1
0–19Y (age group 1)	4	4	1	1
To (i) From (j)	0–19Y (age group 1)	20–39Y (age group 2)	40–59Y (age group 3)	≥60Y (age group 4)

B

Homogeneous transmission with very effective vaccines



C

Homogeneous transmission with effective vaccines

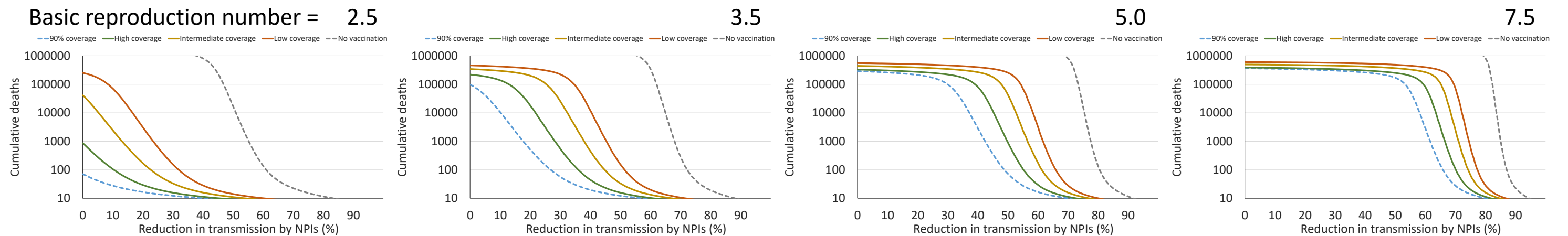


Figure S2. Cumulative numbers of COVID-19 deaths by simulation with a homogeneous model.

(A) Relative values of reproduction numbers from age group j to age group i are shown for the homogeneous and heterogeneous models. In the heterogeneous model, we assumed that transmissions from people aged 20–59 years and transmissions within the same age group were four times higher than the other transmission pairs. Four-fold higher transmissions within the same age group were observed in empirical data from Japan. Four-fold higher transmissions

from those aged 20–59 years were determined in this study as we validated that the coefficient of 4 in our simulation fitted well with the observation data that ~70% of all infected people in Japan were aged 20–59 years before the implementation of the vaccination program. The numbers in the matrices are relative values, and they were proportionally adjusted so that the maximal eigenvalue of the matrix corresponds to a basic reproduction number. (B, C) Cumulative numbers of deaths by vaccination coverage, basic reproduction number, and vaccine efficacy in the simulation for 150 days, assuming a population with homogeneous transmission patterns, are shown. Panel B is for very effective vaccines, and Panel C is for effective vaccines. The x-axis denotes the degree of transmission reduction due to nonpharmaceutical intervention restrictions. The y-axis is in a logarithmic scale. Results for the heterogeneous model are shown in Figure 2.

Supplementary references

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