

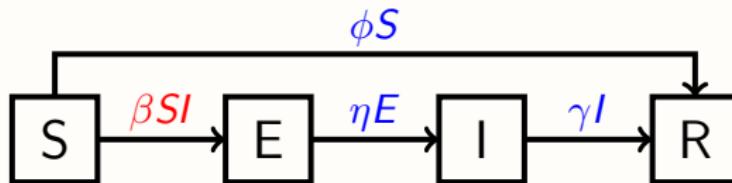
# Modeling Vaccination for a Novel Disease

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## 0. Standard Vaccination Model



$$S' = -\beta SI - \phi S, \quad S(0) = S_0;$$

$$E' = \beta SI - \eta E, \quad E(0) = E_0;$$

$$I' = \eta E - \gamma I, \quad I(0) = I_0.$$

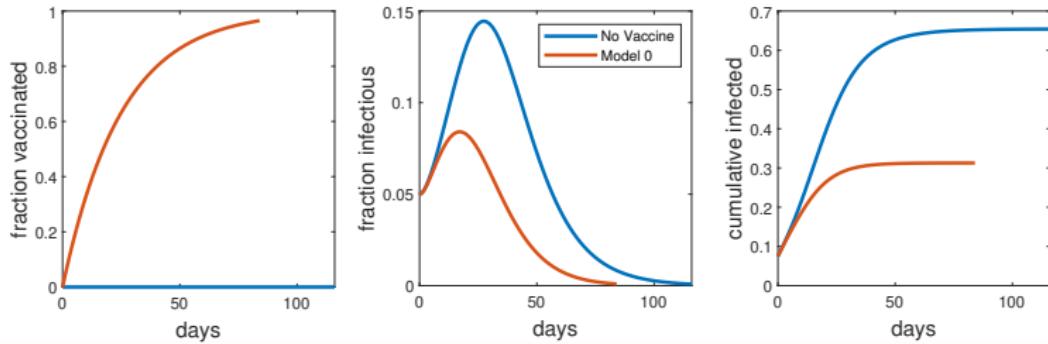
- ▶ Blue processes are instantaneous transitions.
  - Times are exponentially distributed.

## Disclaimer

- ▶ The SEIR model is a little too simple for COVID-19.
- ▶ Nevertheless, we'll use parameters that roughly match COVID-19 in January 2021:
  - 5-day mean incubation
  - 10-day mean duration of infectivity
  - About 30% initial immunity from prior infection
  - Effective basic reproductive number for the delta variant (with masks and social distancing) roughly 2.5 to 4.0, depending on the community.
- ▶ The graphs for a COVID-19 model would be similar.

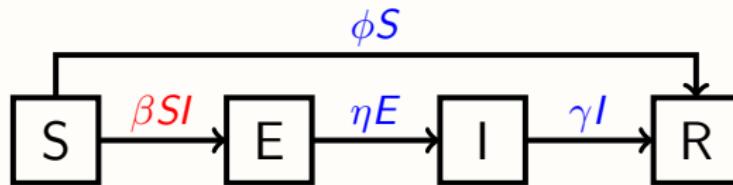
## 0. Vaccination Impact in Standard Model

- ▶  $\phi = 0.04$  — mean time for vaccination is 25 days.
- ▶ Effective reproductive number 4.0, corresponding to delta with limited mask use and social distancing.



- ▶ Vaccination looks really effective! But...

## 0. Standard Vaccination Model

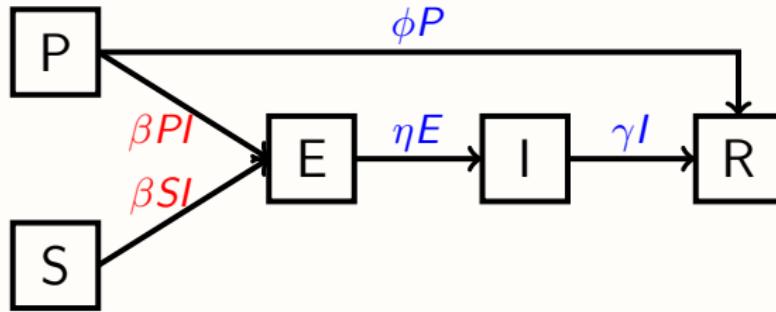


► What is wrong with this model?

1. Nobody refuses the vaccine.
2. Distribution is instantaneous.
3. Supply is unlimited.
4. Vaccine always confers immunity.

## 1. Vaccine Refusal

- ▶ Partition the susceptible class into a vaccine refuser group ( $S$ ) and a prevaccination group ( $P$ ).

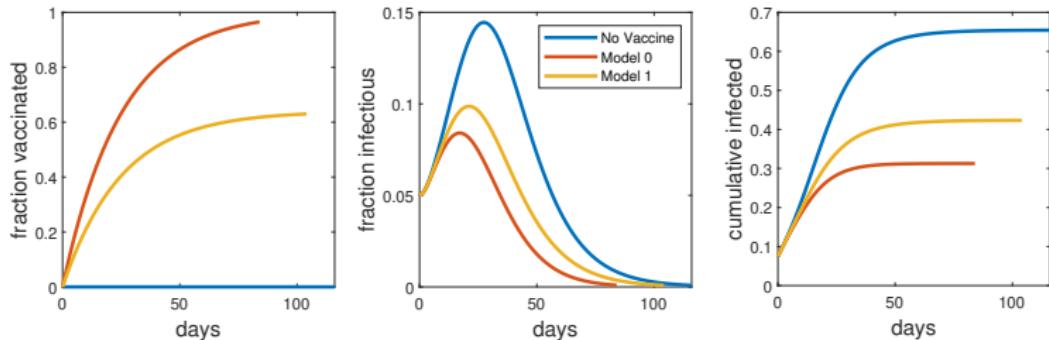


- ▶ Assume the time zero susceptible fraction is  $S_0$  and that a fraction  $r$  of these are vaccine refusers.

$$S(0) = rS_0, \quad P(0) = (1 - r)S_0.$$

## 1. Vaccine Refusal Impact

- Take  $r = 0.36$  for the US average (near 0.1 in Portugal and Chile, and only because the vaccine is age-limited)

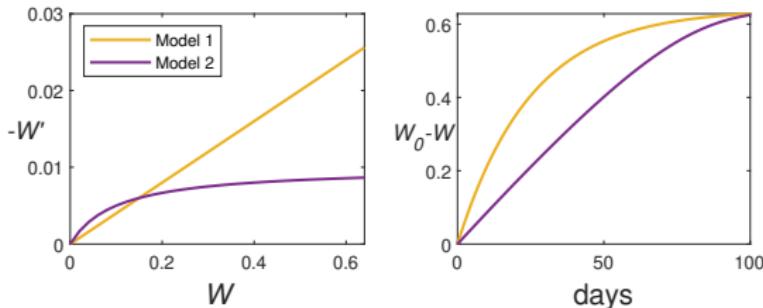


- Vaccine refusal makes a big difference, but....
- The vaccination rate should not depend on  $r$  until prevaccinated people are largely done — we need a better distribution model.

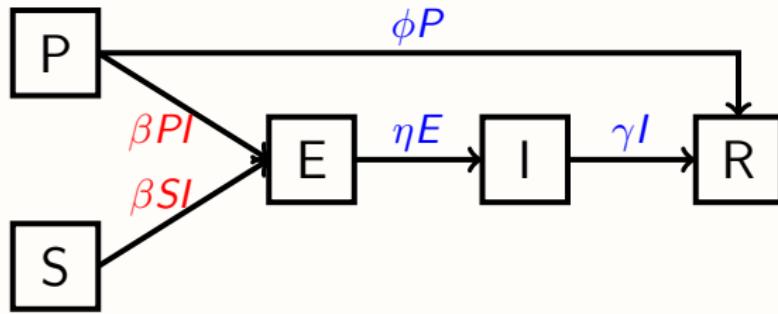
## A Vaccination Model with Limited Distribution

- ▶ Let  $W$  be the fraction of people who want vaccination.
- ▶ Finding recipients for a vaccinator is like finding substrate for a biochemical enzyme (Michaelis-Menten kinetics).
  - $K$  is the level of  $W$  for which the rate is half of the maximum.
  - $V$  is a vaccination rate constant (1/time).

$$W' = -\frac{V KW}{K + W} \equiv -\phi W, \quad W(0) = W_0 = 1 - r.$$



## 2. Limited Distribution Model



$$P' = -\beta PI - \phi(W)P, \quad P(0) = (1 - r)S_0;$$

$$S' = -\beta SI, \quad S(0) = rS_0;$$

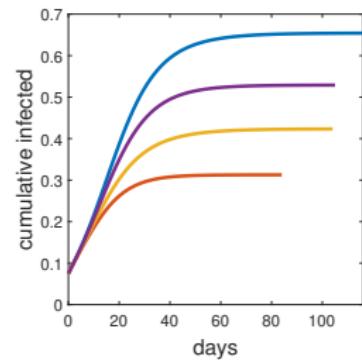
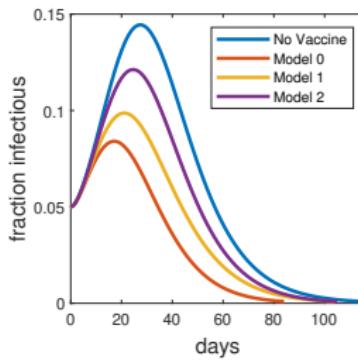
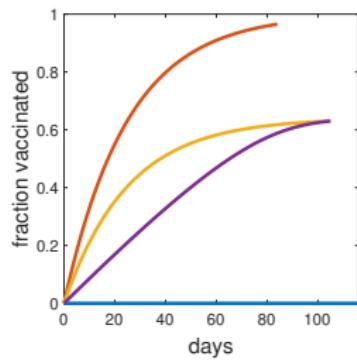
$$E' = \beta(P + S)I - \eta E, \quad E(0) = E_0;$$

$$I' = \eta E - \gamma I, \quad I(0) = I_0;$$

$$W' = -\phi(W)W, \quad \phi(W) = \frac{VK}{K + W}, \quad W(0) = 1 - r.$$

## 2. Impact of Limited Distribution

- ▶  $V = 0.1, K = 0.1$  — max vax rate is 1% per day

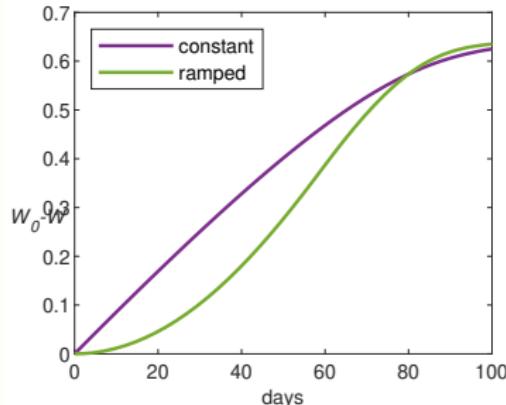


- ▶ Slow distribution significantly decreases vaccine impact.

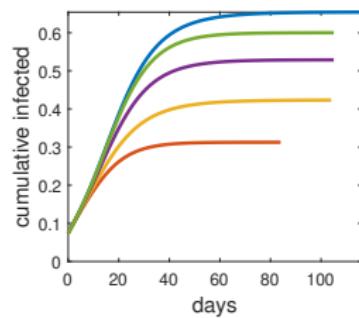
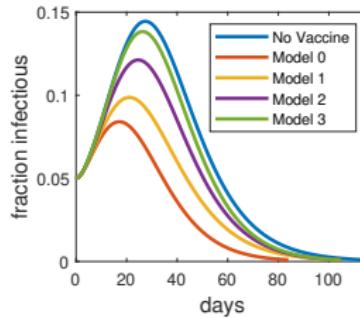
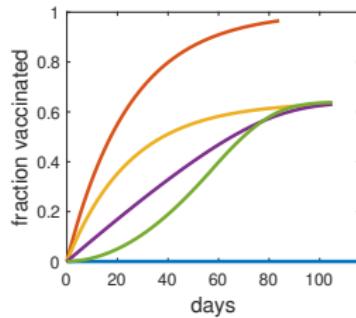
### 3. Limited Supply

- ▶ Vaccine production capacity increases over time.
  - The maximum vaccination rate should ramp up to a maximum.

$$V = \min \left( \frac{V_1 t}{t_1}, V_1 \right).$$

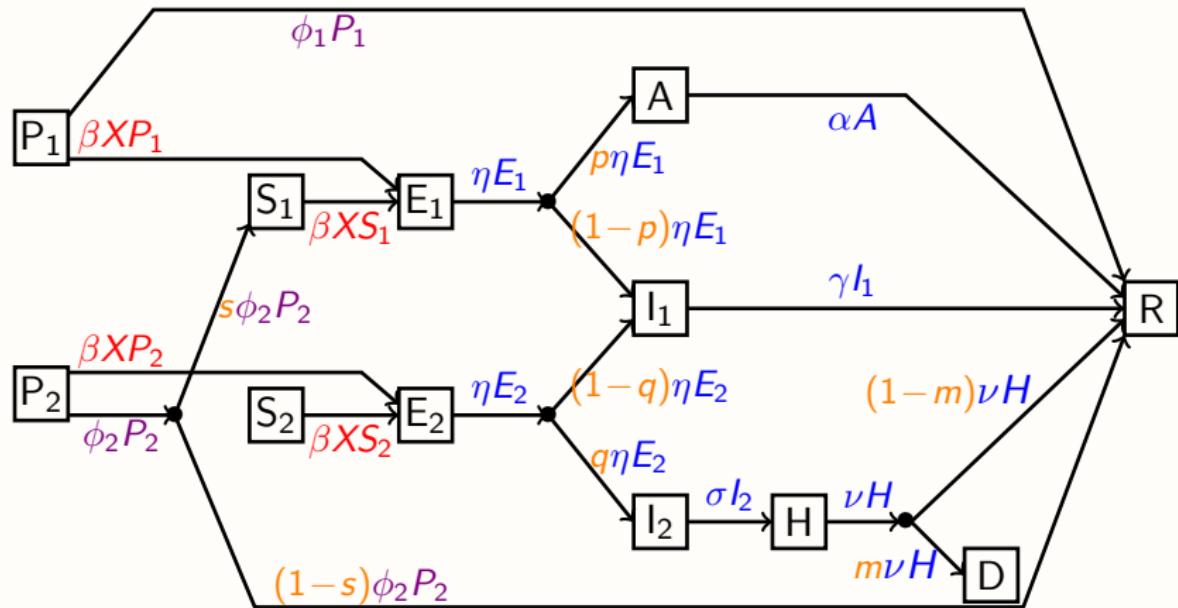


### 3. Impact of Limited Supply



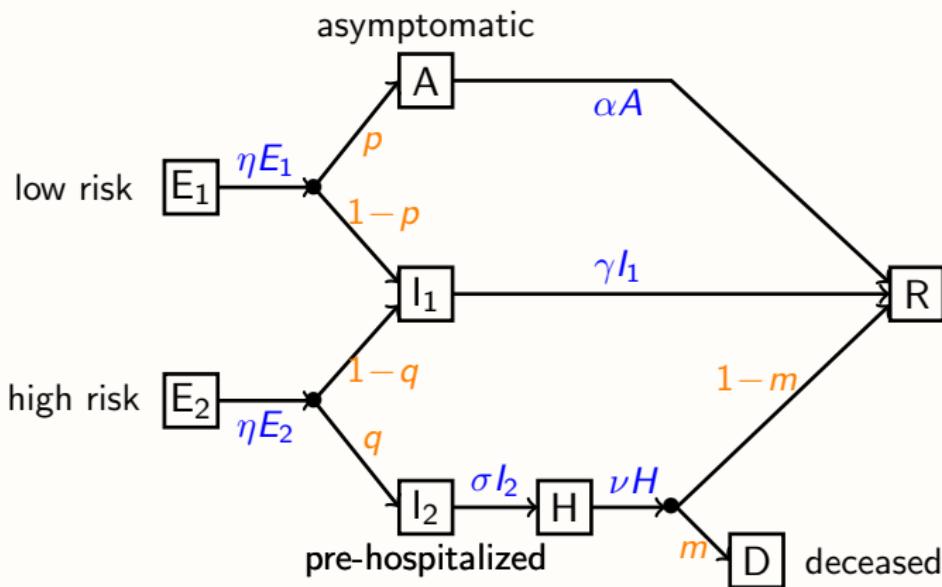
- ▶ If we want meaningful results, we need to use a realistic vaccination model.
- ▶ This scenario assumed a quick relaxing of mitigation strategies (think “Florida”).
  - **Mitigation must be maintained during vaccination.**

# The PSEAIHRD Model (COVID-19, January 2021)

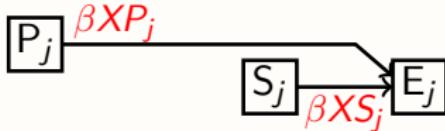


transitions, transmissions, vaccination, probabilities

## The PSEAIHRD Model – Transitions



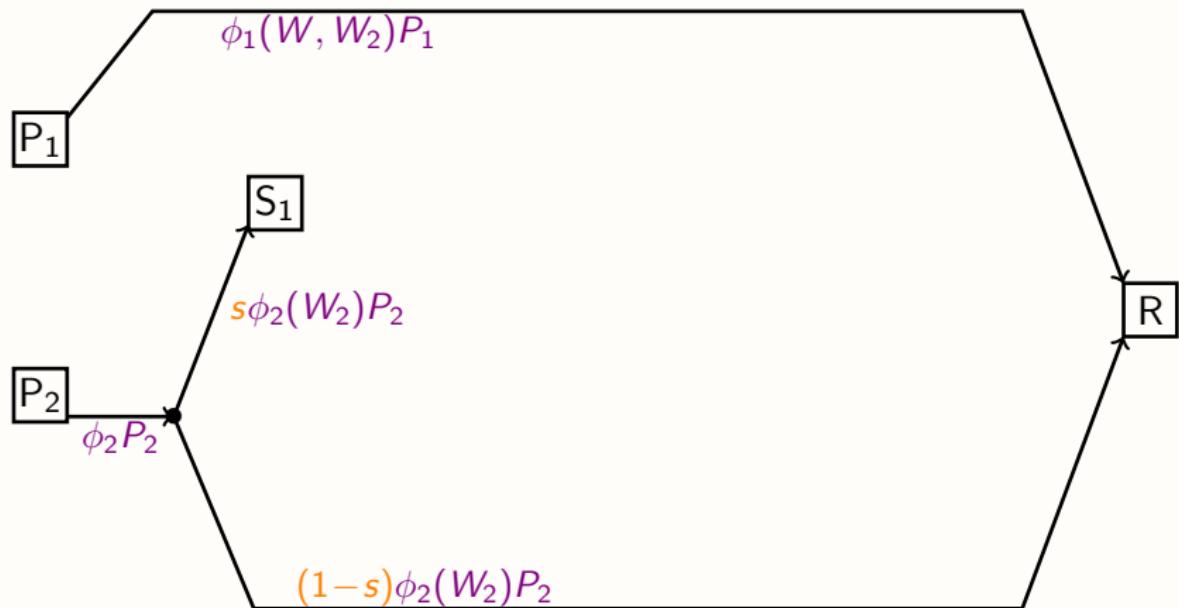
## The PSEAIHRD Model – Transmissions



$$X = f_c(c_i I + c_a A) + \delta[(1 - c_i)I + f_a(1 - c_a)A].$$

- ▶  $c_i$  and  $c_a$  are the fractions of confirmed cases for symptomatic and asymptomatic infectives.
- ▶  $f_c$ ,  $f_a$  are the infectivities of confirmed cases and asymptomatic cases, relative to an unconfirmed symptomatic infective.
- ▶  $\delta$  is a ‘contact factor’ that incorporates physical distancing and mask use for unconfirmed cases.

## The PSEAIHRD Model – Vaccination



- The coefficients  $\phi_i$  depend on the status of the overall ( $W$ ) and high-risk ( $W_2$ ) vaccination programs.

## Two-Class Vaccination Model

- ▶ Suppose a fraction  $h$  of the population is high-risk, with refusal probability  $r_2 \leq r$ .

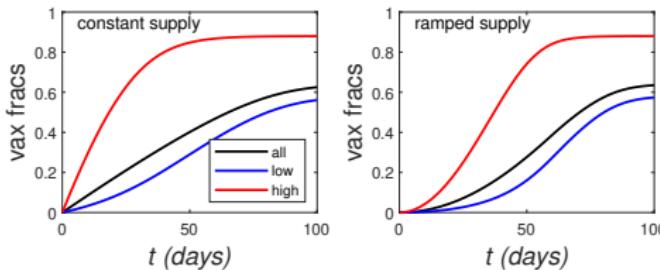
$$W(0) = 1 - r, \quad W_2(0) = (1 - r_2)h.$$

- ▶ Assume the same model for high-risk as for the whole group.

$$W' = -\frac{VK}{K + W} \equiv -VG(W)W, \quad W'_2 = -VG(W_2)W_2.$$

- Then

$$W'_1 = \dots = -VG(W)G(W_2)W_1, \quad W_1(0) = (1 - r_1)(1 - h).$$



## PSEAIHRD Vaccination Details

- We need  $\phi_j$  for

$$P'_j = -\phi_j P_j - \beta X P_j.$$

- We have

$$W'_1 = -VG(W)G(W_2)W_1, \quad W'_2 = -VG(W_2)W_2,$$

- Therefore

$$\phi_1 = VG(W)G(W_2), \quad \phi_2 = VG(W_2).$$

- We need only couple the  $W$  and  $W_2$  equations to the PSEAIHRD system.

## Resources

- ▶ See <https://www.math.unl.edu/SIR-modeling> for
  - Details on a classroom activity in disease modeling;
  - SIR and SEIR teaching modules using Excel and Matlab;
- ▶ See <https://www.math.unl.edu/covid-module> for COVID-19 teaching modules.
- ▶ Ledder and Homp, Using a COVID-19 model in various classroom settings to assess effects of interventions, PRIMUS 2021  
<https://www.tandfonline.com/doi/full/10.1080/10511970.2020.1861143>
- ▶ Ledder and Homp, Mathematical epidemiology, in Mathematics Research for the Beginning Student Volume 1: Accessible Research Projects for First- and Second-Year College and Community College Students before Calculus, ed. E.E. Goldwyn, A. Wootton, S. Ganzell. Birkhauser, in press

## Resources

- ▶ Ledder, Mathematical Modeling for Epidemiology and Ecology, 2ed, Springer, in press
  - 1. Modeling in Biology
  - 2. Empirical Modeling
  - 3. Mechanistic Modeling
    - 3.1 Transition processes (*includes vaccination*)
    - 3.2 Interaction processes
    - 3.3 Compartment analysis: The SEIR epidemic model
    - 3.4 SEIR model analysis
    - 3.5 Two scenarios from the COVID-19 pandemic
    - 3.9 Adding demographics to make an endemic disease model
  - 4. Dynamics of Single Populations
  - 5. Discrete Linear Systems
  - 6. Nonlinear Dynamical Systems
- ▶ Shoot me an email to receive updates or offer feedback!  
[gledder@unl.edu](mailto:gledder@unl.edu)