

Modeling infectious diseases in mixed household structures

Shrawan Parajuli

Mathematisches Institut
Universität Koblenz

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- Emergence of infectious diseases is concerning.
- spread of infectious disease gradually decreases due to immunity or become endemic due to lack of healthcare facilities.
- Study of infectious disease is an interdisciplinary fields is studied with different perspectives and purposes.
- Considering household structure has been an useful way of understanding spread of infectious diseases(for eg. Influenza) and it can be done with different epidemiological models such as SIS (Susceptible-Infectious-Susceptible).
- Also, the dynamics of infectious diseases is considered by studying equilibria.

Reference- [1],[3]

- To analyze equilibria (Disease-Free Equilibrium(DFE) and Endemic Equilibrium(EE)).
- To observe the influence of infection and recovery parameters in Population dynamics.

- Single Household (SHH) using SIS model.
- Master equation describes the time-evolution of P_k , P_k denotes the probability having k infected individuals and $(N-k)$ denotes susceptible in a given household size N ,
- External infection rate, $\epsilon = \tilde{\alpha} \frac{I}{T_p}$,
- Homogeneous household, Expectation: $\mathbb{E}I_H = \sum_{k=0}^N k * P_k$.
- $\frac{I}{T_p} \approx \frac{M_H * \mathbb{E}I_H}{M_H * N} = \frac{\mathbb{E}I_H}{N} = \frac{1}{N} \sum_{k=0}^N k P_k$
- In the SHH-N Model we have

$$\epsilon = \frac{\alpha}{N} \mathbb{E}I_H = \frac{\alpha}{N} \sum_{k=0}^N k P_k$$

Reference- [1]

■ State transition in (SHH)

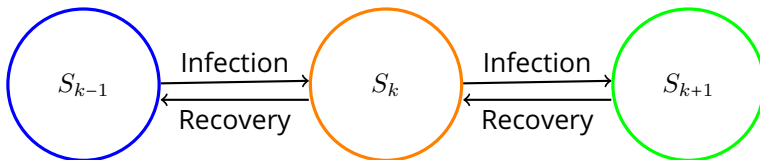


Figure 1: SHH diagram for state transition

$$\begin{aligned} \frac{dP_k}{dt} = & (X_{\{\geq 1\}}(k)) \left(\frac{\alpha}{N} \sum_{k=1}^N k P_k \right) (N - (k - 1)) P_{k-1} \\ & + (X_{\{\geq 2\}}(k)) \beta (N - (k - 1)) P_{k-1} \\ & + (X_{\{\leq N-1\}}(k)) \gamma (k + 1) P_{k+1} \\ & - (X_{\{\leq N-1\}}(k)) \left(\frac{\alpha}{N} \sum_{k=1}^N k P_k \right) (N - k) P_k \\ & - (X_{\{\geq 1\}}(k))(X_{\{\leq N-1\}}(k)) \beta (N - k) P_k \\ & - (X_{\{\geq 1\}}(k)) \gamma (k) P_k \end{aligned}$$

SHH-N

■ Positive Contribution

- (external infection) $S_{k-1} \rightarrow S_k$,
 $k \in \{1, 2, \dots, N\}$,
- (internal infection) $S_{k-1} \rightarrow S_k$,
 $k \in \{2, \dots, N\}$
- (recovery) $S_{k+1} \rightarrow S_k$, $k \in \{0, \dots, N-1\}$,

■ Negative Contribution

- (external infection) $S_k \rightarrow S_{k+1}$,
 $k \in \{0, \dots, N-1\}$,
- (internal infection) $S_k \rightarrow S_{k+1}$,
 $k \in \{1, \dots, N-1\}$
- (recovery) $S_k \rightarrow S_{k-1}$, $k \in \{1, \dots, N\}$,

Model and Method : SHH-N = 1,2

- General Master Equation for SHH in case of household size N=1,(SIS Model)

$$\frac{dP_0}{dt} = \gamma P_1 - \alpha P_1 P_0$$

$$\frac{dP_1}{dt} = \alpha P_1 P_0 - \gamma P_1$$

- General Master Equation for SHH in case of household size N=2,

$$\frac{dP_0}{dt} = \gamma P_1 - 2\left(\frac{\alpha}{2}P_1 + \alpha P_2\right)P_0$$

$$\frac{dP_1}{dt} = 2\left(\frac{\alpha}{2}P_1 + \alpha P_2\right)P_0 + 2\gamma P_2 - \left(\frac{\alpha}{2}P_1 + \alpha P_2\right)P_1 - \beta P_1 - \gamma P_1$$

$$\frac{dP_2}{dt} = \left(\frac{\alpha}{2}P_1 + \alpha P_2\right)P_1 + \beta P_1 - 2\gamma P_2$$

Reference- [1]

Model and Method : Analytical Approach

- Non-linear differential equations is studied with stability analysis, phase portraits and Numerical simulation.

- Total infection rate :

$$\frac{d\mathbb{E}[I_H]}{dt} = \frac{dP_1}{dt} + 2\frac{dP_2}{dt}$$

$$I_{Total} = \int_0^T \mathbb{E}[I_H(t)] dt$$

- Equilibrium

- DFE, $\lim_{t \rightarrow \infty} (P_0(t), P_1(t), P_2(t)) = (1, 0, 0)$.
- In EE, $\lim_{t \rightarrow \infty} (P_0(t), P_1(t), P_2(t)) \neq (1, 0, 0)$ (or $(1, 0, 0)$ does not exist).

- In equilibrium :

$$dP_k/dt = f_k(P_k) = 0$$

Model and Method : Linearization, DFE

- Linearization in SHH-2
- DFE, Jacobian Matrix (J) at $(1, 0, 0)$

$$\begin{pmatrix} 0 & -\alpha + \gamma & -2\alpha \\ 0 & \alpha - \beta - \gamma & 2\alpha + 2\gamma \\ 0 & \beta & -2\gamma \end{pmatrix}$$

- characteristic equation of the form is $|J - \lambda E| = 0$,

$$\begin{pmatrix} \alpha - \beta - \gamma & 2\alpha + 2\gamma \\ \beta & -2\gamma \end{pmatrix}$$

$$\lambda_i = \frac{Tr - \sqrt{Tr^2 - 4D}}{2}$$

$$\lambda_{ii} = \frac{Tr + \sqrt{Tr^2 - 4D}}{2}$$

Reference- [2]

- Let A be 2 X 2 square matrix of form written in

$$\begin{pmatrix} a_{11} & a_{12} \\ a_{21} & a_{22} \end{pmatrix}$$

- $Trace(Tr) = a_{11} + a_{22} = \alpha - \beta - 3\gamma$
- $Determinant(D) = a_{11}a_{22} - a_{12}a_{21} = 2(\gamma^2 - \alpha(\beta + \gamma))$

- EE, endemic equilibrium point (P_0, P_1, P_2) or $(P_0, P_1, 1 - P_0 - P_1)$

- P_1 ,

$$P_1 = \frac{2\alpha P_0(1 - P_0)}{(\gamma + \alpha P_0)} \quad (1)$$

- positive root P_0 ,

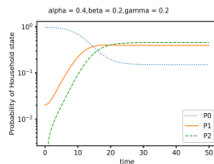
$$P_0 = \frac{\gamma}{2\alpha\beta} (-(\alpha + \beta) + \sqrt{(\alpha + \beta)^2 + 4\beta\gamma}) \quad (2)$$

- jacobian at $(P_0, P_1, 1 - P_0 - P_1)$

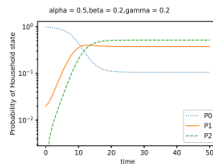
$$\begin{pmatrix} -\alpha(2 - 2P_0 - P_1) & -\alpha P_0 + \gamma & -2\alpha P_0 \\ \alpha(2 - 2P_0 - P_1) & P_0(1 + \alpha) - 1 - \beta - \gamma & 2\alpha P_0 - \alpha P_1 + 2\gamma \\ 0 & \alpha(1 - P_0) + \beta & \alpha P_1 - 2\gamma \end{pmatrix} \quad (3)$$

- Numerical method: Runge-Kutta Method, ($RK45$)
- Initial conditions, P_0 is $P_0(0) = 0.98$, P_1 is $P_1(0) = 0.02$ and P_2 is $P_2(0) = 0$,
- timestep : $[0,50]$
- parameters
 - α : between-household infection rate.
 - β : within-household infection rate.
 - γ : household recovery rate
- vary only one parameter and keep the other two parameter fixed
- regular spaced (0.2,0.3,0.4,0.5,0.6,0.7,0.8) rates
- initial rates , $\alpha = 0.4$, $\beta = 0.2$, $\gamma = 0.2$
- for example : Influenza recovered rate 0.2 (takes 5 days to recover)

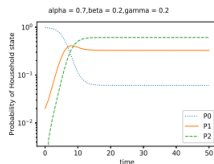
Probability of household state vs time (α)



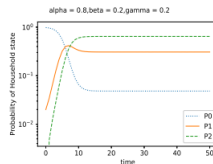
(a) $\alpha = 0.4, \beta = 0.2, \gamma = 0.2$



(b) $\alpha = 0.5, \beta = 0.2, \gamma = 0.2$



(c) $\alpha = 0.7, \beta = 0.2, \gamma = 0.2$



(d) $\alpha = 0.8, \beta = 0.2, \gamma = 0.2$

- Probability of household states(P_k)

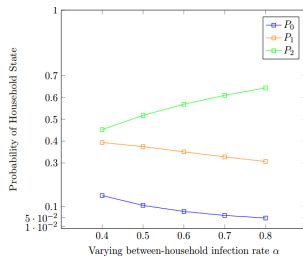
$$|P_{k_{t_{i+1}}} - P_{k_{t_i}}| < 10^{-3}$$

- Total infection rate(I_{Total})

$$|I_{Total_{t_{i+1}}} - I_{Total_{t_i}}| < 10^{-3}$$

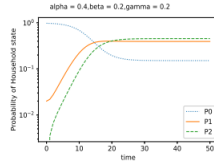
Probabilities of household states	$\alpha = 0.4$	$\alpha = 0.5$	$\alpha = 0.6$	$\alpha = 0.7$	$\alpha = 0.8$
P_0	0.151	0.106	0.078	0.060	0.048
P_1	0.394	0.375	0.351	0.328	0.307
P_2	0.453	0.518	0.569	0.610	0.644

Name	$\alpha = 0.4$	$\alpha = 0.5$	$\alpha = 0.6$	$\alpha = 0.7$	$\alpha = 0.8$
Total infection rate (I_{Total})	(1.290)	(1.402)	(1.482)	(1.543)	(1.592)
Total recovered rate (R_{Total})	(0.258)	(0.280)	(0.296)	(0.308)	(0.318)
Total Timestep	(35)	(28)	(26)	(23)	(19)

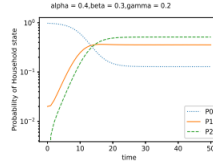


P_0, P_1, P_2 on y-axis and α on x-axis

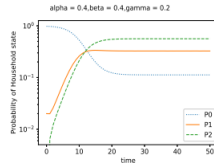
Probability of household state vs time (β)



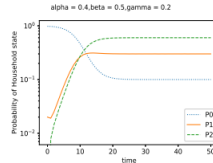
(a) $\alpha = 0.4, \beta = 0.2, \gamma = 0.2$



(b) $\alpha = 0.4, \beta = 0.3, \gamma = 0.2$



(c) $\alpha = 0.4, \beta = 0.4, \gamma = 0.2$



(d) $\alpha = 0.4, \beta = 0.5, \gamma = 0.2$

- Probability of household states(P_k)

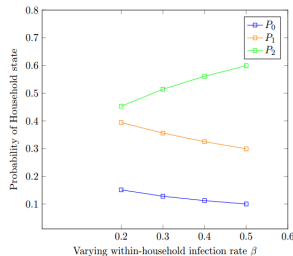
$$|P_{k_{t_{i+1}}} - P_{k_{t_i}}| < 10^{-3}$$

- Total infection rate(I_{Total})

$$|I_{Total_{t_{i+1}}} - I_{Total_{t_i}}| < 10^{-3}$$

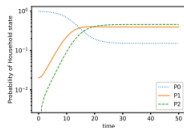
Probability of HouseHold state	$\beta = 0.2$	$\beta = 0.3$	$\beta = 0.4$	$\beta = 0.5$
P_0	0.151	0.128	0.112	0.100
P_1	0.394	0.356	0.325	0.299
P_2	0.453	0.514	0.561	0.599

Name	$\beta = 0.2$	$\beta = 0.3$	$\beta = 0.4$	$\beta = 0.5$
Total infection rate (I_{Total})	(1.290)	(1.375)	(1.441)	(1.493)
Total recovered rate (R_{Total})	(0.258)	(0.275)	(0.288)	(0.298)
Timestep	(35)	(31)	(30)	(27)

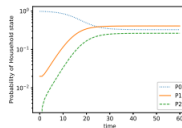


P_0, P_1, P_2 on y-axis and β on x-axis

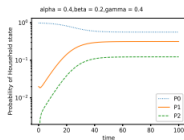
Probability of household state vs time (γ)



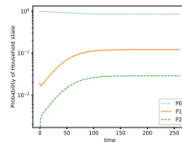
(a) $\alpha = 0.4, \beta = 0.2, \gamma = 0.2$



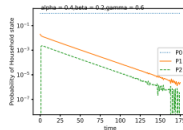
(b) $\alpha = 0.4, \beta = 0.2, \gamma = 0.3$



(c) $\alpha = 0.4, \beta = 0.2, \gamma = 0.4$



(d) $\alpha = 0.4, \beta = 0.2, \gamma = 0.5$



(e) $\alpha = 0.4, \beta = 0.2, \gamma = 0.6$

Probability of household state vs time (γ):cont.

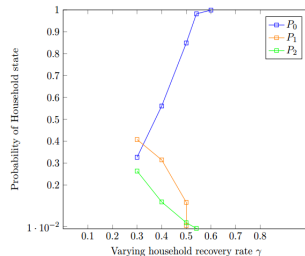
- Probability of household states(P_k)

$$|P_{k_{t_{i+1}}} - P_{k_{t_i}}| < 10^{-3}$$

- Total infection rate(I_{Total})

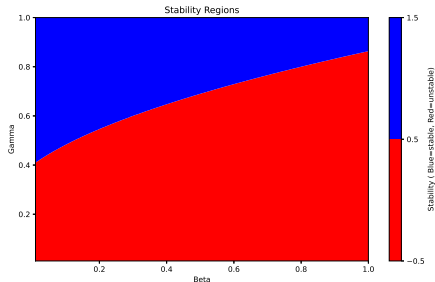
$$|I_{Total_{t_{i+1}}} - I_{Total_{t_i}}| < 10^{-3}$$

Probability of Household state	$\gamma = 0.3$	$\gamma = 0.4$	$\gamma = 0.5$	$\gamma = 0.6$
P_0	0.327	0.561	0.849	0.9999
P_1	0.408	0.315	0.121	-
P_2	0.264	0.123	0.028	-
Name	$\gamma = 0.3$	$\gamma = 0.4$	$\gamma = 0.5$	$\gamma = 0.6$
Total infection rate (I_{Total})	(0.922)	(0.545)	(0.161)	-
Total recovered rate (R_{Total})	(0.276)	(0.218)	(0.080)	-
Timestep	(54)	(74)	(185)	-



P_0, P_1, P_2 on y-axis and γ on x-axis

Disease-Free equilibrium



γ range on y -axis and β range on x -axis
with constant α at 0.4.

- **eigenvalues** : $\lambda_i < 0, \lambda_{ii} < 0$
- **case** : $Tr < 0$ and $Tr > \sqrt{Tr^2 - 4D}$,
where $D > 0$ and $Tr^2 > 4D$
- **Parameter** : $Tr < 0$, means $\alpha < \beta + 3\gamma$
and $D > 0$ means $\alpha < \frac{\gamma^2}{\beta + \gamma}$

Endemic equilibrium: varying α

Probabilities of household states	$\alpha = 0.4$	$\alpha = 0.5$	$\alpha = 0.6$	$\alpha = 0.7$	$\alpha = 0.8$
P_0	0.151	0.106	0.078	0.060	0.048
P_1	0.394	0.375	0.351	0.328	0.307
P_2	0.453	0.518	0.569	0.610	0.644

Runge-Kutta Method: α

HouseHold states	$\alpha = 0.4$	$\alpha = 0.5$	$\alpha = 0.6$	$\alpha = 0.7$	$\alpha = 0.8$
P_0	0.151	0.106	0.078	0.060	0.048
P_1	0.394	0.375	0.351	0.328	0.307
P_2	0.454	0.518	0.569	0.610	0.644

Equilibrium: α

- **Numerical values:** RK method and endemic equilibrium values are similar.
- $P_0 = \frac{\gamma}{2\alpha\beta} (-(\alpha + \beta) + \sqrt{(\alpha + \beta)^2 + 4\beta\gamma})$
- $P_1 = \frac{2\alpha P_0(1-P_0)}{(\gamma + \alpha P_0)}$
- $P_2 = 1 - P_0 - P_1$
- value of P_0 affects value in P_1 and summation of both P_0 and P_1 results change in P_2 .
- **linearization :** Two negative eigenvalues(stable).

Endemic equilibrium: varying β

Probability of HouseHold state	$\beta = 0.2$	$\beta = 0.3$	$\beta = 0.4$	$\beta = 0.5$
P_0	0.151	0.128	0.112	0.100
P_1	0.394	0.356	0.325	0.299
P_2	0.453	0.514	0.561	0.599

Runge-Kutta Method : β

HouseHold states	$\beta = 0.2$	$\beta = 0.3$	$\beta = 0.4$	$\beta = 0.5$
P_0	0.151	0.128	0.112	0.100
P_1	0.394	0.356	0.325	0.300
P_2	0.454	0.514	0.561	0.600

Equilibrium: β

- **Numerical values:** RK method and endemic equilibrium values are similar.
- $P_0 = \frac{\gamma}{2\alpha\beta} (-(\alpha + \beta) + \sqrt{(\alpha + \beta)^2 + 4\beta\gamma})$
- $P_1 = \frac{2\alpha P_0(1-P_0)}{(\gamma + \alpha P_0)}$
- $P_2 = 1 - P_0 - P_1$
- value of P_0 affects value in P_1 and summation of both P_0 and P_1 results change in P_2 .
- **linearization :** Two negative eigenvalues(stable).

Endemic equilibrium: varying γ

Probability of Household state	$\gamma = 0.3$	$\gamma = 0.4$	$\gamma = 0.5$	$\gamma = 0.6$
P_0	0.327	0.561	0.849	0.9999
P_1	0.408	0.315	0.121	-
P_2	0.264	0.123	0.028	-

Runge-Kutta Method : γ

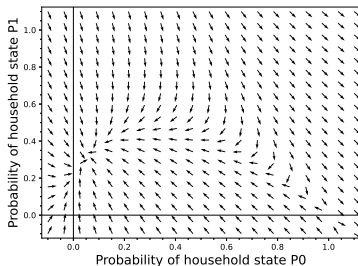
HouseHold states	$\gamma = 0.3$	$\gamma = 0.4$	$\gamma = 0.5$	$\gamma = 0.6$
P_0	0.327	0.561	0.849	-
P_1	0.408	0.315	0.121	-
P_2	0.263	0.123	0.028	-

Equilibrium: γ

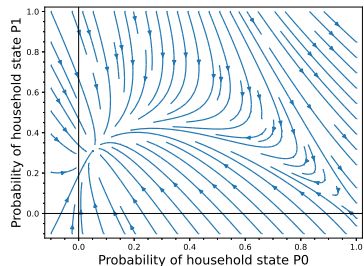
- **Numerical values:** RK method and endemic equilibrium values are similar.
- $P_0 = \frac{\gamma}{2\alpha\beta} (-(\alpha + \beta) + \sqrt{(\alpha + \beta)^2 + 4\beta\gamma})$
- $P_1 = \frac{2\alpha P_0(1-P_0)}{(\gamma + \alpha P_0)}$
- $P_2 = 1 - P_0 - P_1$
- value of P_0 affects value in P_1 and summation of both P_0 and P_1 results change in P_2 .
- **linearization :** Two negative eigenvalues(stable).

$$\begin{aligned}f_0(P_0) &= -\alpha(P_1 + 2(1 - P_0 - P_1))P_0 + \gamma P_1 \\f_1(P_1) &= \alpha(P_1 + 2(1 - P_0 - P_1))P_0 \\&\quad - ((\alpha/2)(P_1 + 2(1 - P_0 - P_1)) + \beta + \gamma)P_1 \\&\quad + 2\gamma(1 - P_0 - P_1)\end{aligned}\tag{4}$$

vector plot and stream plot: α

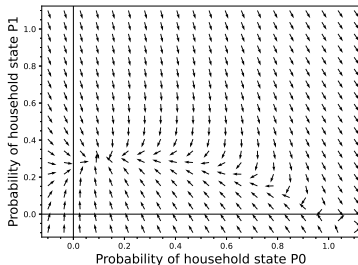


$$\alpha = 0.8, \beta = 0.2, \gamma = 0.2$$

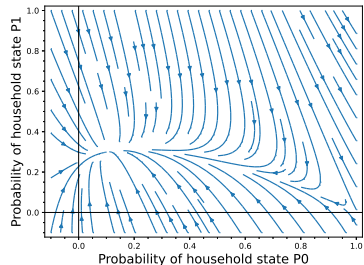


$$\alpha = 0.8, \beta = 0.2, \gamma = 0.2$$

vector plot and stream plot: β

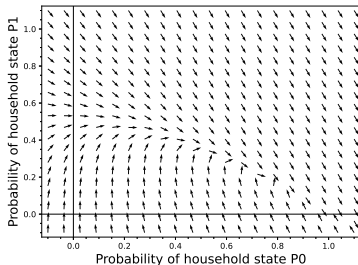


$$\alpha = 0.4, \beta = 0.5, \gamma = 0.2$$

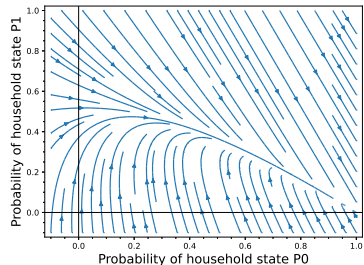


$$\alpha = 0.4, \beta = 0.5, \gamma = 0.2$$

vector plot and stream plot: γ



$$\alpha = 0.4, \beta = 0.2, \gamma = 0.6$$



$$\alpha = 0.4, \beta = 0.2, \gamma = 0.6$$

- Total Timestep:
 - Gradual **increase** in α and β , **decreases** the **timestep** to reach steady values.
 - In **comparison**, there is **decrease** in the timestep for α but an **increase** in the total timestep β .
 - An **increase** in γ **increases** timestep.
- P_0, P_1, P_2 and I_{Total} :
 - An **increase** in α and β leads to a **decrease** in P_0 and P_1 but **increase** in P_2 and I_{Total} .
 - when **comparing increase** in α and β : P_0 **decreases more** in increase in α and **decreases less** in increase in β while P_1 **decreases less** in increase in α and **decrease more** in β whereas P_2 remains similar for both in α and β .
 - while **comparing** for I_{Total} : increase in α **increases more** in I_{Total} but **increases less** in gradual increase in β .
 - increase in γ **increases** P_0 while (P_1, P_2, I_{Total}) **decreases**.

- Stability in gradual increase in α, β and γ (till 0.5), EE.
- Stability seen at $\gamma = 0.6$, at $(P_0, P_1, P_2) = (P_0, 0, 0)$, DFE.
- At Phase portraits,
 - where EE stable (sink), DFE is seen unstable (saddle).
 - where DFE is seen stable (sink), EE is seen unstable (saddle).

- In summary, a transition from EE to DFE is observed, at $\gamma = 0.6$, $\alpha = 0.4$ and $\beta = 0.2$.
- Increase in α and β leads to EE and increase in γ leads to DFE.

- Limitation :
 - Mixed household size rather than same household size.
 - Larger household size rather than only household of size two.
 - Strategies or policies.
 - Household structure (space) and behavioural patterns(movement).
- Further research :
 - Epidemiological models.
 - Higher dimension.
 - Larger population.
 - lower rates in parameters.
 - Intervention strategies.

- [1] Alex Holmes, Mike Tildesley, and Louise Dyson. Approximating steady state distributions for household structured epidemic models. *Journal of Theoretical Biology*, 534:110974, 2022
- [2] Carl P. Simon and John A. Jacquez. Reproduction numbers and the stability of equilibria of si models for heterogeneous populations. *SIAM Journal on Applied Mathematics*, 52(2):541576, 1992
- [3] Fred Brauer. Mathematical epidemiology: Past, present, and future. *Infectious Disease Modelling*, 2(2):113127, 2017

Thank You