

Week 1 - Intro to dynamical Systems & stability

A system of ODEs

$$\dot{x}_1 = f_1(x_1, \dots, x_n, t)$$

⋮

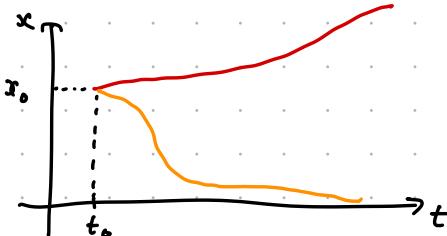
$$\dot{x}_n = f_n(x_1, \dots, x_n, t)$$

$$\frac{dx_i}{dt}$$

Autonomous systems
so our f_1, \dots, f_n
doesn't depend on
 t and $n \leq 3$
Simplification!

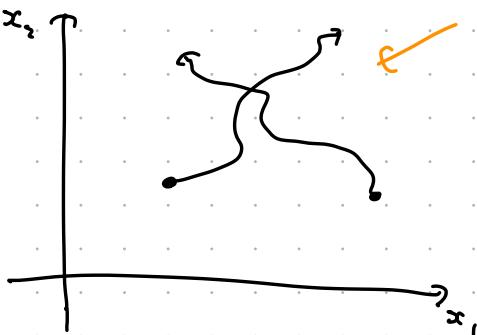
Assume existence and uniqueness of solutions. This means:

• 1D:

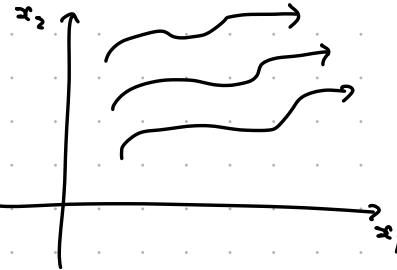


either increasing or
decreasing but
not both
or contradictions.

• 2D:



solutions can't cross in 2d.
(contradicts uniqueness) we get
that trajectories never cross

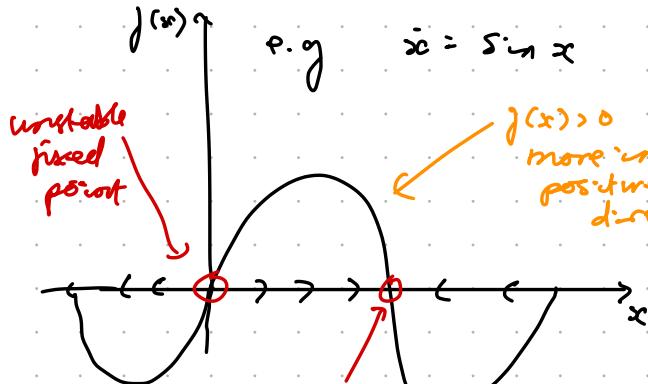


Def: $x_0 \in \mathbb{R}^n$ is an initial condition
& the path given by $x(t) \in \mathbb{R}^n$
is called the trajectory corresponding
to x_0 .

First order Systems

$$\dot{x} = f(x)$$

independent variable
doesn't appear
autonomous.



This is a
fixed point
stable

$f(x) > 0$ so
more in
positive
direction etc

"Phase line" / "vector field"

Phase diagrams.

Def: A fixed point x^* is
stable if all sufficiently small
deviations from x^* damp
out over time.

- ① If all solutions that start near x^* stay near x^* forever, we say that x^* is Lyapunov stable.
Simplest type ↗
- ② If all solutions that start near x^* converge to x^* (as $t \rightarrow \infty$)
then x^* is asymptotically stable.
no mention of how long
- ③ If solutions converge to x^*
at some nonuniform rate, then

Def: x^* is unstable if disturbances grow in time.

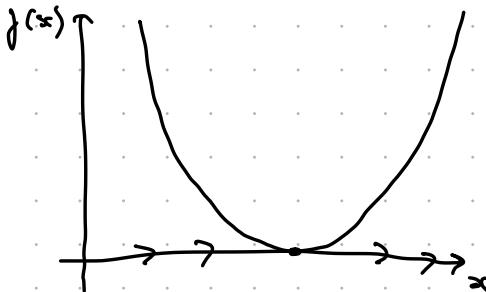
x^* is exponentially stable.

other options!

Saddle node occurs when

$$\dot{x} = f(x)$$

more complex to classify.



stable from one side & unstable from the other.

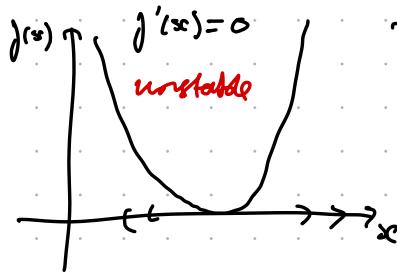
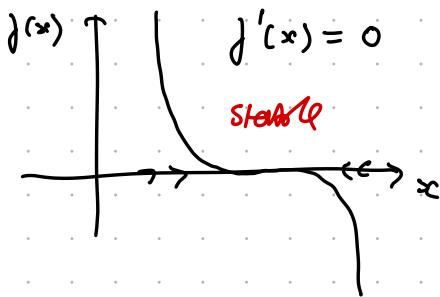
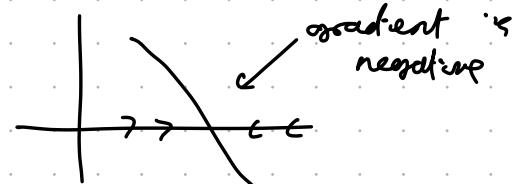
Linear stability Analysis

Look at a system, stable or not???

Thm: Take $n=1$

① x^* is stable if $\frac{df}{dx}(x^*) < 0$

② x^* is unstable if $\frac{df}{dx}(x^*) > 0$



Just draw a picture if you have $f'(x)=0$

Example: (Population growth)

- N is population size, ($N > 0$)
- $r > 0$, $k > 0$

$$\dot{N} = r N \left(1 - \frac{N}{k}\right) = f(N)$$

$$f(N) = 0$$

when?

when $N=0$ or $N=k$

k is carrying capacity

- If $N > k$, then decrease
- If $N < k$, then increase

[This is the logistic equation]

$$\text{Now, want } f'(N) = r - 2r \frac{N}{k}$$

$$f'(0) = r > 0 \Rightarrow \text{unstable}$$

[no one in population so unstable & will grow]

$$f'(k) = -r < 0 \Rightarrow \text{stable}$$

[start near carrying capacity, then will stay]

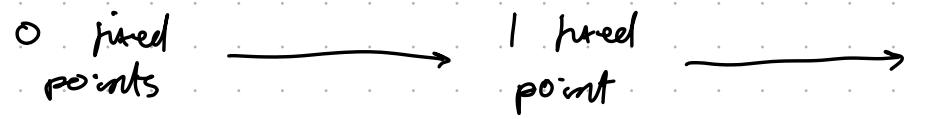
Bifurcations

changes value of a parameter but something qualitatively changes.

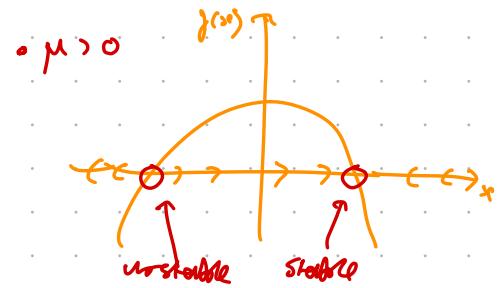
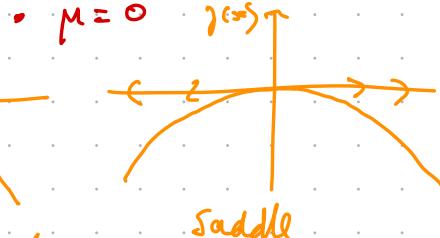
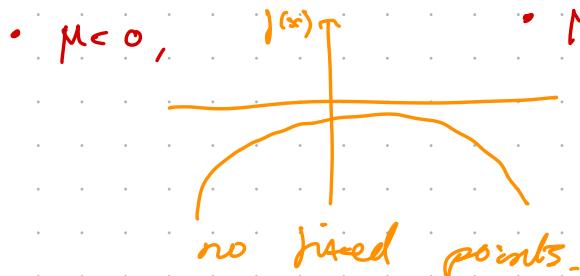
- changes in # of fixed points.
- changes / switch in stability of fixed points.

These are different types.

① Saddle node bifurcation:



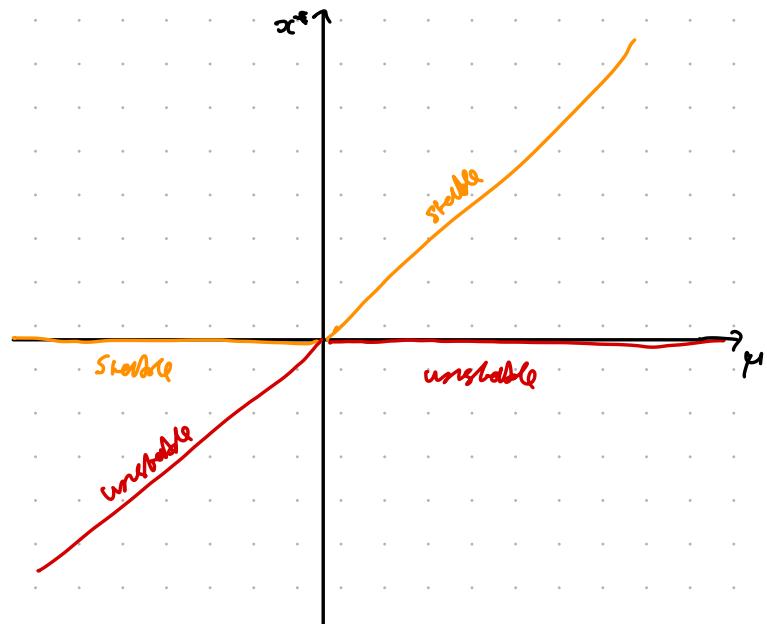
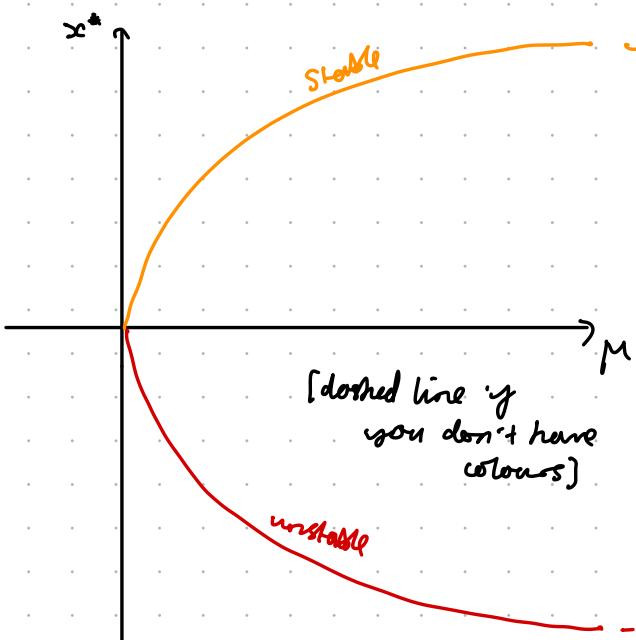
Example: $\dot{x} = \mu - x^2$. Consider different μ



Above is messy, instead, draw a bifurcation diagram, can vary along horizontal axis, then value of fixed points on y

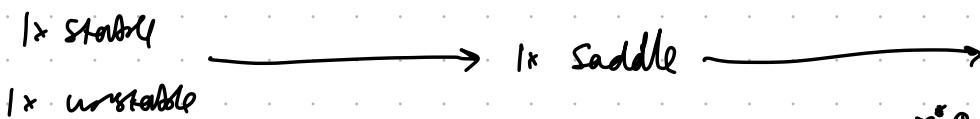
$$\dot{x} = \mu - x^2 = 0$$

$$\sqrt{\mu} = \geq 0$$



Saddle node bifurcation.

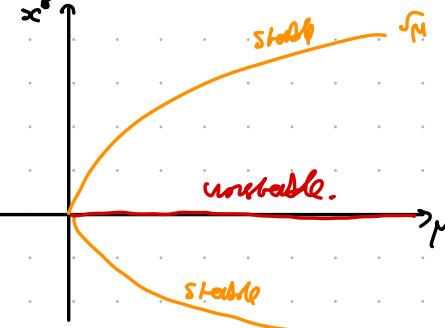
② Transcritical bifurcation



Example: $\dot{x} = \mu x - x^2 = 0$

$x^* = 0, M \Rightarrow$ see picture above.

1x unstable
1x stable



③ Pitchfork bifurcation

Example: $\dot{x} = \mu x - x^3 = x(\mu - x^2)$

$x^* = 0, x^* = \pm \sqrt{\mu}$ so depending on μ , get different # fixed points.
if $\mu > 0$, it $\mu = 0$ one... stable for $\mu > 0$

Second Order Systems

$$\begin{cases} \dot{x} = f(x, y) \\ \dot{y} = g(x, y) \end{cases}$$

Linear Systems

$$\begin{aligned} \dot{x} &= ax + by \\ \dot{y} &= cx + dy \end{aligned}$$

$a, b, c, d \in \mathbb{R}$

zero is always a fixed point in a linear system.

$$\dot{\underline{x}} = A \underline{x}$$

$\underline{x} \in \mathbb{R}^2$ here

$$\underline{x} = (x, y) \quad A = \begin{pmatrix} a & b \\ c & d \end{pmatrix}$$

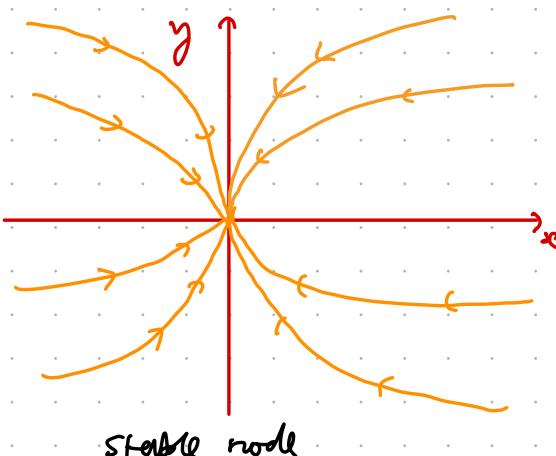
as simple as a system can be

Example: $A = \begin{pmatrix} a & 0 \\ 0 & -1 \end{pmatrix} \Rightarrow \dot{\underline{x}} = A \underline{x} \Rightarrow \begin{cases} \dot{x} = ax \\ \dot{y} = -y \end{cases}$

Now cases:

① $a < 0$

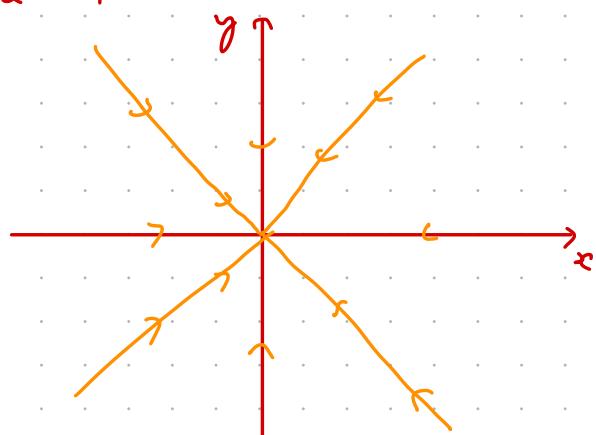
(i) $a < -1$ $x(t) \rightarrow 0$
quicker than $y(t) \rightarrow 0$



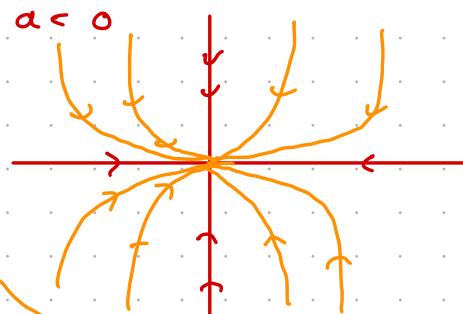
stable node

(ii) $a = -1$

stable star

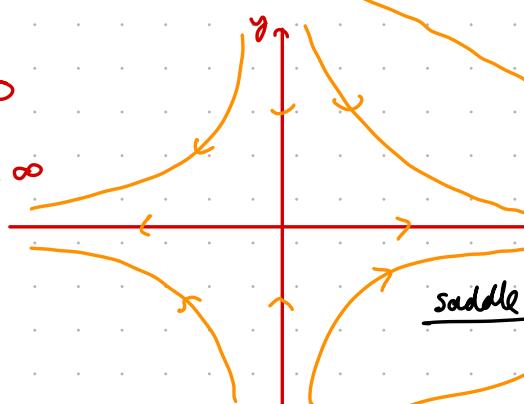


(iii) $-1 < a < 0$



② $a > 0$

$y(t) \rightarrow 0$
 $x(t) \rightarrow \pm\infty$

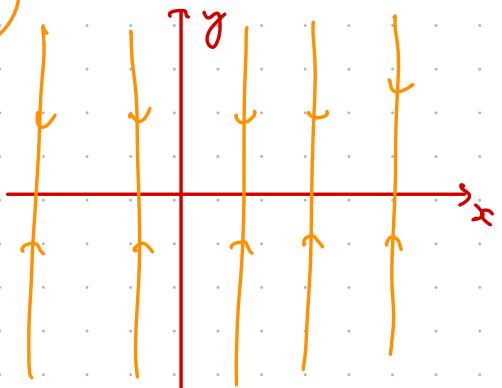


stable manifold is
y axis
unstable manifold
is x axis

[trajectory back in
time to where
stable]

WHY NICE?
Eigenvectors are
just the axes so
nice

line of fixed
points.



General System

$$\dot{\underline{x}} = A \underline{x}, \text{ try } \underline{x} = e^{\lambda t} \underline{v}$$

can write down a general solution to this system.

$$\dot{\underline{x}} = \lambda e^{\lambda t} \underline{v} = A e^{\lambda t} \underline{v}$$

↑ eigenvalue of A .

$$\boxed{\underline{x}(t) = c_1 e^{\lambda_1 t} \underline{v}_1 + c_2 e^{\lambda_2 t} \underline{v}_2}$$

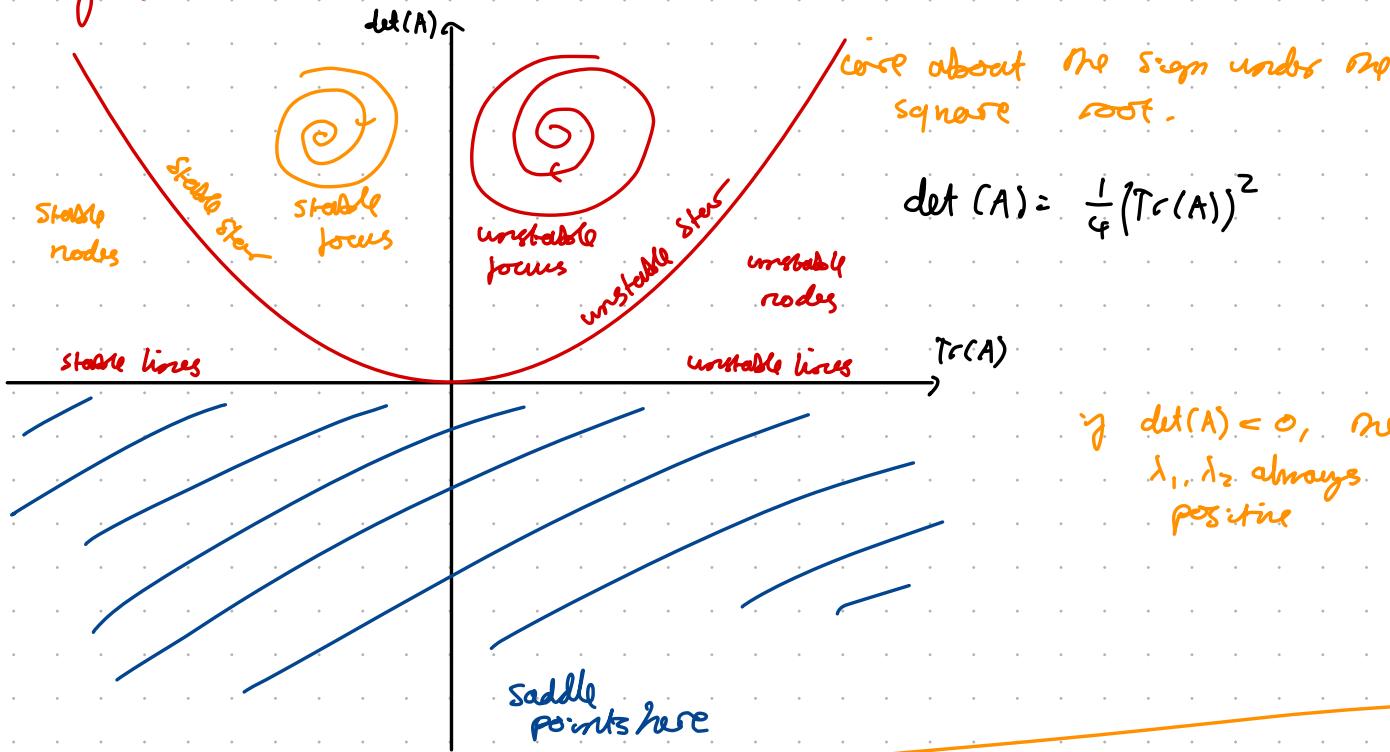
$\underline{v}_1, \underline{v}_2$ are eigenvectors of A

λ_1, λ_2 are eigenvalues.

$$\lambda_1, \lambda_2 \text{ are roots of } c_A(x) = 0 \Rightarrow \lambda^2 - \underbrace{(\alpha + \beta)}_{\text{trace } A} \lambda + \underbrace{(\alpha\beta - \gamma\delta)}_{\det(A)} = 0$$

$$\Rightarrow \lambda_{1,2} = \frac{1}{2} \left(\text{Tr}(A) \pm \sqrt{(\text{Tr}(A))^2 - 4 \det(A)} \right)$$

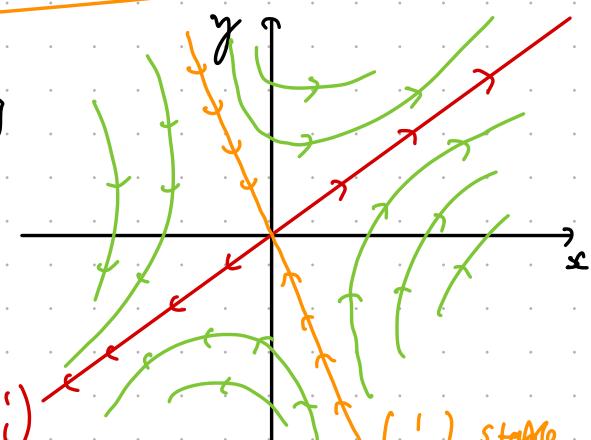
why bother with this? Easier to use tr & det \therefore can see relationships easier.



$$\det(A) = \frac{1}{4} (\text{Tr}(A))^2$$

If $\det(A) < 0$, then λ_1, λ_2 always positive

Example: $A = \begin{pmatrix} 1 & 1 \\ 4 & -2 \end{pmatrix}$ $\lambda_{1,2} = 2, -3$ [saddle]
 $\underline{v}_{1,2} = \begin{pmatrix} 1 \\ 1 \end{pmatrix} \text{ & } \begin{pmatrix} 1 \\ -4 \end{pmatrix}$



$\begin{pmatrix} 1 \\ 1 \end{pmatrix}$
[unstable manifold]

$\begin{pmatrix} 1 \\ -4 \end{pmatrix}$ stable manifold

Non-Linear Systems

$$\begin{aligned}\dot{x}_1 &= f_1(x_1, x_2) \\ \dot{x}_2 &= f_2(x_1, x_2)\end{aligned}$$

or $\dot{\underline{x}} = \underline{f}(\underline{x})$

$$x(0) = x_0 \text{ so}$$

$$\phi(x_0, t)$$

[This is where we start & vary x_0 to get a family, again a flow]

Def: A flow $\phi(x, t)$ is the solution at time t with initial value x at $t=0$.

Trajectories can't intersect or we lose uniqueness.

Def: A fixed point \underline{x}^* is

- ① An attracting fixed point if all trajectories that start near \underline{x}^* approach it as $t \rightarrow \infty$
 → It's orbitally attracting if \underline{x}^* attracts all trajectories.
 [no matter where you start, will always end here].

- ② Lyapunov stable if $\forall \varepsilon > 0 \exists \delta > 0$ s.t. $|x(0) - \underline{x}^*| < \delta$
 Then $|x(t) - \underline{x}^*| < \varepsilon \quad \forall t > 0$

Start close - Stay close stability.

- ③ Asymptotically stable if you're Lyapunov stable and $\exists \delta > 0$ s.t. if

$$|\underline{x}(0) - \underline{x}^*| < \delta \Rightarrow |\underline{x}(t) - \underline{x}^*| \xrightarrow[t \rightarrow \infty]{} 0$$

Start within δ of fixed point. will spread into fixed point.

won't be asked for ε - δ based proof! Context for methods & theory, not math.

Linearisation

$$\frac{dx}{dt} = ax + by = f(x, y) \quad \underline{x} = \begin{pmatrix} a & b \\ c & d \end{pmatrix} \underline{x}$$

$$\frac{dy}{dt} = cx + dy = g(x, y) \quad \text{"the jacobian"}$$

$$\begin{pmatrix} a = \frac{\partial f}{\partial x} & b = \frac{\partial f}{\partial y} \\ c = \frac{\partial g}{\partial x} & d = \frac{\partial g}{\partial y} \end{pmatrix} \leftarrow \begin{matrix} \underline{x} \\ \underline{y} \end{matrix}$$

now take this:

$$\frac{dx}{dt} = f(x, y) \quad \text{fixed point}$$

$$\frac{dy}{dt} = g(x, y) \quad (\underline{x}^*, \underline{y}^*)$$

disturbance

$$\begin{aligned}u &= x - \underline{x}^* \\ v &= y - \underline{y}^*\end{aligned}$$

behaviour around fixed points is the same as linear systems

$$\begin{pmatrix} u \\ v \end{pmatrix} = A \begin{pmatrix} u \\ v \end{pmatrix}$$

Proof: Taylor expansion, in notes...

Thm: If $\dot{x} = f(x)$ has equilibrium x^* and a linearisation $\dot{x} = Ax$, then if A has no zero eigenvalues then the local stability of x^* is entirely determined by the linear system.

Population Models

(a) Predator Prey - 'Lotka Volterra'

Example: x prey y predators

$$(prey) \frac{dx}{dt} = \alpha x - \beta xy \quad (predators) \frac{dy}{dt} = -\gamma y + \delta xy$$

$r_1 \boxed{\frac{x}{K_1}} x, y$

growth of prey unbounded. how much predators eat proportion of max population $\Rightarrow s = 2e$ no food, no prey infinite prey means predators keep growing

(b) Competing species

$$\frac{dx}{dt} = r_1 x \left(1 - \frac{x}{K_1} - \frac{\alpha_1 y}{K_1}\right) \quad \frac{dy}{dt} = r_2 y \left(1 - \frac{y}{K_2} - \frac{\alpha_2 x}{K_2}\right)$$

limits populations $\Rightarrow s = 2e$ competing populations same but reflected.

E.g. rabbits & sheep: (r, s)

$$\begin{aligned} \text{not realistic, } r &= 3r \left(1 - \frac{r}{3} - \frac{2s}{3}\right) \\ \text{real numbers! } s &= 2s \left(1 - \frac{s}{2} - \frac{r}{2}\right) \end{aligned} \Rightarrow \begin{aligned} \dot{r} &= r(3-r-2s) = j \\ \dot{s} &= s(2-r-s) = g \end{aligned}$$

Fixed points: $\dot{r} = \dot{s} = 0 \Rightarrow r = 0 \text{ or } s = 0$

- ① check $r = 0$: $\dot{s} = s(2-s) = 0 \Rightarrow s = 0 \text{ or } s = 2$
- ② Fixed points at $(0,0)$ and $(0,2)$
- ③ check $s = 0$: $\dot{r} = r(3-r) = 0 \Rightarrow r = 0 \text{ or } r = 3$
- ④ Fixed points at $(0,0)$ and $(3,0)$
- ⑤ $3-r-2s = 0$
 $2-r-s = 0$ } \rightarrow fixed point at $(1,1)$

so fixed points are $(0,0), (0,2), (3,0), (1,1)$

want to linearise system to get Df

$$A = \begin{pmatrix} \frac{\partial f}{\partial r} & \frac{\partial f}{\partial s} \\ \frac{\partial g}{\partial r} & \frac{\partial g}{\partial s} \end{pmatrix} = \begin{pmatrix} 3-2r-2s & -2r \\ -s & 2-r-2s \end{pmatrix}$$

evaluate A at fixed points. Prey eigenvalues etc.

Fixed point $(0,0)$: $A = \begin{pmatrix} 3 & 0 \\ 0 & 2 \end{pmatrix} \quad \lambda_{1,2} = 3, 2 \quad v_{1,2} = \begin{pmatrix} 1 \\ 0 \end{pmatrix}, \begin{pmatrix} 0 \\ 1 \end{pmatrix}$

$\lambda_{1,2} > 0 \Rightarrow$ unstable node.

Fixed Point $(0,2)$
just sheep, no rabbits $A = \begin{pmatrix} -1 & 0 \\ -2 & -2 \end{pmatrix} \quad \lambda_{1,2} = -2, -1 \quad v_{1,2} = \begin{pmatrix} 0 \\ 1 \end{pmatrix}, \begin{pmatrix} -1 \\ 2 \end{pmatrix} \neq$

\Rightarrow stable node :: both λ negative

Fixed point $(3, 0)$
just rabbits no sheep
(would stay here)

$$A = \begin{pmatrix} -3 & -6 \\ 0 & -1 \end{pmatrix} \quad \lambda_{1,2} = -3, -1 \quad v_{1,2} = \begin{pmatrix} 1 \\ 0 \end{pmatrix}, \begin{pmatrix} -3 \\ 1 \end{pmatrix}$$

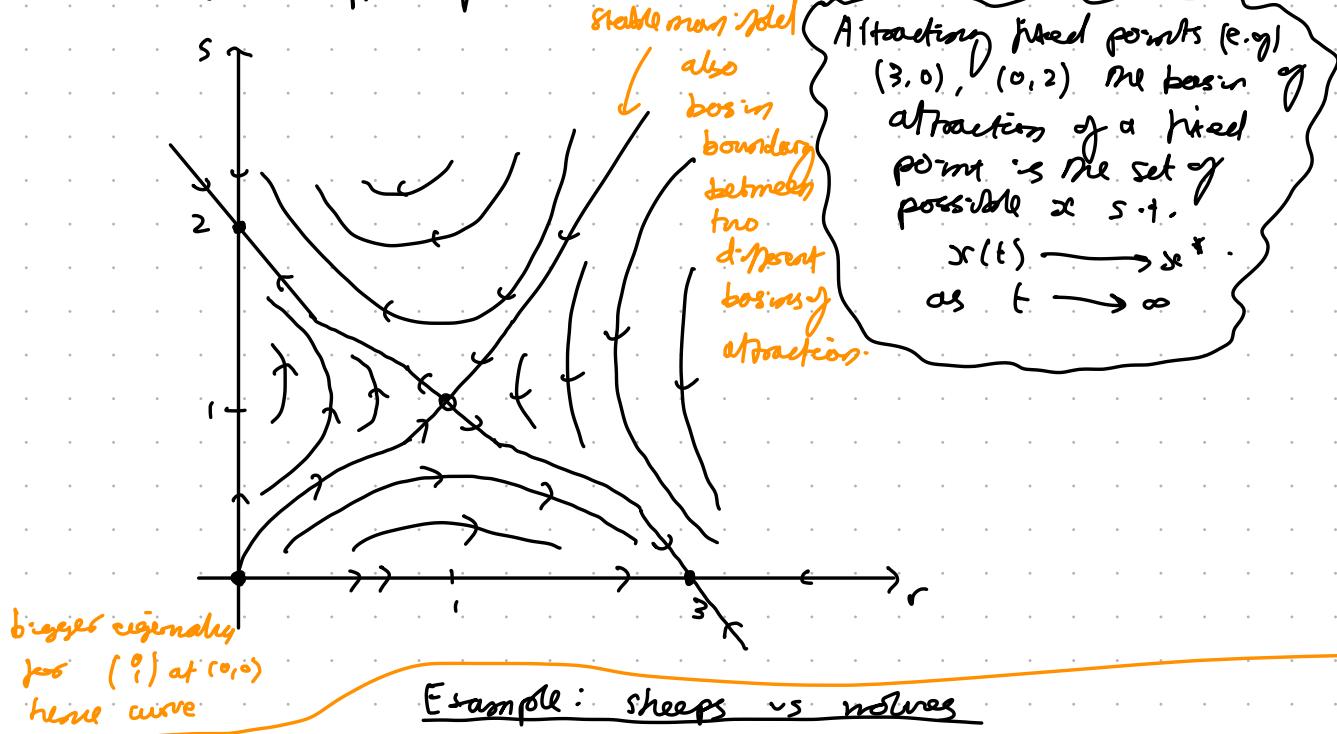
\Rightarrow stable node \because both λ negative

Fixed point $(1, 1)$ $A = \begin{pmatrix} -1 & -2 \\ -1 & -1 \end{pmatrix} \quad \lambda_{1,2} = -\sqrt{2}-1, \sqrt{2}-1$

 $\text{co} > 0 \quad v_{1,2} = \begin{pmatrix} \sqrt{2} \\ 1 \end{pmatrix}, \begin{pmatrix} -\sqrt{2} \\ 1 \end{pmatrix}$

\Rightarrow saddle \because one +ve, one -ve.

Now we do our phase portrait



$w = w(-a + bs) \quad [\text{wolves}]$
 $\dot{s} = s(c - dw) \quad [\text{sheep}]$

$w = \dot{s} = 0 \quad \& \text{ fixed points at } (0, 0), (\frac{c}{d}, \frac{a}{b})$

linearized system: jacobian matrix $A = \begin{pmatrix} -a+bs & bw \\ -ds & c-dw \end{pmatrix}$

Sub in fixed points:

$\textcircled{1} (0, 0) \quad A = \begin{pmatrix} -a & 0 \\ 0 & c \end{pmatrix} \Rightarrow \lambda_{1,2} = -a, c, \quad v_{1,2} = \begin{pmatrix} 1 \\ 0 \end{pmatrix}, \begin{pmatrix} 0 \\ 1 \end{pmatrix}$

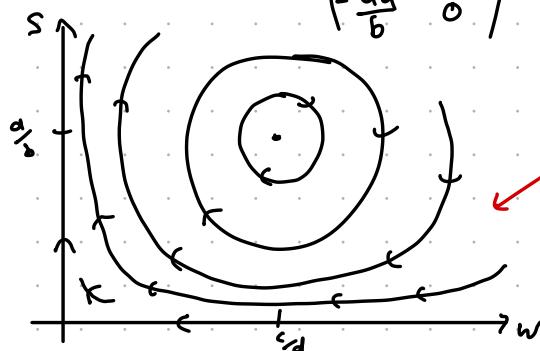
"no wolves so sheep grow exponentially"

stable for wolves,
unstable for sheep.

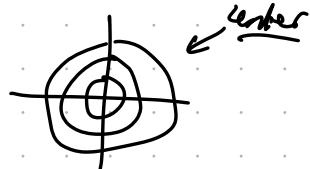
saddle

$\textcircled{2} (\frac{c}{d}, \frac{a}{b}) \quad A = \begin{pmatrix} 0 & \frac{bc}{d} \\ -\frac{da}{b} & 0 \end{pmatrix} \Rightarrow \lambda^2 + ac = 0 \Rightarrow \lambda = \pm i\sqrt{ac}$

so get circular orbits.



get these oscillations.
from this trajectory.



Discrete Time Models

Discrete time can make more sense than continuous time. can get chaos...

1-D maps / difference equations / recursive relations

$x_{n+1} = f(x_n)$, sequence x_0, x_1, x_2, \dots is 1st orbit starting from x_0 .

Def: x^* is a fixed point if $f(x^*) = x^*$

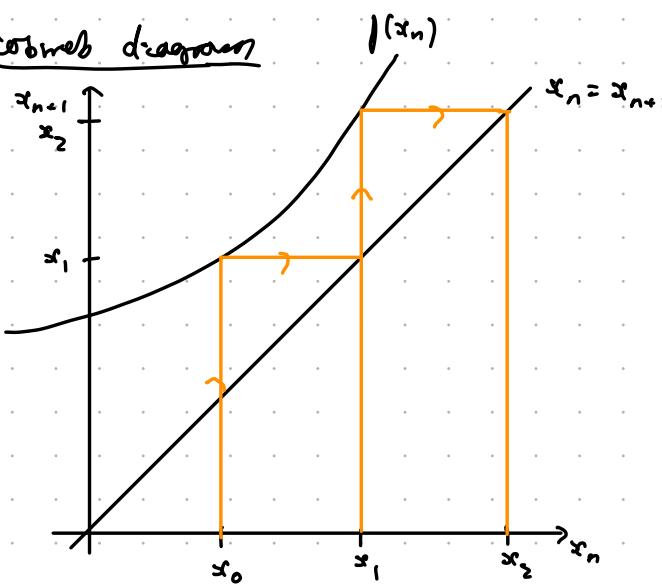
$$x_n = x^* \Rightarrow x_{n+1} = f(x_n) = x^*$$

Stability: write $\lambda = f'(x^*)$ where ' denotes $\frac{dx}{dt}$

- ① If $|\lambda| < 1 \Rightarrow x^*$ is linearly stable
- ② If $|\lambda| > 1 \Rightarrow x^*$ is unstable
- ③ If $|\lambda| = 1 \Rightarrow$ marginal case - not sure...

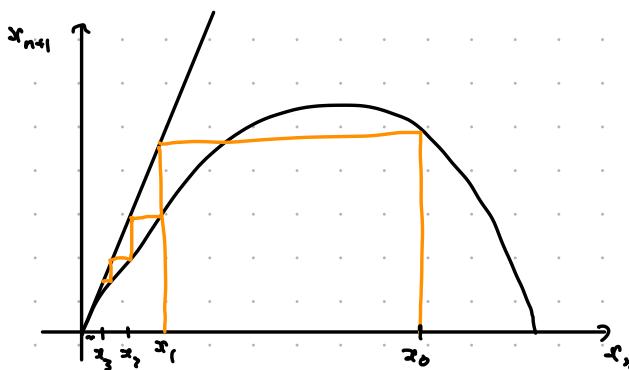
Draw a cobweb diagram to represent this.

cobweb diagram



$$x_{n+1} = f(x_n)$$

Example $f(x_n) = 5\sin x_n$



Logistic Map

$$x_{n+1} = r x_n (1 - x_n)$$

normalized so x is a dimensionless measure

$x=0$ min population
 $x=1$ max population

r is growth rate

$$f(x) = r x (1 - x) = x \quad \leftarrow \text{fixed points}$$

$$\text{If } \boxed{x=0} \text{ fixed, if } x \neq 0, r(1-x)=1 \Rightarrow \boxed{x=1-\frac{1}{r}} \quad \boxed{r>0}$$

Note: this fixed point only exists for $r > 1$
 \Rightarrow bifurcation at $r=1$. new fixed point...

$f(x) = r x (1 - x)$ is a parabola with max at

$$x_n = \frac{r}{4} \quad [\text{when } x_{n-1} = \frac{1}{2}] \Rightarrow \text{care about } 0 \leq r \leq 4 \text{ so } 0 \leq x \leq 1$$

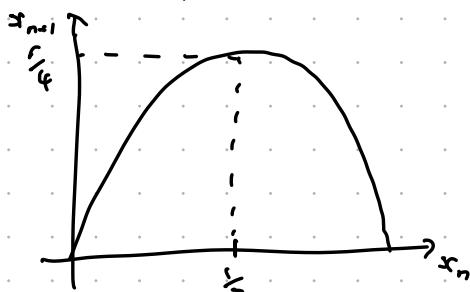
otherwise maximum makes no sense...

Look at stability of fixed points.

If $r < 1$: $x^* = 0$ is fixed point.

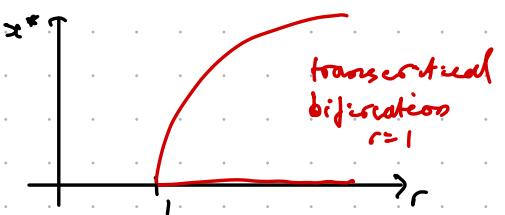
$$f'(x) = r - 2rx \quad |r| < 1$$

$$f'(0) = r < 1 \Rightarrow \text{stable}.$$



$$\text{If } 1 < r < 3: \quad x^* = 0 \quad \& \quad x^* = 1 - \frac{1}{r}$$

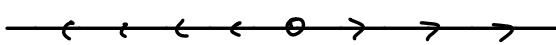
so $f'(0) = r > 1 \Rightarrow \text{unstable}$



$$f'(1 - \frac{1}{r}) = r - 2r + 2 = 2 - r \Rightarrow -1 < 2 - r < 1 \Rightarrow |2 - r| < 1 \Rightarrow \text{stable!}$$

DE

$$\text{If } r > 3:$$



$x^* = 0$ unstable.

$x^* = 1 - \frac{1}{r}$ unstable

} two fixed points & some type of stability.

(cycle / Periodic orbits)

$\exists n \in \mathbb{N} \text{ s.t. } f^n(x_0) = x_0 \text{ & return to start point.}$

"n cycle / periodic orbit of period n"

$$f^2(x) = r f(x)(1 - f(x)) = r^2 x(1-x)(1 - rx(1-x)) \quad \leftarrow \text{quartic!}$$

not $f^2(x) = x \quad \leftarrow \text{analogous to solve...}$

Stay where you start!

For $r > 3$, get one 2 cycle, but very close to 3.

E.g. 2 cycle

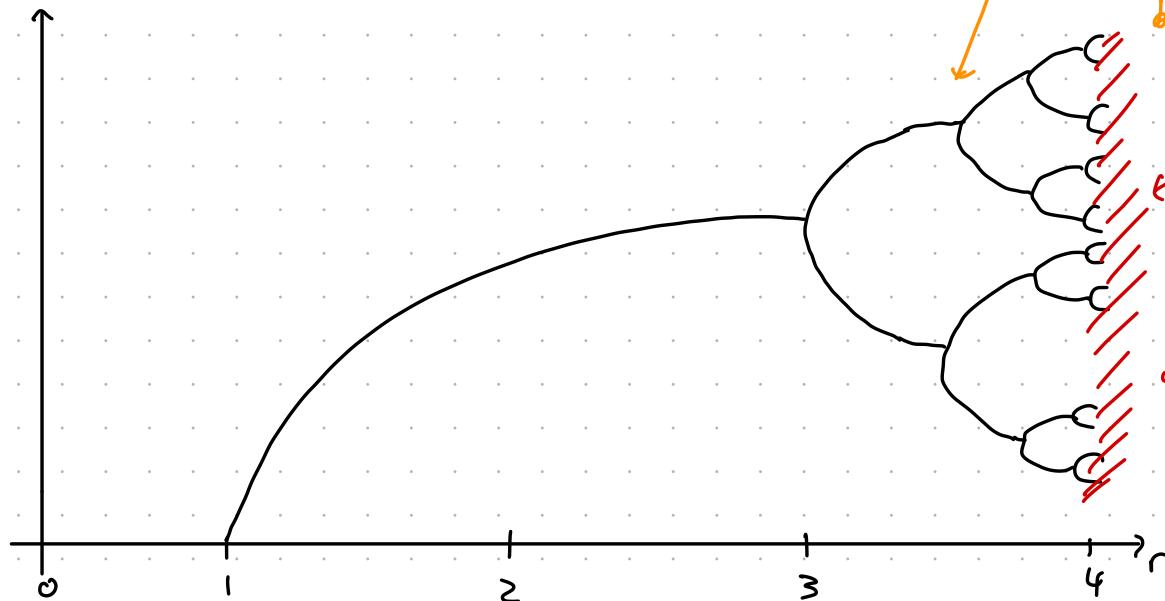
{p, q} s.t.

$$f(p) = q, f(q) = p$$

$$f^2(p) = p.$$

set hands & header to find values...
period 20 orbit is a
true polynomial. too hard.

all fixed
points &
points
that
belong in
a cycle



No bifurcations/period doubling bifurcations

get chaos
as you
approach 4

weird stuff
appearing...

Epidemiological Models

toes up between complexity & ease of solving.

S - susceptible

E - exposed. [have the disease but not symptomatic]

I - infected [and infectious]

R - recovered/removed [might have some immunity]

V - vaccinated individuals

These are the S main compartments might add more based on the biology of the disease

The SIR model

[basic start point]



$$\frac{dS}{dt} = -\beta S I$$

↙ transmission rate
 ↗ become infectious
 ↗ more susceptibles
 ↗ more infected people

$\frac{1}{\gamma}$ = duration of infectiousness

$$\frac{dI}{dt} = \beta S I - \gamma I$$

↗ more into infection

γ = recovery rate

$\beta, \gamma > 0$

$$\frac{dR}{dt} = \gamma I$$

proportion of the population

$$S + I + R = 1$$

everyone belongs to this

$$S(0) > 0, I(0) > 0, R(0) = 0$$

P
people or
susceptible

This is a disease
to start with...

For rate of growth of epidemic, look at

$$\frac{dI}{dt} = \frac{I(\beta S - \gamma)}{\geq 0}$$

for disease to be growing

transmission rate

If $S(0) > \frac{\beta}{\gamma}$ ← critical threshold
The epidemic will grow...

need to keep this smaller than one.

$$\text{If } \gamma > \beta \Rightarrow \frac{\gamma}{\beta} > 1 \Rightarrow \boxed{\frac{\beta}{\gamma} < 1}$$

$\frac{1}{\gamma}$ is duration of infectiousness

Def: The basic reproductive ratio r_0 is the average number of secondary cases arising from an average primary case in an entirely susceptible population.

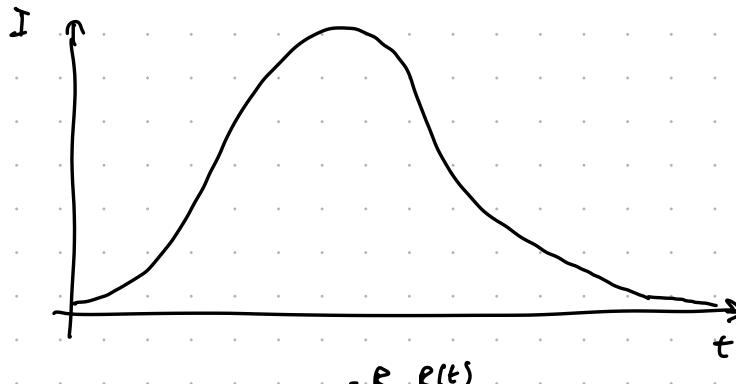
If new disease, previously unseen, then:

$$S(0) \approx 1$$

- $r_0 > 1$: pathogen can invade

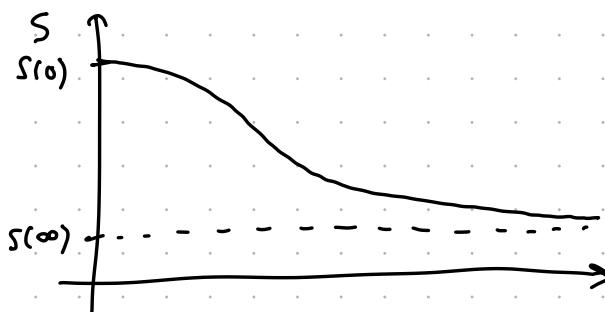
- $r_0 < 1$: pathogen will die out.

In the real world, epidemics look like this.



$$\text{so } S(t) = S(0)e^{-R_0 R(t)}$$

$\rightarrow S(t)$ decreasing since $R(t)$ increasing



$S(\infty) = \lim_{t \rightarrow \infty} S(t)$, so always some susceptible individuals, must have at some point.

$$I(\infty) = 0$$

$$S + I + R = 1 \Rightarrow S(\infty) + I(\infty) + R(\infty) = 1$$

$$\text{so } 1 - R(\infty) = S(\infty)$$

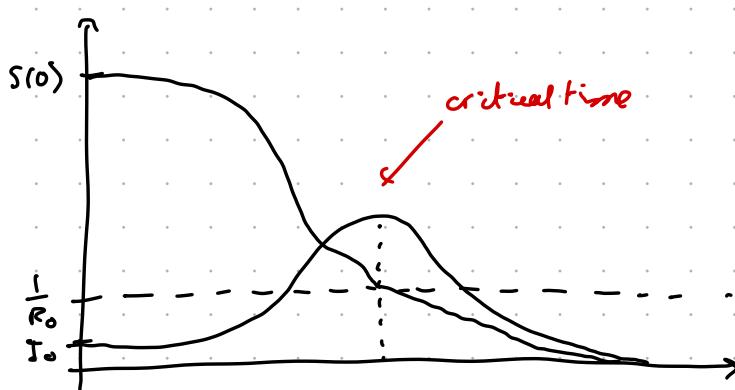
$$\boxed{1 - R(\infty) - S(0)e^{-R_0 R(\infty)} = 0}$$

final epidemic size, final number of people infected...

why do we get epidemic burnout?

Assume $R_0 > 1$ [do get an epidemic], so $\frac{1}{R_0} < 1$

Then $\frac{dI}{dt} > 0 \text{ if } S(0) > \frac{1}{R_0}$



Missing that recovered people can become susceptible again/ add births & deaths into the mix.

only fixed point is at 0 infected. No other fixed points.

SIR with demography

$$\frac{dS}{dt} = -\beta SI - \mu S + M \quad \begin{matrix} \text{death rate} \\ \leftarrow \end{matrix} \quad \frac{1}{\mu} \text{ is average lifespan}$$

$$\frac{dI}{dt} = \beta SI - \gamma I - \mu I$$

$$\frac{dR}{dt} = \gamma I - \mu R$$

γ is death rate
 M is birth rate \times total population
 $S + I + R = 1$
 Births & deaths balance to give constant population

When going?

$$\frac{dI}{dt} = I(\beta S - \gamma - \mu) < 0 \quad \text{if } S < \frac{\mu + \gamma}{\beta}$$

so $R_0 = \frac{\beta}{\mu + \gamma}$ is our threshold for epidemic growth.

Fixed points

$$I=0 \quad \text{or} \quad S = \frac{\mu + \gamma}{\beta} = \frac{1}{R_0}$$

$$\textcircled{1} \quad \underline{I^* = 0} : \quad -\mu S + M = 0 \Rightarrow S^* = 1 \\ \text{Fixed point at } (1, 0, 0)$$

[no infectious people, recovered eventually die off so only susceptibles left]

$$\textcircled{2} \quad S^* = \frac{1}{R_0} : \quad \mu - \beta \left(\frac{\mu + \gamma}{\beta} \right) S - \mu \left(\frac{\mu + \gamma}{\beta} \right) = 0$$

$$\text{so } I^* = \frac{\mu}{\beta} (R_0 - 1) \quad R^* = 1 - S^* - I^*$$

Fixed point at $\left(\frac{1}{R_0}, \frac{\mu}{\beta}(R_0 - 1), 1 - \frac{1}{R_0} - \frac{\mu}{\beta}(R_0 - 1) \right)$ ← endemic equilibrium.

Find jacobian, $\frac{ds}{dt} = f(S, I, R)$, $\frac{dI}{dt} = g(S, I, R)$, $\frac{dR}{dt} = h(S, I, R)$

$$A = \begin{pmatrix} \frac{\partial f}{\partial S} & \frac{\partial f}{\partial I} & \frac{\partial f}{\partial R} \\ \frac{\partial g}{\partial S} & \frac{\partial g}{\partial I} & \frac{\partial g}{\partial R} \\ \frac{\partial h}{\partial S} & \frac{\partial h}{\partial I} & \frac{\partial h}{\partial R} \end{pmatrix} = \begin{pmatrix} -\beta I - \mu & -\beta S & 0 \\ \beta I & \beta S - \mu - \gamma & 0 \\ 0 & \gamma & -\mu \end{pmatrix} \quad \begin{matrix} \text{jacobian} \\ \text{at fixed point} \\ \text{working out} \end{matrix}$$

Now, calculate eigenvalues..

Disease free equilibrium:

$$A = \begin{pmatrix} \mu & \beta & 0 \\ 0 & \beta - \mu - \gamma & 0 \\ 0 & \gamma & \mu \end{pmatrix}$$

$$\lambda_1 = -\mu < 0 \\ \lambda_2 = 0 \\ \lambda_3 = \beta - (\mu + \gamma)$$

Stable if $\lambda_3 < 0$, so $\beta < \mu + \gamma$, $\frac{\beta}{\mu + \gamma} = R_0 < 1$

Endemic equilibrium:

Require $R_0 > 1$, otherwise this won't exist.

Calculate determinant of $A - \lambda I$ before substituting in S & I

$$\det(A - \lambda I_3) = (-\mu - \lambda)(1 - \beta I - \mu - \lambda)(\beta S - (\mu + \gamma) - \lambda) + \beta I \quad \dots$$

$$\lambda_1 = -\mu$$

But, as

S + I + R = 1,
com reduce 3
equations to
one,

Note: $R(0) \neq R_0$

bit dumb!

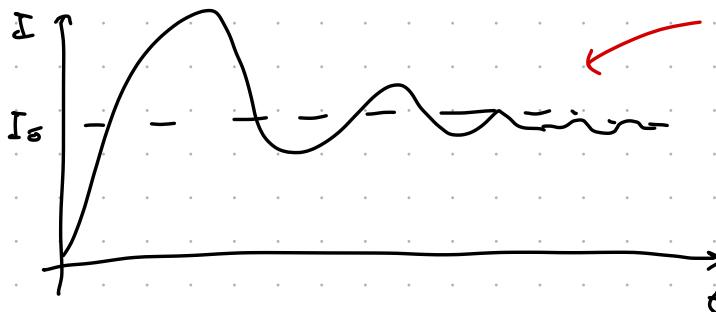
$$\lambda_{2,3} = -\frac{\mu R}{2} \pm \frac{1}{2} \sqrt{(\mu R)^2 - 4(\mu + \gamma)\mu(R_0 - 1)}$$

stable if $\lambda_{1,2,3} < 0$ [all lies below zero]

① if $(\mu R)^2 < 4(\mu + \gamma)\mu(R_0 - 1)$ then get imaginary

$$\therefore \lambda_{2,3} = p \pm iq \quad \text{where } p = -\frac{\mu R_0}{2} < 0$$

\Rightarrow stable, real part negative with oscillations.

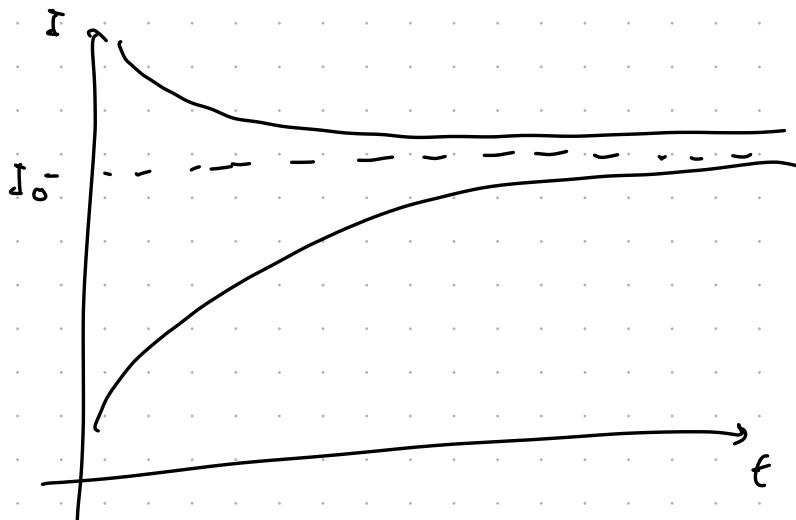


1st & 2nd rows
damped
oscillations.
different cases
for values
of eigenvalues.

② if $(\mu R)^2 > 4(\mu + \gamma)\mu(R_0 - 1) > 0$

$$\Rightarrow \frac{1}{2} \sqrt{(\mu R)^2 - 4(\mu + \gamma)\mu(R_0 - 1)} < \frac{1}{2} \mu R$$

so $\lambda_2, \lambda_3 < 0$, so stable, but no oscillations.



Averaged age for someone getting infected [mean]

Assume death rate $\mu \ll \beta$ [transmissions $\frac{1}{\mu} = 65 \times 365$]

Force of infection = βI $\mu = \frac{1}{65 \times 365}$ lifespan ≈ 1 year

Average time spent in $S = \frac{1}{\beta I}$

At endemic equilibrium, $I^* = \frac{\mu}{\beta} (R_0 - 1)$

$a = \frac{1}{\beta I^*} = \frac{1}{\mu(R_0 - 1)}$ [mean age of infection]

SIS - model

No period of recovery. If you had a cold, can get it again 
 ⇒ only two compartments

$$\frac{dS}{dt} = -\beta SI + \gamma I \quad \left. \begin{array}{l} \\ \end{array} \right] \quad \frac{dS}{dt} = -\frac{dI}{dt} \quad S + I = 1$$

$$\frac{dI}{dt} = \beta SI - \gamma I \quad \left. \begin{array}{l} \\ \end{array} \right] \quad S = 1 - I$$

$$\Rightarrow \frac{dI}{dt} = \beta I(1 - I - \frac{\gamma}{\beta}) = g(I) \quad \text{← look at fixed points...}$$

Stability: If $I^* = 0$, or $I^* = 1 - \frac{\gamma}{\beta}$... fixed points at $(1, 0)$ and $(\frac{1}{R_0}, 1 - \frac{1}{R_0})$

① $I^* = 0$,

$$g'(I) = \beta(1 - \frac{1}{R_0}) - 2\beta I$$

$$g'(0) = \beta(1 - \frac{1}{R_0}) > 0 \quad \text{if } R_0 > 1 \quad [\text{unstable}]$$

② Endemic equilibrium $\Leftrightarrow 0 \quad \text{if } R_0 < 1 \quad [\text{stable}]$

$$g'(\frac{1}{R_0}) = -\beta(1 - \frac{1}{R_0}) < 0 \Rightarrow \text{stable}$$

SEIR model

Exptd: $\frac{1}{\sigma}$ = time taken to become infectious [incubation period]

$$\frac{dS}{dt} = \mu - \beta SI - \mu S \quad S + E + I + R = 1$$

$$\frac{dE}{dt} = \beta SI - \sigma E - \mu E$$

\uparrow people becoming infectious
 \downarrow people can die before passing on disease



$$\frac{dI}{dt} = \sigma E - \gamma I - \mu I$$

$$\frac{dR}{dt} = \gamma I - \mu R$$

How to find R_0 : can calculate, or from definitions.

$$R_0 = \beta \times \left(\frac{\text{time spent infectious}}{\mu + \gamma} \right) \times \left(\frac{\text{proportion that make it to being infectious}}{\frac{\sigma}{\mu + \sigma}} \right)$$

\uparrow transmission rate
 \downarrow $\frac{1}{\mu + \gamma}$
 \downarrow $\frac{\sigma}{\mu + \sigma}$



$$R_0 = \frac{\beta \sigma}{(\mu + \gamma)(\mu + \sigma)}$$

can become infectious & then die

Disease Free Equilibrium: $(1, 0, 0, 0)$

Infection Equilibrium: $\frac{(\mu + \gamma)(\mu + \sigma)}{\beta \sigma} = S^* = \frac{1}{R_0}$

$$\left(\frac{1}{R_0}, \frac{M(\mu + \gamma)}{\beta \sigma}(R_0 - 1), \frac{M}{\beta}(R_0 - 1), 1 - S^* - I^* - E^* \right)$$

Will not exist for 4×4 ^{vector} in our system!

$$A = \begin{pmatrix} -\beta I - M & 0 & -\beta S & 0 \\ \beta I & -(\mu + \sigma) & \beta S & 0 \\ 0 & 0 & -\mu - \gamma & 0 \\ 0 & 0 & \gamma & -M \end{pmatrix} \quad \text{jacobian}$$

Algebraic ...

Eigenvalues: $\lambda_1 = -\mu < 0$,

$$\lambda_2 \approx -(\sigma + \gamma) < 0$$

$$\operatorname{Re}(\lambda_{3,4}) < 0 \quad \text{if } R_0 < 1$$

Stable if $R_0 < 1$

Right structure

high rate & low rate

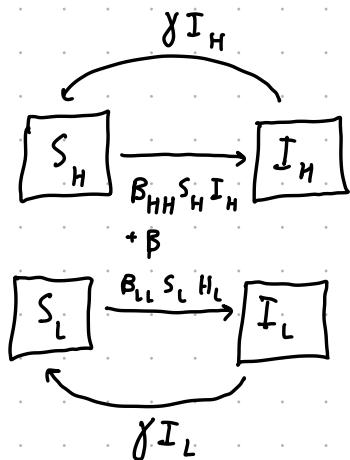
only two types

$n_H + n_L = 1$, transmission rate β_{xy} = transmission rate from y to x

doing the infection

Split into 4 compartments

High rate stay high rate & same w/ low



$$\frac{dS_H}{dt} = -\beta_{HH} S_H I_H - \beta_{HL} S_H I_L + \gamma I$$

$$\frac{dI_H}{dt} = \beta_{HH} S_H I_H + \beta_{KL} S_H I_L - \gamma I$$

$$\frac{dS_L}{dt} = \dots$$

$$\frac{dI_L}{dt} = \dots$$

so $S_H + I_H = n_H$ and $S_L + I_L = n_L$

who acquires infection from whom most? [WAI FW matrix]

$$\begin{pmatrix} \beta_{HH} & \beta_{HL} \\ \beta_{LH} & \beta_{LL} \end{pmatrix} \quad \begin{array}{l} \xleftarrow{\text{infection of H}} \\ \xleftarrow{\text{infection of L}} \end{array}$$

can generally assume
that $\beta_{HL} = \beta_{LH}$

Also $\beta_{HH} > \beta_{LL} > \beta_{KL}$

symmetric interactions...

Example $\begin{pmatrix} 10 & 0.1 \\ 0.1 & 5 \end{pmatrix}$ very little transmission between high resp & low resp

$$n_H = 0.2, n_L = 0.8, \gamma = 1$$

trans. rate is 20%

labeled
is 80%
of populations

$$R_0^H = \frac{\beta_{HH} n_H + \beta_{HL} n_L}{\gamma} = 2.08 > 1$$

$$R_0^L = \frac{\beta_{LL} n_L + \beta_{LH} n_H}{\gamma} = 0.82 < 1$$

Disease free equilibrium

$$S_L = n_L, S_H = n_H$$

$$\dot{J}(I_H, I_L) = \frac{dI_H}{dt} = (\beta_{HH} n_H - \gamma) I_H + \beta_{HL} n_H I_L$$

$$\dot{J}(I_H, I_L) = \frac{dI_L}{dt} = (\beta_{LL} n_L - \gamma) I_L + \beta_{LH} n_L I_H$$

2x2 system of O-Differential Equations...

Jacobian $A = \begin{pmatrix} \beta_{HH} n_H - \gamma & \beta_{HL} n_H \\ \beta_{LH} n_L & \beta_{LL} n_L - \gamma \end{pmatrix} = \begin{pmatrix} 1 & 0.02 \\ 0.08 & -0.2 \end{pmatrix}$

Find eigenvalues. $\lambda_1 = -0.2013, \lambda_2 = 1.0013$

$$e^{\lambda_1 t} + e^{\lambda_2 t}$$

for $t \approx 0$,
longer term
will dominate
at the start...

$\lambda_2 > 0$ dominant eigenvalue \Rightarrow disease is going to invade.

R_0 didn't tell us what was going to happen... This does!

Assuming we know this categories. could have rate depending on age. You then get integral terms etc...

Vaccination

Aim: reduce # susceptibles (proportion), below the critical threshold.
so that $S < \frac{1}{R_0}$ \leftarrow found last time...

Assume: $S(0) + p = 1$ where p is the proportion vaccinated.

$$\text{so want } S - p < \frac{1}{R_0} \Rightarrow p > 1 - \frac{1}{R_0}$$

If $R_0 \geq 1$, need 50% vaccinated to ensure disease can't invade.

Example (Pediatric vaccination)

So vaccinate people at birth & see...

$$\frac{dS}{dt} = \mu(1-p) - \beta SI - \mu S$$

\uparrow
birth rate
 \uparrow
proportion not
vaccinated

$$\frac{dI}{dt} = \beta SI - \gamma I$$

$$\frac{dR}{dt} = \mu p + \gamma I - \mu R$$

\uparrow
those born &
aren't vaccinated
are immediately
recovered!

Change of variables

$$S = S'(1-p)$$

$$I = I'(1-p)$$

$$R = R'(1-p) + p$$

We change the variables to explore the system, setting $S = S'(1-p)$, $I = I'(1-p)$, $R = R'(1-p) + p$. Therefore:

$$(1-p)\frac{dS'}{dt} = \mu(1-p) - (\beta I'(1-p) + \mu)S'(1-p)$$

$$(1-p)\frac{dI'}{dt} = \beta S' I'(1-p)^2 - (\gamma + \mu)I'(1-p)$$

$$(1-p)\frac{dR'}{dt} = \gamma I'(1-p) + \mu p - \mu R'(1-p) - \mu p$$

which reduces to

[SIR model with
 $\beta' = \beta(1-p)$]

$\frac{dS'}{dt} = \mu - (\beta(1-p)I' + \mu)S'$	nice
$\frac{dI'}{dt} = \beta(1-p)S'I' - (\gamma + \mu)I'$	
$\frac{dR'}{dt} = \gamma I' - \mu R'$	

$$\text{Require } R_0 < 1 \Rightarrow (1-p)R_0 < 1 \Rightarrow p > 1 - \frac{1}{R_0} = p_c$$

Measles needs 97%
 vaccinated so depends
 on public behaviour...

critical number of
 vaccinations so that
 disease doesn't break
 out...

Isolations [Q for quarantine]

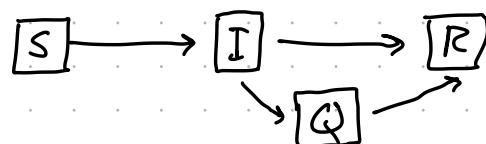
Assume: detection rate of

$$\text{time spent in } Q = \frac{1}{\tau}$$

In whole demography... births - infections - deaths.

$$\frac{dS}{dt} = \mu - \beta IS - \mu S$$

$$\frac{dI}{dt} = \beta IS - dI - \gamma I - \mu I$$



$$\frac{dQ}{dt} = dI - \tau Q (-\mu Q)$$

$$\frac{dR}{dt} = \gamma I + \tau Q - \mu R$$

no deaths in
 quarantine

$$R_0 = \frac{\beta}{\delta + \gamma + \mu} = 1 \Rightarrow d = \beta - \mu - \gamma$$

← critical rate
to increase
infection doesn't
take off...

Setting all $\frac{d}{dt} = 0$, Equilibrium at

$$I^* = 0 \quad \text{or} \quad S^* = \frac{1}{R_0} = \frac{\delta + \gamma + \mu}{\beta}$$

Disease free: $(1, 0, 0, 0)$

Epidemic eq: $\left(\frac{1}{R_0}, \frac{\mu}{\beta}(R_0 - 1), \frac{\delta}{\tau_B}(R_0 - 1), \frac{R_0 - 1}{\beta}(\gamma + \delta) \right)$ if $R_0 > 1$

\uparrow susceptibles \uparrow quarantine

Big assumption: no limitation on ability to quarantine people.

What if we had that $Q = Q_c$ some upper bound on quarantine.

$\{dI \geq \tilde{\epsilon} Q_c\}$

$\frac{dS}{dt}$ stays same.

$$\frac{dI}{dt} = \beta SI - (\gamma + \mu)I - \tilde{\epsilon} Q_c$$

$$\frac{dQ}{dt} = \tau_B Q_c - \tilde{\epsilon} Q_c = 0 \quad \text{so } Q = Q_c \text{ constant!}$$

so $\frac{dR}{dt} = \tilde{\epsilon} Q_c + \gamma I - \mu R$

At threshold, jumps to a different model
so got weird discontinuities

$$R_0^c = \frac{\beta}{\gamma + \frac{\tilde{\epsilon} Q_c}{I^*} + \mu} > R_0$$

← before we hit critical threshold

If $Q < Q_c$ and $R_0 > 1$, we get ① a stable endemic equilibrium
② unstable disease free equilibrium.

If $Q = Q_c$ $\Rightarrow R_0^c > R_0 > 1$, set all equal to zero, get quadratic
in $I = 0$, get one low equilibrium
& one branches

- ① I_{low}^*
- ② I_{high}^*

This means

Discrete Time Models

$S + I + R = N$ \leftarrow population size . Assume no demography (no births & deaths)

Assume a timestep $n = \Delta t$

At each timestep two possible events

- ① $S \rightarrow S-1, I \rightarrow I+1$ [infection]
- ② $I \rightarrow I-1, R \rightarrow R+1$ [recovery]

Write down difference equations

Euler's method: if $\frac{dx}{dt} = f(x)$, then $x_{n+1} = x_n + f(x) \Delta t$

apply
↓

$$g(S_n, I_n, R_n) = S_{n+1} = S_n - \frac{\beta S_n I_n}{N} \Delta t + \mu \Delta t N - \mu \Delta t S_n$$

$$g(S_n, I_n, R_n) = I_{n+1} = I_n + \frac{\beta S_n I_n}{N} \Delta t - (\gamma + \mu) \Delta t I_n$$

$$h(S_n, I_n, R_n) = R_{n+1} = R_n + \gamma \Delta t I_n - \mu \Delta t R_n$$

Susceptibles at timestep $n+1 \rightarrow$

Fixed points: $S_{n+1} = S_n, I_{n+1} = I_n, R_{n+1} = R_n$

Disease free Equilibrium: $S_n^* = N, I_n^*, R_n^* = 0$

Endemic Equilibrium: $S_n^* = \frac{N}{R_0}, I_n^* = \frac{\mu N}{\beta} (R_0 - 1), R_n^* = \frac{\gamma N}{\beta} (R_0 - 1)$

Now want to consider stability. So construct a master equation.

$P_I(t)$ = probability of I infectious individuals at time t

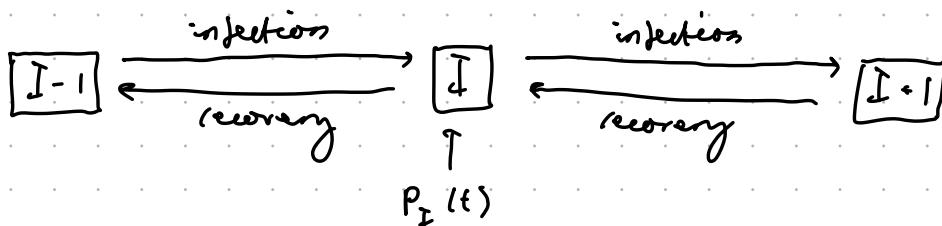
$\{P_I(t)\}_{I=0}^N$ ← this is our distribution...

Example SIS model w/ no demography (no births/deaths)

Two events:

- | | |
|--|--|
| ① $S \rightarrow S-1, I \rightarrow I+1$
infection
rate: $\frac{\beta S I}{N}$ | ② $S \rightarrow S+1, I \rightarrow I-1$
recovery
rate: γI |
|--|--|

4 events / interactions that impact probability that I ppl infections



- ① Situation w/ I infected, has recovery at rate γI additional sickness
- ② Situation w/ $I-1$ ppl infected, has infections at rate $\frac{\beta(S+1)(I-1)}{N}$
 $= \frac{\beta(N-I+1)(F-1)}{N}$
- ③ Situation w/ I ppl infected, has a new infection = $\frac{\beta SI}{N} = \frac{\beta(N-1)I}{N}$
- ④ Situation w/ $I+1$ ppl infected, has recovery at rate $\gamma(I+1)$

Master equations: P_I [rate of event A] = prob of being in state I and event A occurs.

$$\begin{aligned}\frac{dP_I}{dt} = & -P_I[\gamma I] - P_I\left[\frac{\beta(N-1)I}{N}\right] \\ & + P_{I-1}\left[\frac{\beta(N-I+1)(I-1)}{N}\right] + P_{I+1}\left[\gamma(I+1)\right]\end{aligned}$$

We want the distribution of all possible scenarios & their probability

$$\frac{dP_I}{dt} = 0$$

Balance occurs when recovery from $I+1$ balances infections from I

$$P_{I-1}^* [\gamma(I+1)] = P_I^* \left[\frac{\beta(N-I)I}{N} \right]$$

Reasoning

$$P_{I-1}^* = P_I^* \frac{\beta(N-I)I}{N\gamma(I+1)} = P_{I-1}^* \left(\frac{\beta(N-F)I}{N\gamma(F-1)} \right) \left(\frac{\beta(N-I+1)(I-1)}{N\gamma I} \right)$$

I to I-1

keep going & iterating

$$P_{I+1}^* = P_I^* \prod_{j=1}^I \left(\frac{\beta(N-j)j}{N\gamma(j+1)} \right)$$

so can work out equilibrium distribution $\{P_i\}_{i=1}^N$ if we know P_I^*

Simplify:

$$P_I^* = P_1 \frac{(N-1)!}{(N-I)! I} \left(\frac{\beta}{\gamma N}\right)^{I-1}$$

Need P_1 .

Extinction rate: only happens when $I=1 \rightarrow F=0$

rate = γP_1 ← probability only one infectious individual

$$\sum_{I=1}^N P_I = P_1 \sum_{I=1}^N \frac{(N-1)!}{(N-I)! I} \left(\frac{\beta}{\gamma N}\right)^{I-1} = 1 \quad \text{← } R_0$$

Rearrange:

$$P_1 = \left(\sum_{I=1}^N \frac{(N-1)!}{(N-I)! I} R_0^{I-1} \right)^{-1}$$

can now calculate

P_1 so know

the entire distribution
of all possible outcomes.

Extinction rate = γP_1 ,
is rate of extinction for
SIS model without
demography

master equation method is a
powerful way to get any
possible distribution.

Better than DEs :- you can say exactly when does each extinction.

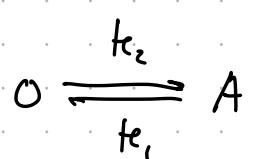
Before, dies out at infinity. Now can say when it ends exactly.

Chapter 3 - Modelling molecular networks

We now consider within-host biological models. Now into the micro scale. Chemical reactions converting one molecule into another.

Simple example

Assume a molecule A is produced & degraded at a constant rate such that



k_1 = production rate

k_2 = degradation rate

Production of A assumed constant over time.

$$\frac{d[A]}{dt} = k_1 - k_2 [A]$$

$[A]$ = concentration of molecule A

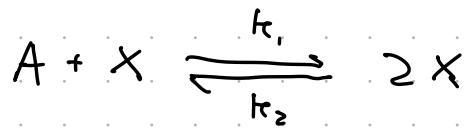
Fixed points occur when $\frac{d[A]}{dt} = 0$. Assume $k_1, k_2 > 0$

$$\text{so equilibrium when } [A] = \frac{k_1}{k_2}$$

$$\begin{aligned} \text{so } \frac{d[A]}{dt} > 0 \text{ when } [A] < \frac{k_1}{k_2} \\ \frac{d[A]}{dt} < 0 \text{ when } [A] > \frac{k_1}{k_2} \end{aligned} \quad \left. \begin{array}{l} \text{fixed point is globally} \\ \text{stable...} \end{array} \right\}$$

Section 3.1 - Autocatalysis

System w/ a molecule X which induces its own production with the addition of a molecule A.



Use law of mass action: States that the rate of a reaction is proportional to

Assume d surplus on A \Rightarrow A's concentration is constant.

This gives us

$$\begin{aligned} j([X]) &= \frac{d[X]}{dt} = k_1 [A][X] - k_2 [X][X] \\ &= [X](k_1 - k_2 [X]) \quad \text{for } k_1 = k_1 [A] \end{aligned}$$

Fixed points occur when $\frac{d[x]}{dt} = 0 \Rightarrow [x] = 0 \text{ or } [x] = \frac{k}{k_2}$

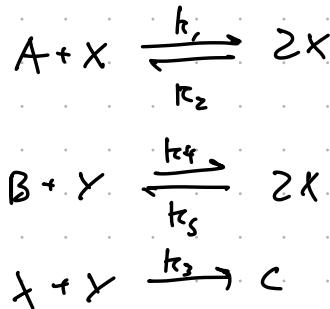
Dynamically:

$$\frac{\partial f}{\partial [x]} = k - 2k_2 [x]$$

When $[x] = 0 \Rightarrow \frac{\partial f}{\partial [x]} > 0 \Rightarrow \text{unstable}$

...

Example: Two molecules X & Y form a complex. Induce their own production in presence of A, B . The system can be described as



$$f([x], [y]) = \frac{d[x]}{dt} = k_1[A][X] - k_2[A][X] - k_3[X][Y]$$

$$\stackrel{\text{by small}}{=} [x](k_1 - k_2[x] - k_3[x])$$

$$k_1 = k_1[x]$$

$$\text{Similarly: } g([x], [y]) = \frac{d[y]}{dt} = [y](k_2 - k_5[y] - k_3[x])$$

Equilibrium when $\frac{d[x]}{dt} = \frac{d[y]}{dt} = 0$

Then find fixed points... (see rates I too tired...)

To find stability, compute one jacobian J

$$J = \begin{pmatrix} \frac{\partial f}{\partial x} & \frac{\partial f}{\partial y} \\ \frac{\partial g}{\partial x} & \frac{\partial g}{\partial y} \end{pmatrix} =$$

Eigenvalues at $(0, 0)$

Stable / unstable directions

law of mass action: rate of chemical reaction.



Rate $r_{12} =$

Example

Species A & B, form a complex C.

