

week 1 - Intro to dynamical systems & stability

A system of ODEs

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

\vdots

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

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

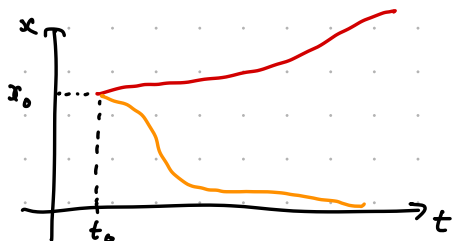
Autonomous systems

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

Simplification!

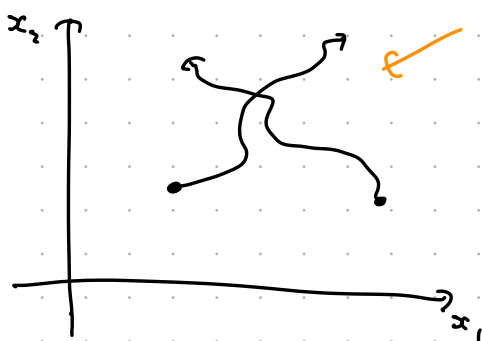
Assume existence and uniqueness of solutions, this means:

• 1D:



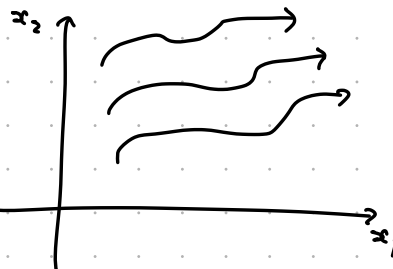
either increasing or decreasing but not both or contradictions.

• 2D:



solutions can't cross in 2d.

(contradicts uniqueness) we get that trajectories never cross



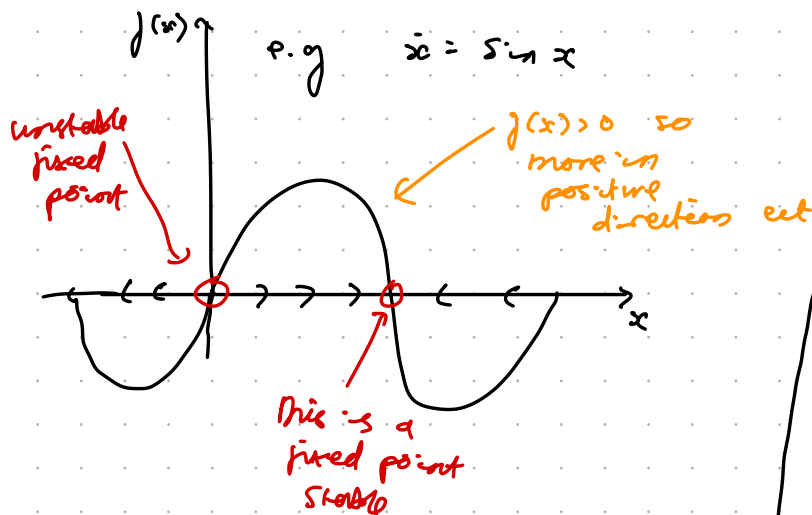
Def: $x_0 \in \mathbb{R}^n$ is an initial condition & the path given by $x(t) \in \mathbb{R}^n$ is called the trajectory corresponding to x_0 .

First order Systems

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

independent variable doesn't appear \therefore autonomous.

e.g. $\dot{x} = \sin x$



"phase line" / "vector field"
phase diagram.

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 minimum rate, then

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

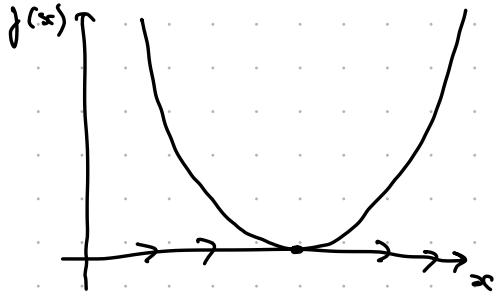
x^* is exponentially stable.

other options!

Saddle node occurs when

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

more complex to classify.



stable from one side & unstable from the other.

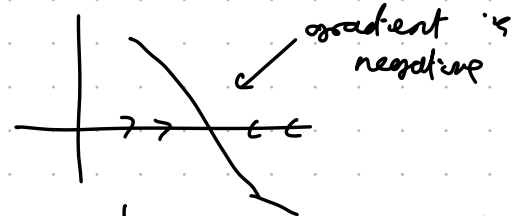
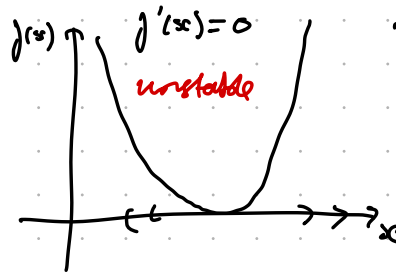
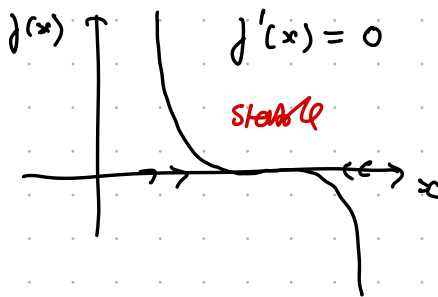
Linear stability Analysis

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, ($N > 0$)
- $r > 0$, $k > 0$

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

$$f(N) = 0 \quad \leftarrow \text{when?}$$

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

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

$$f'(0) = r > 0 \Rightarrow \text{unstable} \quad [\text{no one in population so unstable \& will grow}]$$

$$f'(k) = -r < 0 \Rightarrow \text{stable} \quad [\text{start near carrying capacity, then will stay}]$$

k is carrying capacity

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

[This is the logistic equation]

Bifurcations

change value of a parameter but something qualitatively changes.

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

There are different types.

① saddle node bifurcation:

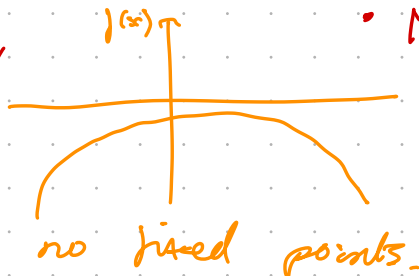
0 fixed points

1 fixed point

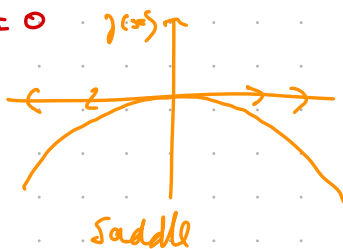
2 fixed points
one stable &
one unstable.

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

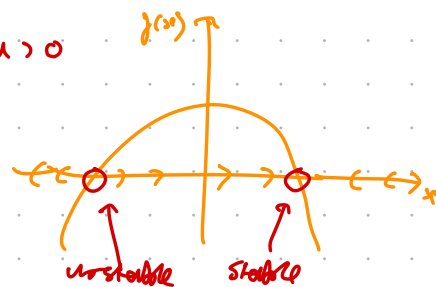
• $\mu < 0$,



• $\mu = 0$



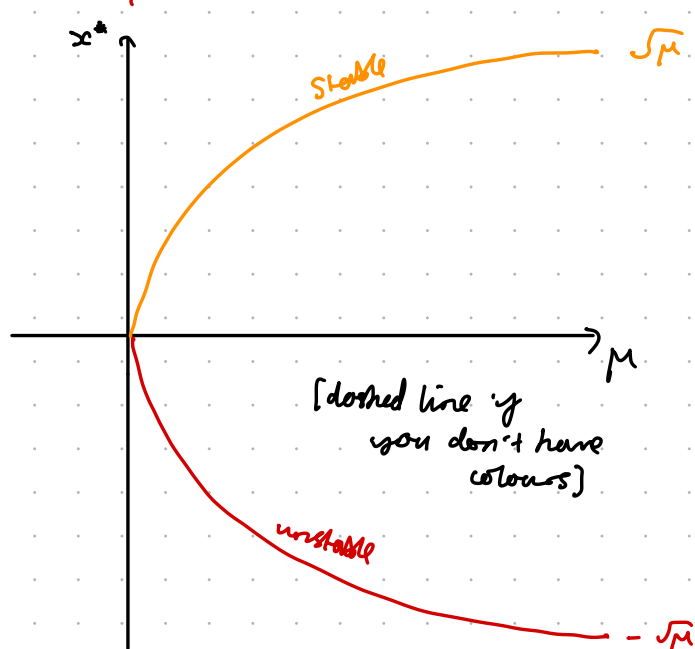
• $\mu > 0$



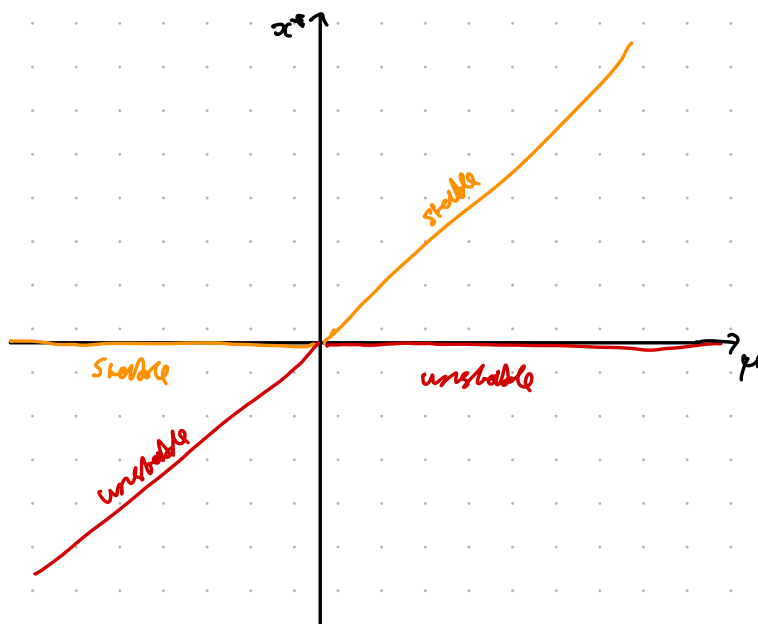
Above is messy, instead, draw a bifurcation diagram, can vary along horizontal axis, then value of fixed points on y

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

$$\sqrt{\mu} = x$$



saddle node bifurcation.



transcritical bifurcation.

② transcritical bifurcation

1x stable

1x unstable

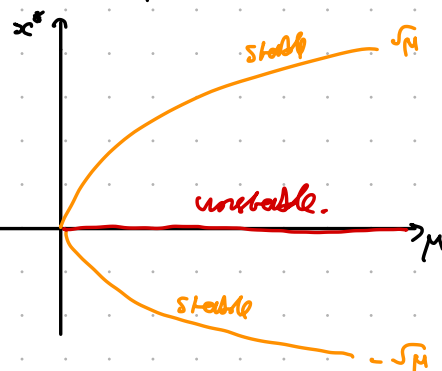
1x saddle

1x unstable

1x stable

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

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



③ Pitchfork bifurcation (looks like one!)

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

$$x^* = 0, x^* = \pm \sqrt{\mu}$$

if $\mu > 0$, if $\mu = 0$ one...

so depending on μ , get different # fixed points.
stable for $\mu > 0$

Second Order Systems

System now because 2.

where will we start from initial point?

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

$$\underline{x}(x, y) = (\dot{x}, \dot{y}) = (f(x, y), g(x, y))$$

zero is always a fixed point in a linear system.

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

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

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

as simple as a system can be

Linear Systems

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

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

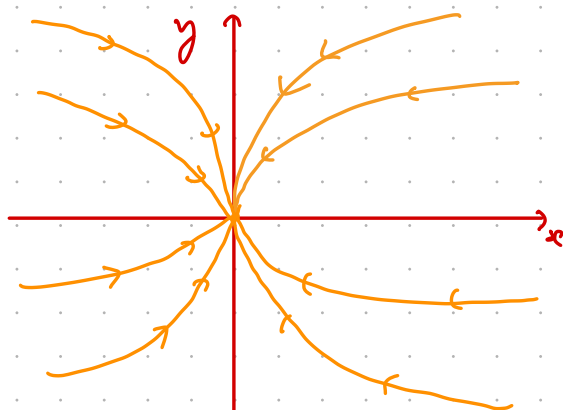
Example: $A = \begin{pmatrix} a & 0 \\ 0 & -1 \end{pmatrix} \Rightarrow \dot{\underline{x}} = A \underline{x} \Rightarrow \boxed{\dot{x} = ax \quad \dot{y} = -y}$

$$x(t) = x_0 e^{at} \quad y(t) = y_0 e^{-t}$$

Now cases:

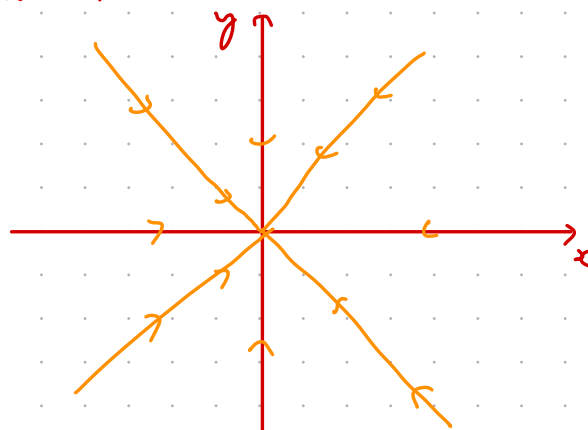
① $a < 0$

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

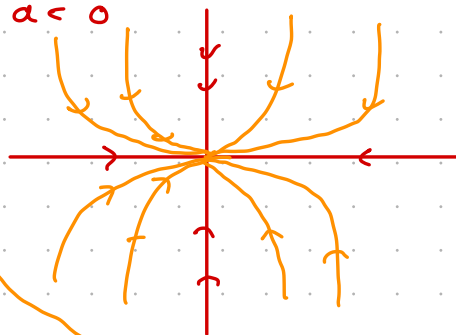


stable node

(ii) $a = -1$ stable star



(iii) $-1 < a < 0$



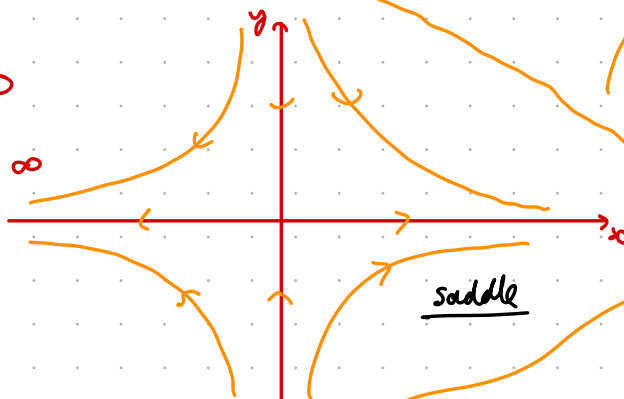
② $a > 0$

$$y(t) \rightarrow 0$$

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

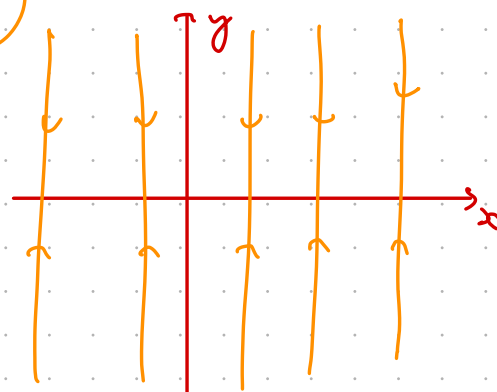
stable manifold is y axis

unstable manifold is x axis



saddle

③ $a = 0$



line of fixed points.

[trajectories back in time to where stable]

WHY NICE?
Eigenvectors are just the axes so nice

General System

$$\dot{x} = Ax, \quad \text{try } x = e^{\lambda t} v$$

can write down a general solution to this system.

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

eigenvalue of A.

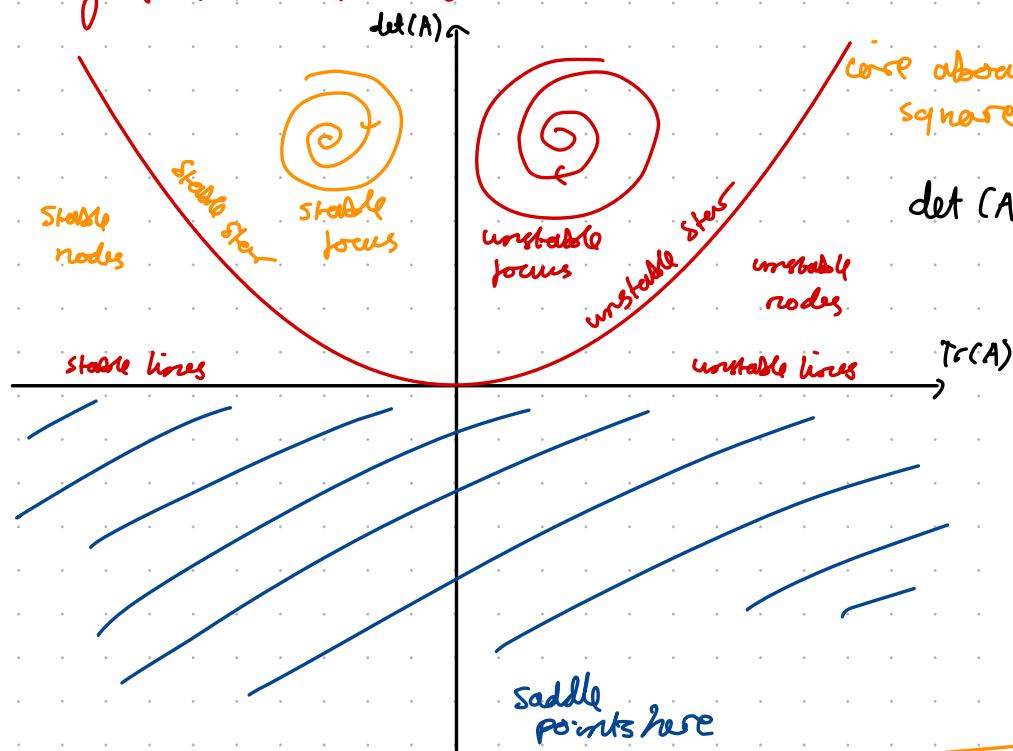
$$x(t) = c_1 e^{\lambda_1 t} v_1 + c_2 e^{\lambda_2 t} v_2$$

v_1, v_2 are eigenvectors of A
 λ_1, λ_2 are eigenvalues.

$$\lambda_1, \lambda_2 \text{ are roots of } \chi(\lambda) = 0 \Rightarrow \lambda^2 - \underbrace{(a+d)}_{\text{trace } A} \lambda + \underbrace{(ad-bc)}_{\det(A)} = 0$$

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

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

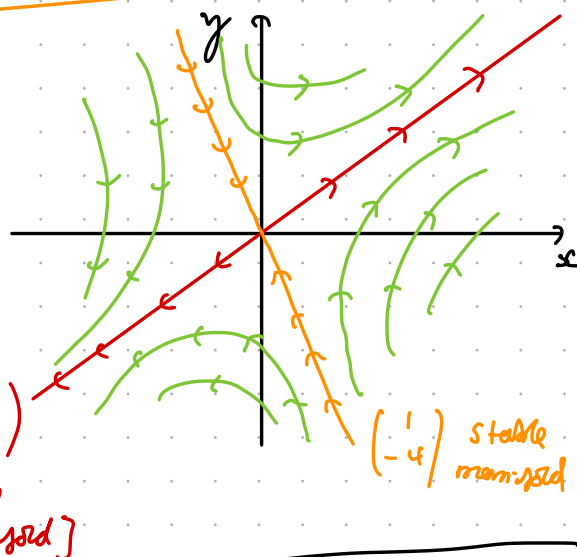


care about the sign under the square root.

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

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]
 $v_{1,2} = \begin{pmatrix} 1 \\ 1 \end{pmatrix}$ & $\begin{pmatrix} 1 \\ -4 \end{pmatrix}$



Non-Linear Systems

$$\dot{x}_1 = f_1(x_1, x_2)$$

$$\dot{x}_2 = f_2(x_1, x_2)$$

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

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

$$\phi(x_0, t)$$

↑ 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.

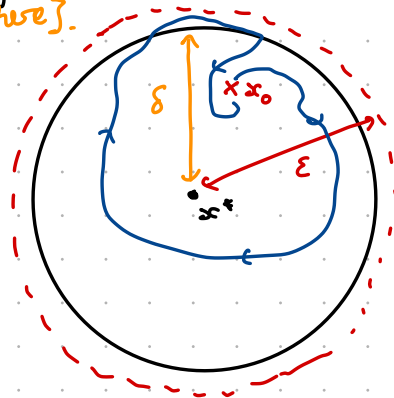
Def: A fixed point x^* is

① An attracting fixed point if all trajectories that start near x^* approach it as $t \rightarrow \infty$

→ It's globally attracting if x^* attracts all trajectories.
[no matter where you start, will always end there].

② Lyapunov stable if $\forall \epsilon > 0 \exists \delta > 0$ s.t. $|x(0) - x^*| < \delta \Rightarrow |x(t) - x^*| < \epsilon \quad \forall t > 0$

↑ where you start
↑ fixed point



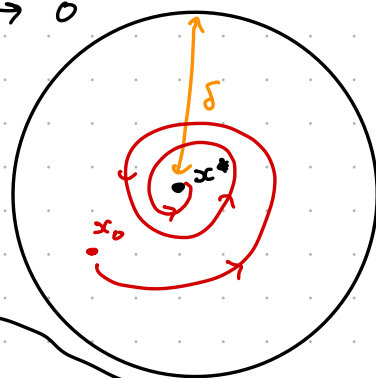
Start close - stay close stability.

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

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

Start within δ of fixed point.

will spiral into fixed point.



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

Linearisation

$$\frac{dx}{dt} = ax + by = f(x, y)$$

$$\dot{\underline{x}} = \begin{pmatrix} a & b \\ c & d \end{pmatrix} \underline{x}$$

$$\frac{dy}{dt} = cx + dy = g(x, y)$$

"the Jacobian"

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

now take this:

$$\frac{dx}{dt} = f(x, y)$$

$$\frac{dy}{dt} = g(x, y)$$

fixed point
 (x^*, y^*)

disturbance

$$u = x - x^*$$

$$v = y - y^*$$

Proof: Taylor's expansion, in notes...

behaviour around fixed points is the same as linear system

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

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 - 'Lottka Volterra'

Example: x prey y predators

(prey) $\frac{dx}{dt} = \alpha x - \beta xy$

growth of prey unhindered.

how much predators eat

proportion of max population $\propto x$

$r_1 \left(\frac{x}{K_1} \right) x, y$

(predators) $\frac{dy}{dt} = -\gamma y + \delta xy$

no food, no prey

infinite prey means predators keep growing

(b) Competing species

$\frac{dx}{dt} = r_1 x \left(1 - \frac{x}{K_1} - \frac{\alpha_1 y}{K_1} \right)$

limits populations $\propto x$

competing population

$\frac{dy}{dt} = r_2 y \left(1 - \frac{y}{K_2} - \frac{\alpha_2 x}{K_2} \right)$

same but reflected.

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

not realistic, real numbers!

$$\begin{aligned} \dot{r} &= r \left(1 - \frac{r}{3} - \frac{2s}{3} \right) \\ \dot{s} &= 2s \left(1 - \frac{s}{2} - \frac{r}{2} \right) \end{aligned} \Rightarrow \begin{aligned} \dot{r} &= r(3-r-2s) = f \\ \dot{s} &= s(2-r-s) = g \end{aligned}$$

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

① check $r=0$: $\dot{s} = s(2-s) = 0 \Rightarrow s=0$ or $s=2$

Fixed points at $(0,0)$ and $(0,2)$

② check $s=0$: $\dot{r} = r(3-r) = 0 \Rightarrow r=0$ or $r=3$

Fixed points at $(0,0)$ and $(3,0)$

③ $\begin{cases} 3-r-2s=0 \\ 2-r-s=0 \end{cases} \rightarrow$ fixed point at $(1,1)$

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

want to linearise system to get A

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

evaluate A at fixed points. Then 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 (should stay here)

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

\Rightarrow stable node \because both λ negative

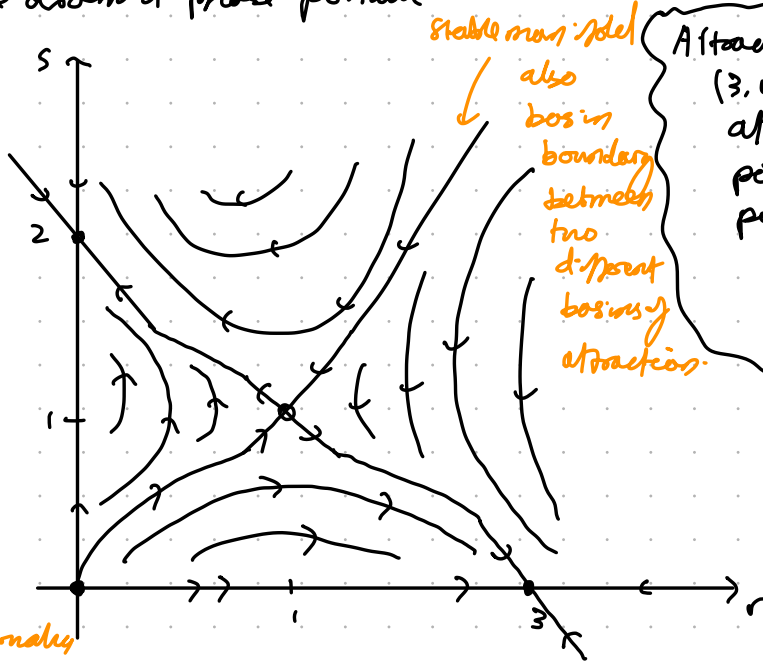
Fixed point (3,0)
just rabbits no sheep
(should stay here)

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

\Rightarrow stable node \because both λ negative

Fixed point (1,1) $A = \begin{pmatrix} -1 & -2 \\ -1 & -1 \end{pmatrix} \quad \lambda_{1,2} = -\sqrt{2}-1, \sqrt{2}-1$
 $\begin{matrix} < 0 & > 0 \end{matrix} \quad v_{1,2} = \begin{pmatrix} \sqrt{2} \\ 1 \end{pmatrix}, \begin{pmatrix} -\sqrt{2} \\ 1 \end{pmatrix}$
 \Rightarrow saddle \because one λ +ve, one -ve.

Now we draw a phase portrait



Attracting fixed points (e.g.) (3,0), (0,2) the basin of attraction of a fixed point is the set of possible x s.t.
 $x(t) \xrightarrow{\text{as } t \rightarrow \infty} x^*$

biggest eigenvalue for (0) at (0,0) hence curve

Example: sheeps vs wolves

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

$$\dot{s} = s(c - dw) \quad [\text{sheep}]$$

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

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

Sub in fixed points:

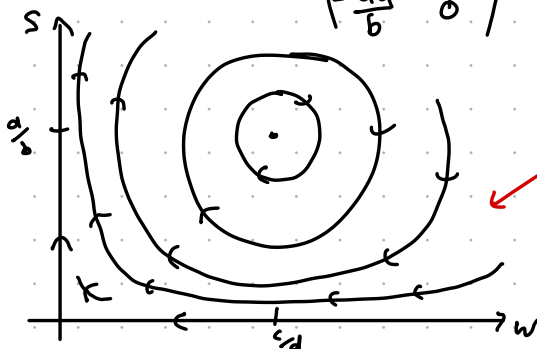
① (0,0) $A = \begin{pmatrix} -a & 0 \\ 0 & c \end{pmatrix} \Rightarrow \lambda_{1,2} = -a, c \quad v_{1,2} = \begin{pmatrix} 1 \\ 0 \end{pmatrix}, \begin{pmatrix} 0 \\ 1 \end{pmatrix}$

"no wolves so sheep grow exponentially"

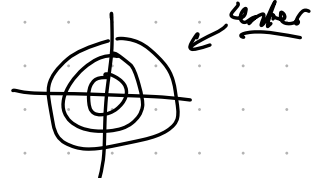
stable for wolves, unstable for sheep.

Saddle

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. lion this trajectory.



jacobian matrix

wolves \uparrow sheep \uparrow

center

Discrete Time Models

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

1-D maps / difference equations / recurrence relations

$x_{n+1} = f(x_n)$, sequence x_0, x_1, x_2, \dots is the 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 $\frac{df}{dx}$

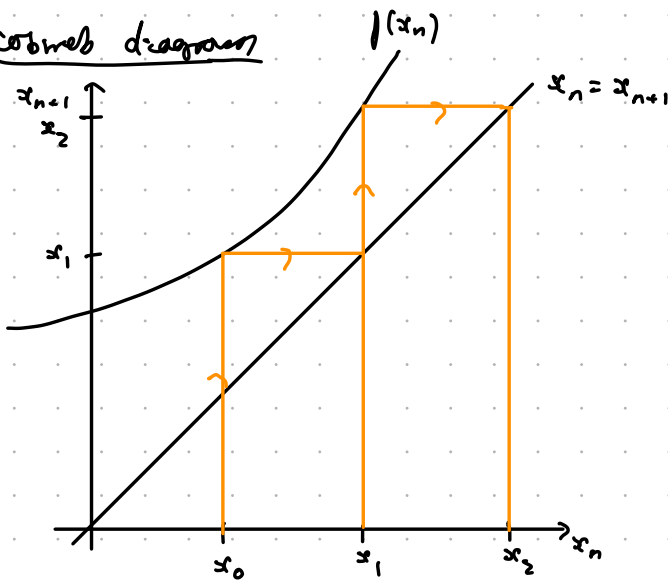
Draw a cobweb diagram to represent this.

① If $|\lambda| < 1 \Rightarrow x^*$ is linearly stable

② If $|\lambda| > 1 \Rightarrow x^*$ is unstable

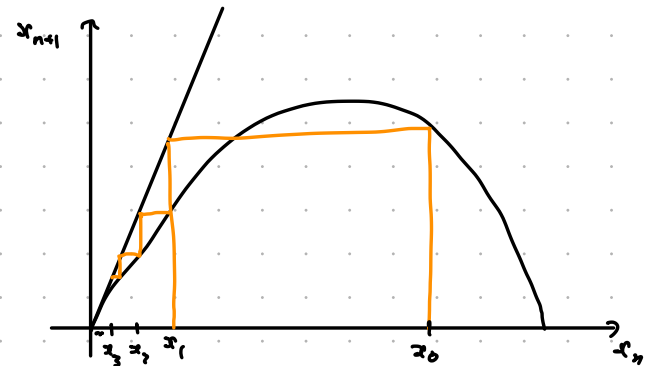
③ If $|\lambda| = 1 \Rightarrow$ marginal case - not sure...

cobweb diagram



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

Example $f(x_n) = \sin x_n$



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

$f(x) = r x (1 - x) = x$ ← fixed points

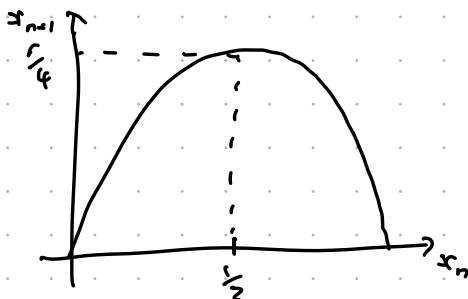
If $x = 0$ fixed, if $x \neq 0$, $r(1-x) = 1 \Rightarrow x = 1 - \frac{1}{r}$

fixed points. $r > 0$

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

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

$x_n = \frac{r}{4}$ [when $x_{n-1} = \frac{1}{2}$] \Rightarrow care about $0 \leq r \leq 4$ so $0 \leq x \leq 1$ otherwise maximum makes no sense...



look at stability of fixed points.

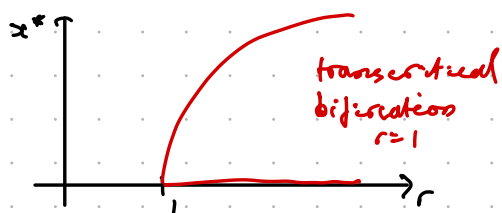
If $r < 1$: $x^* = 0$ is fixed point.

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

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

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

So $f'(0) = r > 1 \Rightarrow$ unstable



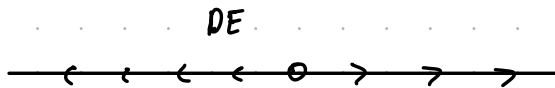
$$f'(1 - \frac{1}{r}) = r - 2r + 2 = 2 - r \Rightarrow -1 < 2 - r < 1 \Rightarrow |2 - r| < 1 \Rightarrow \text{stable!}$$

If $r > 3$:

$x^* = 0$ unstable.

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

} two fixed points & same type of stability.



Cycle / Periodic orbits

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

"n cycle / periodic orbit of period n"

E.g. 2 cycle

$\{p, q\}$ s.t.

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

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

$$f^2(x) = r f(x)(1 - f(x)) = r^2 x(1-x)(1 - rx(1-x))$$

↙ quartic!

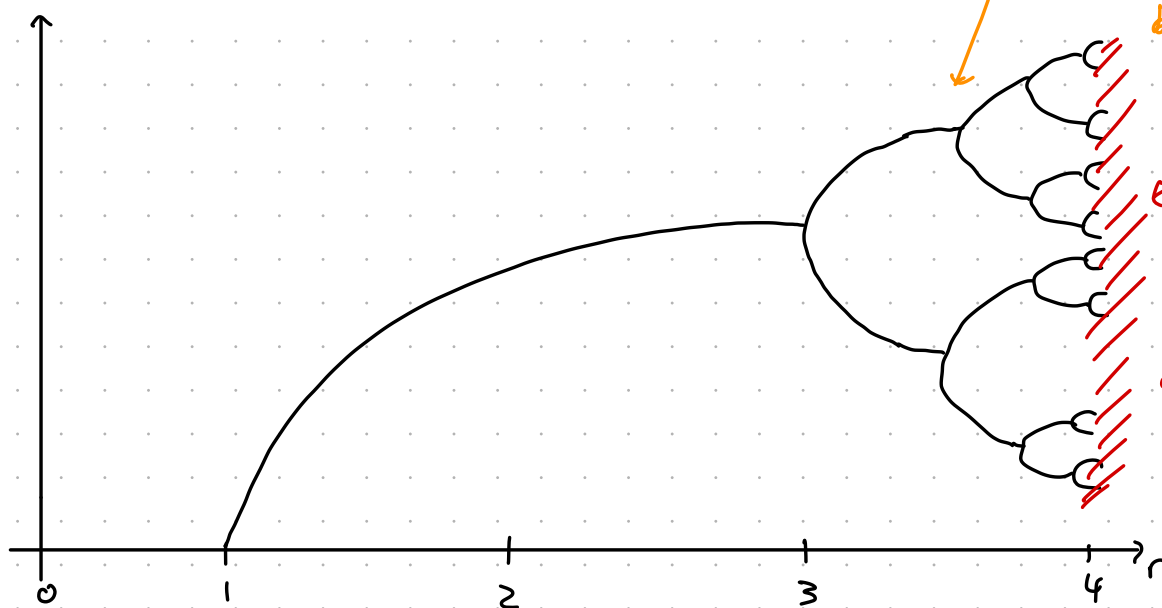
set hands & head
to find values...
period 20 orbit is a
huge polynomial. Too hard.

next $f^2(x) = x$ ↙ annoying to solve...

stay where you start!

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

all fixed points & points that belong in a cycle



tip bifurcation / period doubling bifurcation

get chaos as you approach 4
↓
weird stuff appearing...

Epidemiological Models

ties 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]



$$\frac{dS}{dt} = -\beta SI$$

Annotations for $\frac{dS}{dt} = -\beta SI$:

- β : transmission rate
- S : more susceptible people
- I : more infected people
- $-$: become infectious

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

γ = recovery rate

$\gamma, \beta > 0$

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

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

- βSI : more into infections

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

$$S + I + R = 1$$

proportion of the population

everyone belongs to this

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

people are susceptible

this is a disease to start with...

For rate of growth of epidemic, look at

$$\frac{dI}{dt} = \underbrace{I(\beta S - \gamma)}_{> 0?}$$

for disease to be growing

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

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

$$\boxed{\frac{\beta}{\gamma} = R_0}$$

Annotations for $\frac{\beta}{\gamma} = R_0$:

- β : transmission rate
- $\frac{1}{\gamma}$: duration of infectiousness

need to keep this smaller than one.

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.



$$\frac{dS}{dt} = - \frac{dR}{dt}$$

new 1st order

ODE

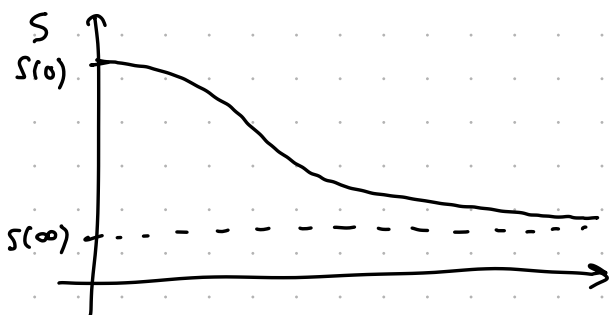
$$\frac{dS}{dR} = - \frac{\beta S I}{\gamma I} = - R_0 S$$

Integrate: $S(R) = A e^{-R_0 R(t)}$
initial conditions,

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

So $S(t) = S(0) e^{-R_0 R(t)}$

$\Rightarrow S(t)$ decreasing since $R(t)$ increasing [$R(0)=0, S(0)>0$]



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

$$I(\infty) = 0$$

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

So $1 - R(\infty) = S(\infty)$

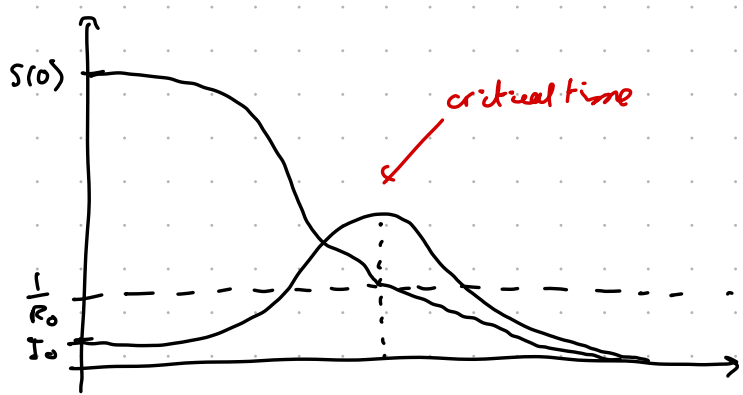
final epidemic size, final number of people infected...

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

Why do we get epidemic burnout?

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

Then $\frac{dI}{dt} > 0$ 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 with births/deaths

death rate

$\frac{1}{\mu}$ is average lifespan

$$\frac{dS}{dt} = -\beta S I - \mu S + \mu \quad \leftarrow \text{birth rate} \times \underbrace{\text{total population}}_{S+I+R=1}$$

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

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

Births & deaths balance to give constant population

When growing?

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

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

Fixed points

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

① $I^* = 0$: $-\mu S + \mu = 0 \Rightarrow S^* = 1$
Fixed point at $(1, 0, 0)$

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

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

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

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

$$A = \begin{pmatrix} \frac{\partial f}{\partial S} & \frac{\partial f}{\partial I} & \frac{\partial f}{\partial R} \\ \frac{\partial g}{\partial S} & \frac{\partial g}{\partial I} & \frac{\partial g}{\partial R} \\ \frac{\partial h}{\partial S} & \frac{\partial h}{\partial I} & \frac{\partial h}{\partial R} \end{pmatrix} = \begin{pmatrix} -\beta I - \mu & -\beta S & 0 \\ \beta I & \beta S - \mu - \gamma & 0 \\ 0 & \gamma & -\mu \end{pmatrix}$$

← jacobian matrix. working out

Now, calculate eigenvalues...

Disease free equilibrium:

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

$$\lambda_1 = -\mu < 0$$

$$\lambda_2 = 0$$

$$\lambda_3 = \beta - (\mu + \gamma)$$

But, as

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

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 substiting in S & I

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

$$\lambda_1 = -\mu$$

note: $R(0) \neq R_0$

but doesn't!

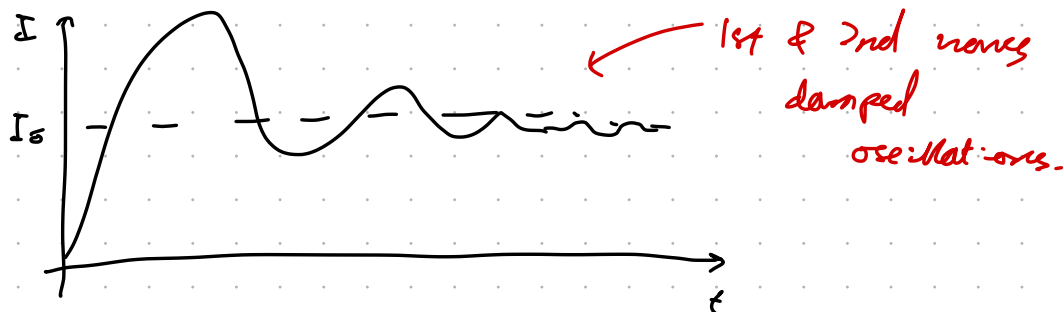
$$\lambda_{2,3} = -\frac{\mu R}{2} \pm \frac{1}{2} \sqrt{(\mu R)^2 - 4(\mu + \gamma)\mu(R_0 - 1)}$$

Stable if $\lambda_{1,2,3} < 0$ [all less than zero]

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

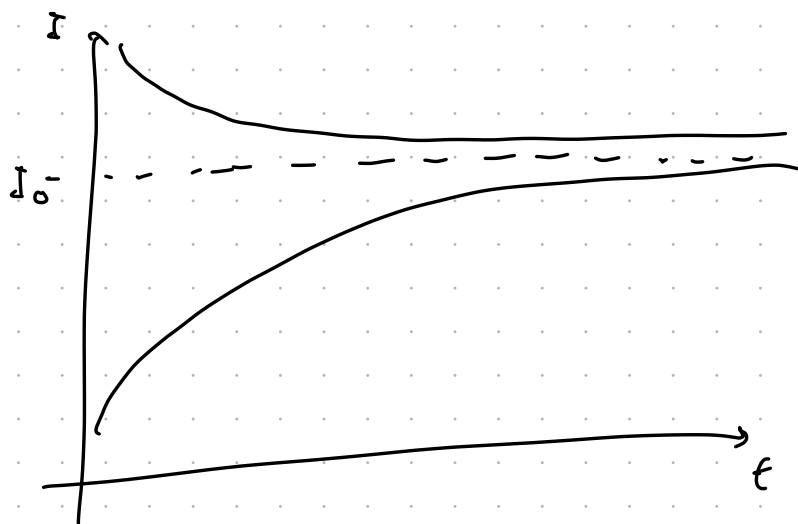


different cases for values of eigenvalues.

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.



Average age for someone getting infected [mean]

Assume death rate $\mu < \beta$

[transmission $\frac{1}{\mu} = 65 \times 365$]

Force of infection = βI


$\mu = \frac{1}{\text{big lifespan}} \sim 1 \text{ year}$

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 
 \Rightarrow only two compartments

$$\left. \begin{aligned} \frac{dS}{dt} &= -\beta SI + \gamma I \\ \frac{dI}{dt} &= \beta SI - \gamma I \end{aligned} \right\} \begin{aligned} \frac{dS}{dt} &= -\frac{dI}{dt} & S+I &= 1 \\ S &= 1-I \end{aligned}$$

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

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 < 0 if $R_0 < 1$ [stable]

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

SEIR model

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

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

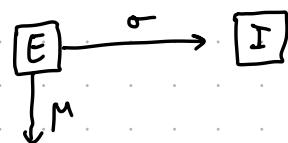
$$\frac{dE}{dt} = \beta SI - \sigma E - \mu E$$

\uparrow deaths
 \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 definition.

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

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

\leftarrow can become infectious & then die

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

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

Endemic 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 ask for λ in an exam!

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

Algebra ...

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

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

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

Stable if $R_0 < 1$

R-str structure

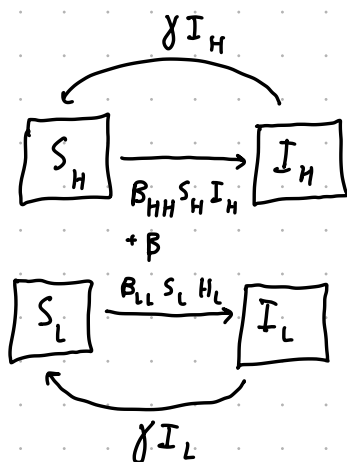
high risk & low risk ← only two types

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

← doing the infecting

Split into 4 compartments

High risk stay high risk & same w/ low



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

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

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

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

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

who acquires infection from whom matrix [WAIFW matrix]

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

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

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

Symmetric interactions...

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

$$\eta_H = 0.2, \quad \eta_L = 0.8, \quad \gamma = 1$$

high risk is 20%

low risk is 80% of population

$$R_0^H = \frac{B_{HH} \eta_H + B_{HL} \eta_L}{\gamma} = 2.08 > 1$$

infection takes off

$$R_0^L = \frac{B_{LL} \eta_L + B_{LH} \eta_H}{\gamma} = 0.82 < 1$$

infection dies off

But no! mixing between populations leads to a takeoff everywhere...

Disease free equilibrium

$$S_L = \eta_L, \quad S_H = \eta_H$$

$$f(I_H, I_L) = \frac{dI_H}{dt} = (B_{HH} \eta_H - \gamma) I_H + B_{HL} \eta_H I_L$$

$$g(I_H, I_L) = \frac{dI_L}{dt} = (B_{LL} \eta_L - \gamma) I_L + B_{LH} \eta_L I_H$$

2x2 system of Differential Equations...

$$\text{Jacobian } A = \begin{pmatrix} B_{HH} \eta_H - \gamma & B_{HL} \eta_H \\ B_{LH} \eta_L & B_{LL} \eta_L - \gamma \end{pmatrix} = \begin{pmatrix} 1 & 0.02 \\ 0.08 & -0.2 \end{pmatrix}$$

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

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

for $t \approx 0$, larger 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 $S < \frac{1}{R_0}$ found last time...

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

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

If $R_0 = 2$, need 50% vaccinated to ensure disease can't invade.

Example (Pediatric vaccination)

so vaccinate people at birth & school...

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

↑
birth rate
↑
proportion not vaccinated

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

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

↑
those born & then 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 $\beta' = \beta(1-p)$]

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

← nice

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

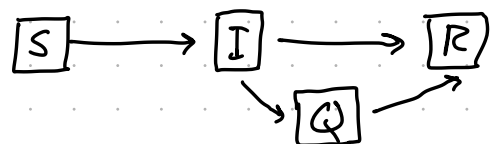
measles needs 97% vaccinated so depends on public behaviour...

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

Isolation [Q for quarantine]

Assume: detection rate d

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



Include 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

$$R_0 = \frac{\beta}{d + \gamma + \mu} = 1 \Rightarrow d = \beta - \mu - \gamma \quad \leftarrow \text{critical rate to ensure infection doesn't take off...}$$

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

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

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

$$\text{Endemic eq: } \left(\frac{1}{R_0}, \frac{\mu}{\beta}(R_0 - 1), \frac{d + \mu}{\tau\beta}(R_0 - 1), \frac{R_0 - 1}{\beta}(\gamma + d) \right) \quad \left[\text{if } R_0 > 1 \right]$$

\uparrow susceptibles

\uparrow quarantine

only exists if $R_0 > 1$

Big assumption: no limitation on ability to quarantine people.

what if we had that $Q = Q_c$ \leftarrow some upper bound on quarantine.

$$dI \geq \tau Q_c$$

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

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

not dependent on # infectious people...

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

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

At threshold, jump to a different model so get weird discontinuities

$$R_0^c = \frac{\beta}{\gamma + \frac{\tau Q_c}{I^*} + \mu} > R_0 \quad \leftarrow \text{before we hit critical threshold}$$

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, set quadratic in $I = 0$, get one low equilibrium & one higher

This means

① I_{low}^*

② I_{high}^*

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
↓ ↓

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

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

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

Successorables at timestep $n+1$

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

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

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

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

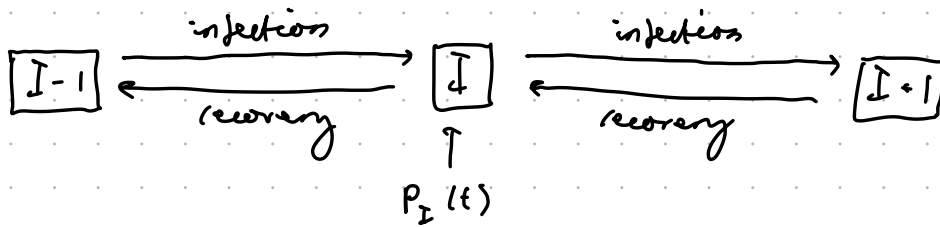
Example SIS model w/ no demography (no births/deaths)

Two events:

① $S \rightarrow S-1, I \rightarrow I+1$
infection
rate: $\frac{\beta SI}{N}$

② $S \rightarrow S+1, I \rightarrow I-1$
recovery
rate: γI

4 events/interactions that impact probability that I ppl infectious



① Situation w/ I infected, has recovery at rate γI additional successful

② Situation w/ I-1 ppl infected, has infection at rate $\frac{\beta(S+1)(I-1)}{N}$
 $= \frac{\beta(N-I+1)(I-1)}{N}$

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

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

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

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

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

keep going & iterating

$$P_{I+1}^* = P_1^* \prod_{j=1}^I \left(\frac{\beta(N-j)j}{N\gamma(j+1)} \right)$$

So can work out equilibrium distribution $\{P_i\}_{i=1}^N$ if we know P_1^*

Simplify:

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

Need P_I .

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

rate = γP_1 ← probability only one infectious individual

$$\underbrace{\sum_{I=1}^N P_I}_{=1} = P_I \sum_{I=1}^N \frac{(N-1)!}{(N-I)! I} \left(\frac{\beta}{\gamma N} \right)^{I-1}$$

$\leftarrow R_0$

Rearrange:

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

can now calculate

P_I 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 distributions.

Better than DEs \because you can say exactly when you reach extinction.

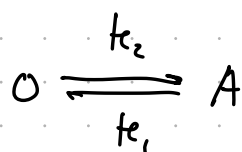
Before, dies out at infinity. Now can say when it ends exactly.

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 $k_1, k_2 > 0$

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

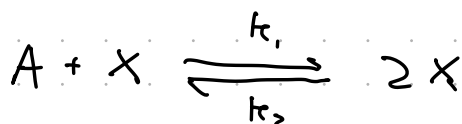
So $\frac{d[A]}{dt} > 0$ when $[A] < \frac{k_1}{k_2}$

$\frac{d[A]}{dt} < 0$ when $[A] > \frac{k_1}{k_2}$

} fixed point is globally stable...

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 a surplus of A \Rightarrow A's concentration is constant.

This gives us

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

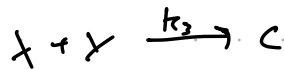
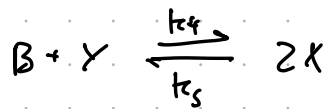
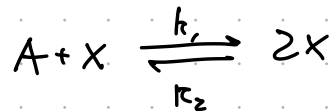
$$= [X](k - k_2[X]) \quad \text{for } k = k_1[A]$$

Fixed points occur when $\frac{d[x]}{dt} = 0 \Rightarrow [x] = 0$ or $[x] = \frac{k_1}{k_2}$

Differentiate: $\frac{\partial f}{\partial [x]} = k_1 - 2k_2 [x]$

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

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



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

$\begin{matrix} \text{big} & \text{small} \\ \nwarrow & \swarrow \\ k_1 = k_1 [x] \end{matrix} = [X][k_1 - k_2 [X] - k_3 [Y]]$

Similarly: $g([x], [y]) = \frac{d[y]}{dt} = [Y](k_4 - k_5 [Y] - k_3 [X])$

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

Then find fixed points... (set rates to zero...)

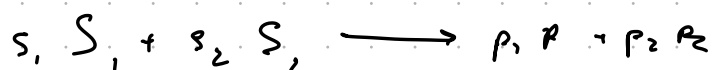
To find stability, compute the jacobian J

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

Eigenvalues at $(0, 0)$

Stable / unstable system

Law of mass action: rate of chemical reactions. —



Rate p_1, p_2 —

Example

Species A, B, form a complex C.

