

# mid problem sheet 4 answers

(1)

1/

$$\dot{H} = \delta_H (H_0 - H) + \lambda_H T_S - \mu_H T_P$$

$$C_a \dot{p}_a = \frac{-(p_a - p_v)}{R_c} + H \Delta V$$

$$c_v \dot{p}_v = \frac{p_a - p_v}{R_c} - \frac{p_v}{R_v}$$

$$T_S = g[p_a(t - \tau)] \quad T_P = 1 - g(p_a)$$

$$g(p) = \frac{1}{1 + \left(\frac{p}{p_0}\right)^n}$$

$$H = H_0 h, \quad p_a = p_0 p, \quad p_v = \frac{R_v p_0}{R_c} \tau \quad t \sim \tau \Rightarrow g = \frac{1}{1 + p^n}$$

$$\Rightarrow \frac{H_0}{\tau} \dot{h} = \delta_H H_0 (1 - h) + \lambda_H T_S - \mu_H T_P$$

$$\Rightarrow \frac{1}{\delta_H \tau} \dot{h} = 1 - h + \frac{\lambda_H}{\delta_H H_0} T_S - \frac{\mu_H}{\delta_H H_0} T_P$$

$$\Rightarrow \varepsilon \dot{h} = 1 - h + \lambda g(p_a) - \mu [1 - g(p)]$$

$$\text{where } \varepsilon = \frac{1}{\delta_H \tau}, \quad \lambda = \frac{\lambda_H}{\delta_H H_0}, \quad \mu = \frac{\mu_H}{\delta_H H_0}$$

Also

2

$$\frac{C_a p_0}{\tau} \dot{p} = - \frac{\left[ p_0 p - \frac{R_v p_0}{R_c} q \right]}{R_c} + H_0 \Delta V h$$

$$\frac{C_v \frac{R_v p_0}{R_c}}{\tau} \dot{q} = \frac{p_0 p - \frac{R_v p_0}{R_c} q}{R_c} - \frac{\frac{R_v p_0}{R_c} q}{R_v}$$

$$\Rightarrow \dot{p} = - \frac{\tau}{R_c C_a} \left( p - \frac{R_v}{R_c} q \right) + \frac{H_0 \Delta V \tau}{C_a p_0} h$$

$$\dot{q} = \frac{\tau}{C_v R_v} \left[ p - \frac{R_v}{R_c} q - q \right]$$

$$\Rightarrow \dot{p} = \kappa \left[ -(p - \delta q) + \sigma h \right]$$

$$\dot{q} = \omega \left[ p - (1 + \delta) q \right]$$

$$\text{if } \omega = \frac{\tau}{C_v R_v}, \delta = \frac{R_v}{R_c}, \kappa = \frac{\tau}{R_c C_a}, \sigma = \frac{H_0 \Delta V R_c}{p_0}$$

$$\delta_H = 1.7 s^{-1}$$

$$\tau = 10 s$$

$$\lambda_H = 0.84 s^{-2}$$

$$\mu_H = 1.17 s^{-2}$$

$$H_0 = 1.7 s^{-1}$$

$$R_v = 0.06 \text{ m}^3 \text{ kg}^{-1} s^{-1}$$

$$R_c = 1.2 \text{ " " " "}$$

$$p_0 = 77 \text{ m}^3 \text{ kg}^{-1}$$

$$C_a = 1.5 \text{ ml m}^3 \text{ kg}^{-1}$$

$$C_v = 50 \text{ ml m}^3 \text{ kg}^{-1}, \Delta V = 70 \text{ ml}$$

$$\Rightarrow \varepsilon = \frac{1}{\delta_H \tau} \sim 0.06 \quad \lambda = \frac{\lambda_H}{\varepsilon_H H_0} \sim \frac{0.84}{2.89} \sim 0.3$$

$$\mu = \frac{\mu_H}{\varepsilon_H H_0} \sim \frac{1.17}{2.89} \sim 0.4, \omega = \frac{\tau}{C_v R_v} \sim \frac{10}{50 \cdot 0.06} \sim 3.3$$

$$\delta = \frac{R_v}{R_c} = \frac{0.06}{1.2} = 0.05, \kappa = \frac{\tau}{R_c C_a} = \frac{10}{1.8} \sim 5.5$$

$$\sigma = \frac{H_0 \Delta V R_c}{p_0} \sim \frac{1.7 \cdot 70 \cdot 1.2}{77} \sim 1.85$$

(3)

Summary

$$\underset{0.06}{\varepsilon} \dot{h} = 1 - h + \underset{0.3}{\lambda} g(p) - \underset{0.4}{\mu} [1 - s(p)]$$

$$\dot{p} = \underset{5.5}{\kappa} [- \underset{0.05}{(p - \delta \varepsilon)} + \underset{1.85}{\sigma h}]$$

$$\dot{q} = \underset{3.3}{\omega} [p - \underset{0.05}{(1 + \delta) \varepsilon}] \quad , \quad g = \frac{1}{1 + p^n}$$

$\delta \ll 1$  so we drop the  $\delta \varepsilon$  term in the  $p$  equation

So  $\delta \downarrow 0$ . Steady state  $\dot{p} \approx \sigma h$

$$1 - h + \lambda g - \mu + \mu g = 0$$

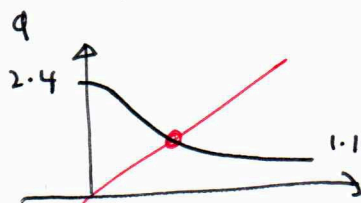
$$\text{so } h = 1 - \mu + (\lambda + \mu)g$$

$$\text{so } p = \sigma(1 - \mu) + \sigma(\lambda + \mu)g$$

$$\equiv q(p) = A + \frac{B}{1 + p^n}$$

$$A = \sigma(1 - \mu) \approx 1.85 \times 0.6 \approx 1.1$$

$$B \approx \sigma(\lambda + \mu) \approx 1.85 \times 0.7 \approx 1.3$$

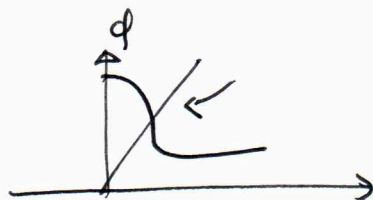


$\hookrightarrow$  unique positive solution

(4)

Which you require depends on  $|\phi'(p^*)|$

1 guess  $n=7$



$|\phi'| > 1$

so try  $p_{n+1} = f^{-1}(p_n)$  etc

$$p_n = A + \frac{B}{1 + p_{n+1}^n}$$

$$\text{so } 1 + p_{n+1}^n = \frac{B}{p_n - A}$$

$$p_{n+1} = \left[ \frac{B}{p_n - A} - 1 \right]^{\frac{1}{n}}$$

(more stable with  $p_0 > A$ )

on a calculator,  $n=7$ ,  $A=1.1$ ,  $B=1.3$

$$p_0 = 1.5$$

$$p_1 = 1.1228 \dots$$

$$p_2 = 1.777 \dots \text{ oops}$$

$$p_3 =$$

$$\text{try } p_{n+1} = A + \frac{B}{1 + p_n^n}, n=7$$

$$p_0 = 1.5$$

$$p_1 = 1.1718 \dots$$

$$p_2 = 1.422 \dots$$

$$p_3 = 1.2018 \dots$$

better

converging slowly.

$$\text{just test } p = A + \frac{B}{1 + p^n} \rightarrow \underline{p_\infty \approx 1.29}$$



(5)

Now if we expect  $p^*$  (already size) = 100 mmHg

$$\Rightarrow p \text{ (mm-d)} = \frac{180}{77} = 1.2987 \dots$$

So then

$$p = A + \frac{B}{1+p^n}$$

$$\Rightarrow 1+p^n = \frac{B}{p-A}$$

$$n \ln p = \ln \left[ \frac{B}{p-A} - 1 \right]$$

$$\text{So } n = \frac{\ln \left[ \frac{B}{p-A} - 1 \right]}{\ln p}$$

$$\text{with } A = 1.1 \\ B = 1.3 \\ \text{and } p = 1.2987$$

$$\Rightarrow \underline{n \approx 6.552}$$

2

(i) Consider  $f(\sigma) = \frac{e^{-\sigma}}{\sigma + \beta} = -\frac{1}{\gamma}$

$f$  has an isolated essential singularity at  $\infty$

$f \neq 0$  anywhere

$\therefore$  Picard  $\Rightarrow f = -\frac{1}{\gamma}$  in any neighbourhood of  $\infty$

(ii) If  $\sigma \rightarrow \infty$

then  $-ye^{-\sigma} \rightarrow \infty \Rightarrow \operatorname{Re} \sigma \rightarrow -\infty$

(iii)  $\sigma = -\beta - ye^{-\sigma}$

$$\frac{d\sigma}{dy} = \sigma' = -e^{-\sigma} + ye^{-\sigma} \sigma'$$

$$\Rightarrow \sigma' = \frac{-e^{-\sigma}}{1 - ye^{-\sigma}} = \frac{-ye^{-\sigma}}{y[1 - ye^{-\sigma}]} = \frac{\sigma + \beta}{y[1 + \beta + \sigma]}$$

or:  $\gamma(\sigma)$  is analytic  $\Rightarrow \sigma(\gamma)$  is also except where  $\gamma' = 0$

(iv)  $\gamma \ll 1$  [If  $\sigma \approx 0(1)$   $\sigma \approx -\beta$ ]

Alternatively Suppose  $\operatorname{Re} \sigma > 0$

then  $\sigma + \beta = -ye^{-\sigma}$

$$|\sigma + \beta| > \operatorname{Re} \sigma + \beta > \beta$$

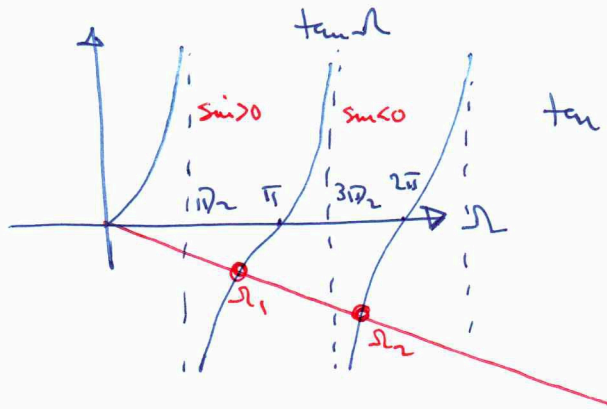
$$\Rightarrow \beta < \gamma |e^{-\sigma}| = \gamma e^{-\operatorname{Re} \sigma} < \gamma$$

$$\Rightarrow \operatorname{Re} \sigma < 0 \text{ for } \gamma < \beta.$$

(v) Hence instability occurs if  $\sigma = i\Omega$  (not  $\sigma = 0$  as never a root)

$$\Rightarrow i\Omega = -\beta - \gamma(\cos \Omega - i \sin \Omega)$$

$$\Rightarrow \gamma = \frac{\Omega}{\sin \Omega}, \quad -\beta = \gamma \cos \Omega = \frac{\Omega}{\tan \Omega}$$



$$\tan \Omega = -\frac{\Omega}{\beta}$$

(7)

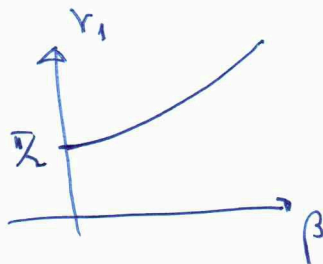
$\infty$  roots  $\Omega_1 \in (\pi/2, \pi)$   
 $\Omega_2 \in (3\pi/2, 2\pi)$   
 $\Omega_3 \in (5\pi/2, 3\pi)$  etc.

of which  $\gamma_k > 0$  for  $k=1, 3, \dots$

[For large  $k$  odd  $\Omega_k \approx (k - \frac{1}{2})\pi$

$$\gamma_k \approx (k - \frac{1}{2})\pi$$

Instability first at  $\gamma_1$



[see fig 6.11 of notes]

To check transversality:

$$\sigma' = \frac{\beta + i\Omega}{\gamma(1 + \beta + i\Omega)}$$

$$= \frac{(\beta + i\Omega)(1 + \beta - i\Omega)}{\gamma[(1 + \beta)^2 + \Omega^2]}$$

$$= \frac{\beta(1 + \beta) + \Omega^2 + i\Omega}{\gamma[(1 + \beta)^2 + \Omega^2]}$$

$\text{Re } \sigma' > 0$  for  $\beta, \gamma > 0$

so roots cross successively & remain unstable.

$$\gamma_{2n+1} \approx \text{Roots near } (2n + \frac{1}{2})\pi \quad n=0, 1, \dots$$

[Note  $\tan \Omega = -\frac{\Omega}{\beta}$

$$\Rightarrow 1 + \frac{\Omega^2}{\beta^2} = 1 + \tan^2 \Omega = \sec^2 \Omega$$

$$\cos^2 \Omega = \frac{\beta^2}{\beta^2 + \Omega^2}$$

$$\sin^2 \Omega = \frac{\Omega^2}{\beta^2 + \Omega^2}$$

$$\sin \Omega = \frac{\Omega}{(\beta^2 + \Omega^2)^{1/2}}$$

$$\gamma = (\beta^2 + \Omega^2)^{1/2} \quad \text{increases with } \Omega$$

[Note:

$$\gamma = \frac{\Omega}{\sin \Omega}, \quad \frac{\Omega}{\tan \Omega} = -\beta \Rightarrow -\frac{\beta}{\gamma} = \cos \Omega$$

$$\gamma'(\beta) = \frac{\gamma'(\Omega)}{\beta'(\Omega)} = \frac{\frac{1}{\sin \Omega} - \frac{\Omega \cos \Omega}{\sin^2 \Omega}}{-[\frac{1}{\tan \Omega} - \frac{\Omega \sec^2 \Omega}{\tan^2 \Omega}]}$$

$$= \frac{\sin \Omega - \Omega \cos \Omega}{-[\cos \Omega - \Omega]} = \frac{1 + \beta}{-[\cos \Omega - \frac{\Omega}{\gamma}]}$$

$$\Rightarrow \gamma' = \frac{\gamma(1 + \beta)}{\beta + \gamma^2} > 0$$

from graph at top,  $\beta \rightarrow 0, \Omega_1 \rightarrow \pi/2, \gamma_1 \rightarrow \pi/2$

$$\beta \rightarrow \infty, \Omega_1 \rightarrow \pi, \Omega_1 = \pi - \delta, \frac{\pi}{\delta} \approx \beta$$

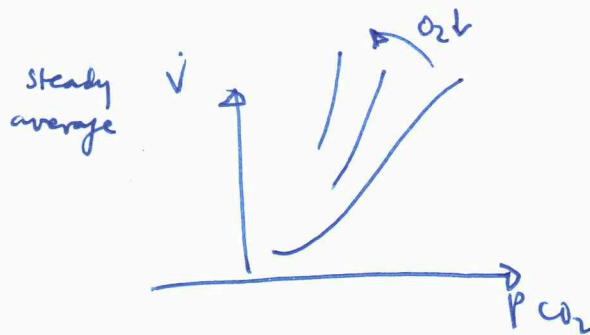
$$\gamma_1 = \frac{\pi}{\delta} \Rightarrow \gamma_1 \sim \beta \text{ as } \beta \rightarrow \infty$$

### 3. Minute ventilation:

volume of air breathed in a minute

Central chemoreceptor responds to  $H^+$  & thus effectively to  $CO_2$  via the bicarbonate buffering system.

Peripheral chemoreceptors also respond to  $CO_2$  but the response is amplified if  $O_2$  is lowered



Haldane-Glass

$$K \dot{V} = M - p \dot{V}$$

$$\dot{V} = \dot{V}(p_r)$$

$K$  tissue volume

$M$  metabolic production rate

$\dot{V}$  ventilation

1st eq. conservation of  $CO_2$

2nd controller

$$\dot{V} = G[p - p_0]_+$$

$$M = 200 \text{ mm Hg l (BTFS) min}^{-1}$$

$$p_0 = 35 \text{ mm Hg}$$

$$K = 40 \text{ l (BTFS)}$$

$$G = 2 \text{ l (BTFS) min}^{-1} \text{ mmHg}^{-1}$$

$$\tau = 0.2 \text{ min}$$

$$K \dot{p} = M - \overbrace{p G (p - p_0)}^{\dot{V}}_+$$

Define  $p = p_0 + \Delta p$   $\Rightarrow v = [\Delta p]_+$

$$\dot{V} = G \Delta p v$$

$$t \sim \tau$$

and (drop  $\dot{p}$ )

$$\frac{K \Delta p}{\tau} \dot{p} = M - \underbrace{p_0 \left(1 + \frac{\Delta p}{p_0} p\right)}_{\dot{V}} G \Delta p v$$

choose  $M = p_0 G \Delta p$  i.e.  $\Delta p = \frac{M}{p_0 G} \sim \frac{200 \text{ mmHg l min}^{-1}}{35 \text{ mmHg} \cdot 2 \text{ l min}^{-1} \text{ mmHg}^{-1}} \approx 3 \text{ mmHg}$

$$\mu = \frac{\Delta p}{p_0} = \frac{M}{p_0^2 G}$$

then  $\dot{p} = \frac{M \tau}{K \Delta p} [1 - (1 + \mu p) v]$

$$= \alpha [1 - (1 + \mu p) v]$$

$$\alpha = \frac{M \tau}{K \Delta p} = \frac{G p_0 \tau}{K}$$

so  $\alpha \sim \frac{2 \text{ l min}^{-1} \text{ mmHg}^{-1} \cdot 35 \text{ mmHg} \cdot 0.2 \text{ min}}{40 \text{ l}}$

$$\approx \frac{1}{3} \quad \& \quad \mu \approx \frac{3}{35} \approx 0.08$$

conversion factor from wet saturated body temperature to dry NTP:  $863 = 760 \times \frac{310}{273}$  (10)

4

$$K_L \dot{P}_{aCO_2} = -\dot{V} P_{aCO_2} + 863 K_{CO_2} Q [P_{vCO_2} - P_{aCO_2}] \quad (1)$$

lung volume      ventilatory loss      solubility coefficient      blood flow      venous      arterial

$$K_{CO_2} K_B \dot{P}_{bCO_2} = MR_{TbCO_2} + K_{CO_2} Q_B [P_{aCO_2}(t - \tau_{aB}) - P_{bCO_2}] \quad (2)$$

brain volume      production rate in ~~brain~~ brain      brain blood flow      arterial delay to brain

$$K_{CO_2} K_T \dot{P}_{tCO_2} = MR_{TtCO_2} + (Q - Q_B) K_{CO_2} [P_{aCO_2}(t - \tau_{aT}) - P_{tCO_2}] \quad (3)$$

tissue volume      tissue production rate      delay to tissues

$$Q P_{vCO_2} = Q_B P_{bCO_2}(t - \tau_{vB}) + (Q - Q_B) P_{aCO_2}(t - \tau_{aT}) \quad (4)$$

delay from tissues

(1): concentration of arterial/lung  $CO_2$

(2): .. brain  $CO_2$

(3): .. tissue  $CO_2$

(4): venous blood flow  $CO_2$  is the sum of the flow from the brain & that from other tissues

Response time scales (green time)

$$(1) \quad t_a = \frac{K_L}{863 K_{CO_2} Q} \quad (\text{since } 863 K_{CO_2} Q \gg \dot{V})$$

$\sim \frac{3}{26} \approx 0.12 \text{ min}$

$$(2) \quad t_B = \frac{K_B}{Q_B} \sim \frac{1}{0.75} = \frac{4}{3} \text{ min} = 80 \text{ s}$$

$$(3) \quad t_T = \frac{K_T}{Q} \sim \frac{39}{6} \sim 6.5 \text{ min}$$



Therefore on a time scale of a minute  $P_{aCO_2}$  is rapid  $\rightarrow$

$P_{aCO_2} \rightarrow$  quasi-equilibrium.

$t_T$  is slow  $\Rightarrow P_{TCO_2}$  changes slowly

$$(4) \Rightarrow P_{aCO_2} = P_{TCO_2} + \frac{Q_B}{Q} [P_{BCO_2} - P_{TCO_2}]$$

and since  $\frac{Q_B}{Q} \approx 0.15$ ,

$P_{aCO_2} \approx P_{TCO_2}$  is slowly varying

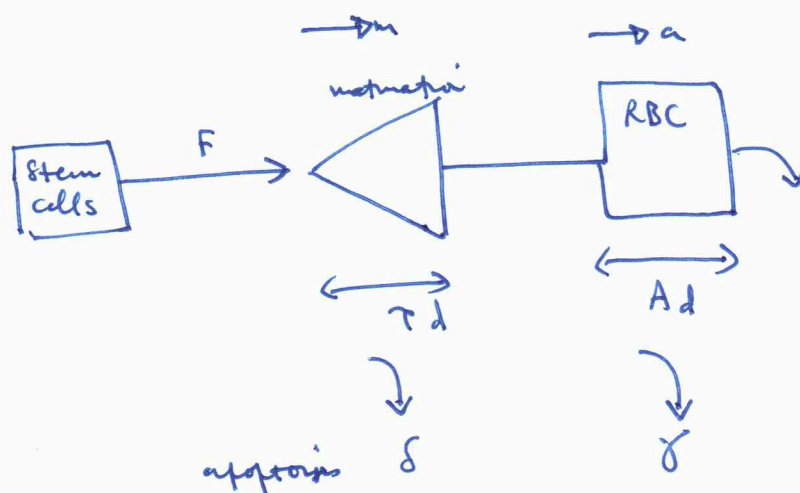
but also 
$$-\frac{\dot{V}}{863 K_{CO_2} Q} P_{aCO_2} + P_{VCO_2} - P_{aCO_2} \approx 0$$

$$\Rightarrow P_{aCO_2} \approx \frac{P_{VCO_2}}{1 + \frac{\dot{V}}{863 K_{CO_2} Q}} \approx \frac{P_{TCO_2}}{1 + \frac{\dot{V}}{863 K_{CO_2} Q}} \quad \text{gives } P_{aCO_2}$$

~~$P_{aCO_2}$  is slowly varying~~

~~Not always  $P_{aCO_2} \approx P_{TCO_2}$  is all exactly equal~~

5/



Let  $p(t, m)$  be the proliferative cell density ( $m$  is maturation time)

$$\Rightarrow \frac{\partial p}{\partial t} + \frac{\partial p}{\partial m} = -\delta p, \quad 0 < m < \tau$$

Let  $e(t, a)$  be circulating RBC density as function of age  $a$

$$\Rightarrow \frac{\partial e}{\partial t} + \frac{\partial e}{\partial a} = -\gamma e, \quad 0 < a < A$$

Now with  $P = \int_0^\tau p \, dm$

$$\text{we have } \frac{dP}{dt} + p|_{m=\tau} - p|_{m=0} = -\delta P$$

$$\text{i.e. } \frac{dP}{dt} = p|_{m=0} - \delta P - p|_{m=\tau}$$

clearly  $p|_{m=0} = \text{supply rate} = F$

$\hookrightarrow p|_{m=\tau} = \text{loss rate to circulation}$

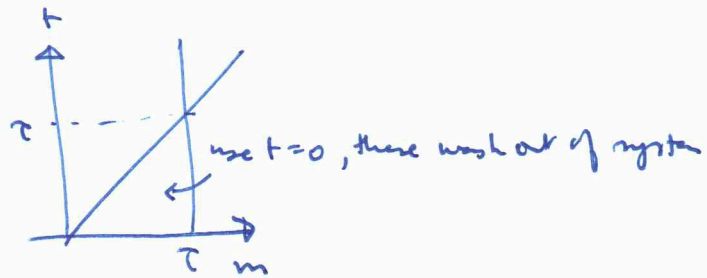
$$(\text{thus } e|_{a=0} = p|_{m=\tau})$$



Characteristics :

$$\dot{p} = -\delta p$$

$$\dot{m} = 1$$



Use boundary conditions in  $m=0$

$$m=0$$

$$t=s$$

$$p = F(s) \quad (\text{specifically } F[E(s)], \quad E(t) = \int_0^A e(t, a) da)$$

$$\Rightarrow p = F(s) e^{-\delta(t-s)}$$

$$m = t-s$$

$$\text{So } p = F(t-m) e^{-\delta m} \quad \forall m \text{ if } t > \tau$$

$$\Rightarrow p|_{m=\tau} = e|_{a=0} = F(t-\tau) e^{-\delta \tau}$$

Next

$$\dot{e} = -\gamma e$$

$$\dot{a} = 1$$

$$e = F(\xi - \tau) e^{-\delta \tau} e^{-\gamma(t-\xi)}$$

$$a = t - \xi$$

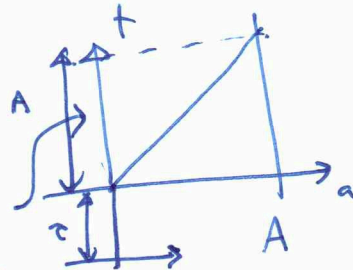
$$\Rightarrow e = F(t-a-\tau) e^{-\delta \tau - \gamma a}$$

and works for all  $a$  if  $t > A + \tau$

Thus with  $E = \int_0^A e da$

$$\dot{E} = -\gamma E + e|_{a=0} - e|_{a=A}$$

$$= -\gamma E + F[E(t-\tau)] e^{-\delta \tau} - F[E(t-A-\tau)] e^{-\delta \tau - \gamma A}$$



If there is no age limit, then  $A \rightarrow \infty$

$$\hookrightarrow \text{just } \dot{E} = -\gamma E + F(E_1)e^{-\delta\tau}$$

Such reduction occurs if  $\gamma A \gg 1$  for example

Non-d  $F = F_0 f$

$$\hookrightarrow \text{scale } E \sim E_0 = \frac{F_0 \tau}{\gamma}, \quad t \sim \tau$$

$$\text{then } \frac{E_0}{\tau} \dot{E} = -\gamma E_0 E + F_0 f(E_1) e^{-\delta\tau} - F_0 f\left[\frac{E}{E_0} E_0 \left(t - 1 - \frac{A}{\tau}\right)\right] e^{-\delta\tau - \gamma A}$$

$$\text{thus } \frac{1}{\gamma\tau} \dot{E} = -E + f(E_1) - e^{-\gamma A} f\left[E \left(t - 1 - \frac{A}{\tau}\right)\right]$$

$$\text{define } \mu = \gamma\tau, \quad \Lambda = \frac{A}{\tau} \Rightarrow \gamma A = \mu\Lambda$$

$$\hookrightarrow \dot{E} = \mu \left[ -E + f(E_1) - e^{-\mu\Lambda} f(E_{1+\Lambda}) \right]$$

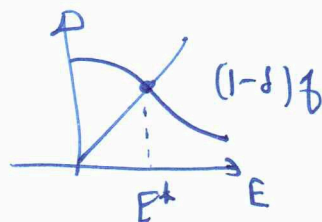
$$A = 120 \text{ d}, \quad \tau = 6 \text{ d}, \quad \Rightarrow \Lambda = 20$$

If  $\mu$  is not small then the finite lifetime is irrelevant - so why would  $A$  be so big. Also we would hope  $E$  would be relatively constant & this is facilitated by  $\mu \ll 1$ .

write  $\delta = e^{-\lambda} < 1$

steady state  $E = (1-\delta) f(E)$

$f$  is decreasing  $\Rightarrow$  unique steady state



Linearize  $E = E^* + \eta$

$$\Rightarrow \dot{\eta} \approx \mu [-\eta + f' \eta_1 - \delta f' \eta_{1+\lambda}]$$

$\eta = e^{\sigma t}$

$$\Rightarrow \sigma = \mu [-1 - |f'| \{e^{-\sigma} - \delta e^{-\sigma(1+\lambda)}\}]$$

$$\Rightarrow \sigma = -\mu [1 + |f'| e^{-\sigma} \{1 - \delta e^{-\sigma\lambda}\}]$$

Suppose that  $\text{Re } \sigma > 0$ :

this requires  $|f'| |e^{-\sigma} \{1 - \delta e^{-\sigma\lambda}\}| > 1$

but LHS  $< |f'| |1 - \delta e^{-\sigma\lambda}|$

since  $\delta < 1$ ,  $|e^{-\sigma\lambda}| < 1$ , certainly  $|1 - \delta e^{-\sigma\lambda}| < 2$

so LHS  $< 2|f'|$

so  $E^*$  is stable if  $|f'| < \frac{1}{2}$

since  $\lambda \gg 1$ ,  $|1 - \delta e^{-\sigma\lambda}| \approx 1$  & this becomes  $|f'| < 1$

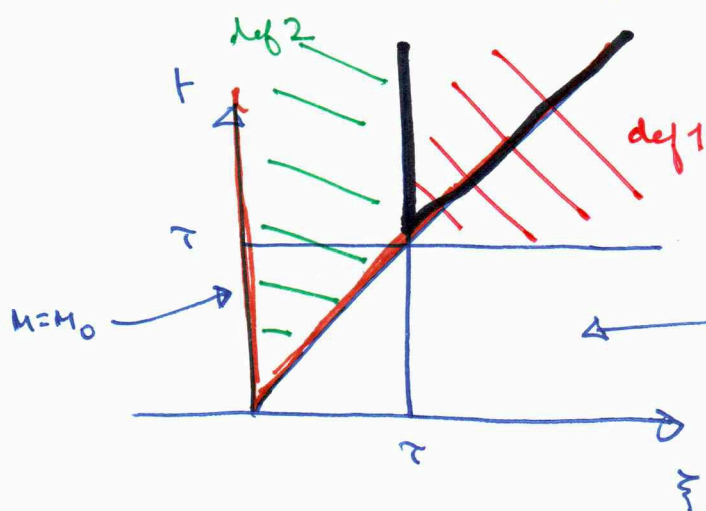
[unless  $\text{Re } \sigma \approx 0 \dots$ ]

6.  $M_t + M_{\xi} = -RM + Q$

$$Q = 2e^{-\gamma\tau} R(t-\tau, \xi-\tau) M(t-\tau, \xi-\tau) \quad \xi > \tau, t > \tau \quad (1)$$

$$= 2e^{-(\gamma_0 + \nu_0)\tau} e^{(\gamma_0 + \nu_0 - \gamma)\xi} \nu_0 R_0(t-\tau) N_0(t-\tau), \quad (2)$$

$$\xi < \tau, t > \xi$$



The definition of  $Q$  will depend on the initial ( $t=0$ ) condition but is washed out of the system.

We focus on the region  $t > \xi$ .

Using  $R = (1+\lambda)R_0$ ,  $\lambda = 2e^{-\gamma\tau} - 1$ ,  $\gamma_0 + \nu_0 = \gamma$ ,  $N_0 \nu_0 = M_0$  we have

$$Q = (1+\lambda)RM_{\tau,\tau} \quad (1)$$

$$RM_0 \quad (2)$$

and  $M = M_0$  on  $\xi = 0, t > 0$ .

In  $t > \xi, \xi < \tau$  (def 2) we have

$$M_t + M_{\xi} = -RM + RM_0, \quad M = M_0 \text{ on } \xi = 0$$

& the solution is obviously  $M \equiv M_0$

Therefore to solve the model in  $t > \xi, t > \tau$  (outlined in black)

we have the characteristic equations

$$\begin{aligned} \dot{t} &= -RM + (1+\lambda)RM_{\tau,\tau} \\ \dot{\xi} &= 1 \end{aligned}$$

$$\begin{cases} M = M_0 \\ t = s > \tau \\ \xi = \tau \end{cases}$$

Thus  $\xi = t - s + \tau$ .

Since the initial condition is independent of  $t$ , it is clear that the solution is a function of  $\xi$  only

Defining  $\eta = \frac{\xi - \tau}{\tau}$ ,  $M = M_0 u(\eta)$

we have  $\eta = \frac{t-s}{\tau}$  and thus  $\dot{M} = \frac{M_0}{\tau} u' = -R M_0 u + (1+\lambda) R u(\eta-1)$ ,

since  $M_{t,\tau} = M(t-\tau, \xi-\tau) = M_0 u\left[\frac{\xi-\tau-\tau}{\tau}\right] = M_0 u(\eta-1)$

i.e.

$$u' = -\alpha u - \Gamma u_1$$

where  $\alpha = R\tau$ ,  $\Gamma = -(1+\lambda)R\tau$

Note that applies for  $\eta \geq 0$  ( $\xi > \tau$ ), and the initial function is  $u = 1$  for  $\eta \in [-1, 0)$

Next, define the Laplace transform

$$U(p) = \int_0^\infty u(\eta) e^{-p\eta} d\eta$$

$$\text{Note } \hat{u}_1 = \int_0^\infty u(\eta-1) e^{-p\eta} d\eta$$

$$[\eta = 1+s] = \int_{-1}^\infty u(s) e^{-p} e^{ps} ds$$

$$= e^{-p} U + e^{-p} \int_{-1}^0 e^{-ps} ds$$

$$= e^{-p} U + \frac{e^{-p}}{p} [e^p - 1] = e^{-p} U + \frac{(1 - e^{-p})}{p}$$

So we have from  $u' = -\alpha u - \Gamma u$ ,  $u \neq 1$ ,  $\gamma < 0$

$$pU - 1 = -\alpha U - \Gamma \left[ e^{-p} U + \frac{(1-e^{-p})}{p} \right]$$

$$\text{or } U = \frac{1 - \Gamma \left( \frac{1-e^{-p}}{p} \right)}{p + \alpha + \Gamma e^{-p}}$$


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$$\text{Hence } U = \frac{p - \Gamma + \Gamma e^{-p}}{p(p + \alpha + \Gamma e^{-p})} = \frac{p + \alpha + \Gamma e^{-p} - (\alpha + \Gamma)}{p(p + \alpha + \Gamma e^{-p})}$$

$$= \frac{1}{p} - \frac{\alpha + \Gamma}{p(p + \alpha + \Gamma e^{-p})}$$

$$= \frac{1}{p} + \frac{(-\alpha - \Gamma)e^p}{p[(p + \alpha)e^p + \Gamma]} = \frac{1}{p} + \frac{\Lambda e^p}{p[(p + \alpha)e^p + \Gamma]}$$

$$\text{where } \Lambda = -\alpha - \Gamma = -R\tau + (1 + \lambda)R\tau = \lambda R\tau$$

$$\text{To invert this, we have } u = \frac{1}{2\pi i} \int_{\gamma-i\infty}^{\gamma+i\infty} U(p) e^{p\eta} dp$$

with all singularities of  $U$  in  $\text{Re } p < \gamma$ .



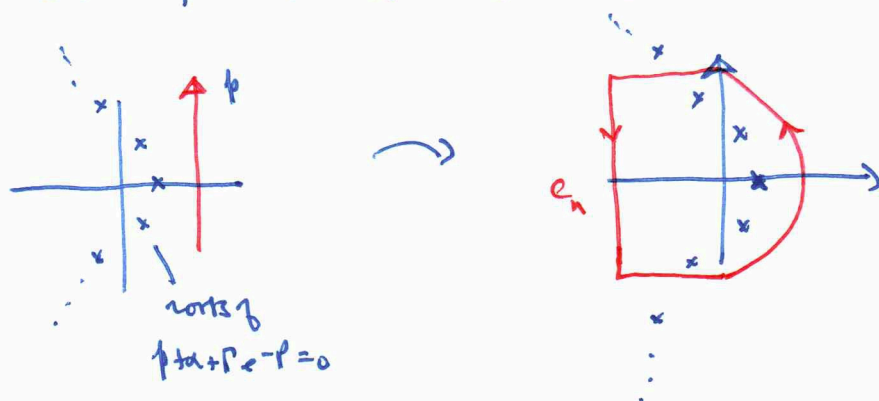
The singularities of  $U$  are at  $p=0$  (removable, so does not contribute)

where  $p = -\alpha - Pe^{-P}$

— let us know all about this! (see question 5)

For  $-P > \alpha$  as here, there is one pair of complex roots, with a finite number to the right of the imaginary axis.

So the inversion contour can be taken as the limit of  $C_n$  as  $n \rightarrow \infty$ , where  $C_n$  is as shown



where the left hand part is the left side of a square of half-length  $(n + \frac{1}{2})\bar{n}$ . The integral round this part vanishes as  $n \rightarrow \infty$  by a version of Jordan's lemma, provided  $p + \alpha + Pe^{-P}$  is bounded away from zero on top + bottom (clearly it goes to  $\infty$  on the left side).

But for  $p = r + i(n + \frac{1}{2})\bar{n}$ ,  $p + \alpha + Pe^{-P} = r + \alpha + i[(n + \frac{1}{2})\bar{n} + |P|e^{-r}]$  if  $\alpha$  is even &  $P < 0$ , so  $|p + \alpha + Pe^{-P}| > n\bar{n} \rightarrow \infty$  in fact.



Therefore

$$u(\gamma) = \lim_{n \rightarrow \infty} \frac{1}{2\pi i} \int_{C_n} U(p) e^{p\gamma} dp$$

$$= \sum_j c_j e^{p_j \gamma},$$

where  $c_j$  are the residues of  $U$  at the roots ~~of  $p + \alpha + \Gamma e^{-p} = 0$~~   
 of  $p + \alpha + \Gamma e^{-p} = 0$

Since  $U = \frac{\Lambda}{p[p + \alpha + \Gamma e^{-p}]} + \frac{1}{p}$

$\hookrightarrow (p + \alpha + \Gamma e^{-p})' = 1 - \Gamma e^{-p}$ , there are

$$\frac{1}{p_j} \cdot \frac{\Lambda}{1 - \Gamma e^{-p_j}}$$
~~$$= \frac{\Lambda}{p_j (1 + p_j + \alpha)}$$~~
~~$$= \frac{\Lambda}{p_j (1 + p_j + \alpha)}$$~~

[Note:  $p$  the roots  $p + \alpha + \Gamma e^{-p} = 0$

satisfy  $p_n = \ln\left(\frac{\Gamma + \alpha}{\Gamma}\right) = \ln p_n - \ln \Gamma + \ln\left(1 + \frac{\alpha}{p_n}\right) + 2\pi i n$

or  $p_n = \frac{2\pi i n \left[ 1 + \frac{\ln p_n - \ln \Gamma + \frac{\alpha}{p_n} \dots}{2\pi i n} \right]}{2\pi i n}$

$\hookrightarrow \ln p_n = \ln 2\pi i n + \frac{\ln p_n}{2\pi i n} \dots$

$= (\ln 2\pi i n) \left( 1 + O\left(\frac{1}{n}\right) \right)$

thus  $p_n \approx 2\pi i n \left[ 1 + \frac{\ln 2\pi i n}{2\pi i n} \dots \right] = 2\pi i n + \ln 2\pi i n + \dots$

$\hookrightarrow p_j \sim O(j) \text{ as } j \rightarrow \infty.$