

3(f) f1) Assuming fast equilibrium for eq<sup>ns</sup> 3-5  
We are left with

$$\frac{dR_i^o}{dt} = \frac{\beta^u}{K^u + N_i^{*u}} - \gamma_R R_i^o \quad (6)$$

∴ For cell 1

$$\frac{dR_1}{dt} = \frac{\beta^u}{K^u + N_1^{*u}} - \gamma_R R_1$$

$$\frac{dR_2}{dt} = \frac{\beta^u}{K^u + N_2^{*u}} - \gamma_R R_2$$

From (3)-(5) at fast eq<sup>m</sup>, we get

$$K_f L R_i^o - K_r R_i^{*u} = 0 \quad (3)$$

$$K_f^{ND} N_i^o D_j^o - K_r^{ND} N_i^{*u} = 0 \quad (4)$$

$$K_D R_i^o - \gamma_D D_i^o = 0 \Rightarrow R_i^o = \frac{\gamma_D D_i^o}{K_D} \Rightarrow D_i^o = \frac{K_D}{\gamma_D} R_i^o \quad (5)$$

$$\therefore N_i^{*u} = \frac{K_f^{ND} \gamma_D}{K_r^{ND} K_D} N_i^o$$

∴ From (4) we get

$$N_i^{*u} = \frac{K_f^{ND} N_i^o D_j^o}{K_r^{ND}} = \frac{K_f^{ND} N_i^o K_D}{K_r^{ND} \gamma_D} R_j^{*u} = A_j R_j^o L$$

$$\text{where } A_j = \frac{K_f^{ND} N_i^o K_D}{K_r^{ND} \gamma_D L}$$

Equation

and

from (6)

$$\therefore \frac{dR_i}{dt} = \frac{\beta^n}{K^n + M_j^n R_j^n L^n} - \gamma_R R_i^n$$

Now, let  $u = \frac{R_1}{K}$ ,  $v = \frac{R_2}{K}$ ,  $\pm \gamma_R = \gamma$

$$\therefore K \gamma \frac{du}{d\tau} = \frac{\beta^n}{K^n + K^n M_j^n u^n L^n} - \gamma_R K u$$

$$\Rightarrow \frac{du}{d\tau} = \frac{\frac{\beta^n}{K^{n+1} \gamma_R}}{1 + M_j^n v^n L^n} - u$$

$$\Rightarrow \boxed{\frac{du}{d\tau} = \frac{\alpha}{1 + A^n u^n} - u} \quad \text{where } \alpha = \frac{\beta^n}{K^{n+1} \gamma_R}$$

$A = M_j^n L$

|| by

$$\boxed{\frac{dv}{d\tau} = \frac{\alpha}{1 + A^n u^n} - v}$$

For stability we have Jacobian of the form

$$J = \begin{bmatrix} -1 & -n A^n (1 + A^n v^n)^{-2} \\ -n A^n (1 + A^n u^n)^{-2} & -1 \end{bmatrix}$$

$$\begin{aligned} \text{Det } |J| &= 1 - n^2 A^{2n} (1 + A^n v^n)^{-2} \cdot n^2 A^{2n} (1 + A^n u^n)^{-2} \\ &= 1 - \frac{n^2 M_j^{2n} L^{2n}}{1 + M_j^{2n} L^{2n} v^{2n}} \cdot \frac{n^2 M_j^{2n} L^{2n}}{1 + M_j^{2n} L^{2n} u^{2n}} \end{aligned}$$

$\therefore$  For stability  $\text{Det } |J| \propto L^{-4n}$  and depends strongly on conc<sup>n</sup> of Ligand for stability.  
The degree of cooperativity,  $n$  can be manipulated for instability.