

where the dimensionless parameter $Cl_{rN} = Cl_r K_r / (k_p A)$ is a measure of the magnitude of the removal rate from the receptor phase (Cl_r) relative to transport through the membrane ($k_p A$), and $V_{rN} = V_r K_r / V_m K_m$ is the dimensionless receptor volume defined as the ratio of the amount of drug in the receptor phase and membrane ($C_r V_r / C_m V_m$), assuming equilibrium exists between phases.

Figure 2.7 shows the effect of receptor volume (as defined by V_{rN}) and clearance of solution from the receptor phase (as defined by Cl_{rN}) on the $J_s(t)$ -time profile.

The steady-state approximation of Equation (2.31) is:

$$Q(t) \approx AJ_{ss} (t - \text{lag}) \tag{2.33}$$

where:

$$J_{ss} = k_p C_v \frac{Cl_r}{Cl_r + (k_p A / K_r)} = \frac{k_p C_v}{1 + (1 / Cl_{rN})} \tag{2.34}$$

and:

$$\text{lag} = \frac{t_d}{6} \left[1 + \frac{2Cl_{rN} - 6V_{rN}}{Cl_{rN} (Cl_{rN} + 1)} \right] \tag{2.35}$$

We note that if Equation (2.29) is substituted into Equation (2.28), the expression for J_{ss} is identical to Equation (2.34). We also note that when $Cl_r \rightarrow \infty$ (infinite sink), J_{ss} and lag reduce to Equations (2.4) and (2.12), respectively.

The corresponding solution for the receptor/epidermal concentration with the noted boundary conditions is:

$$\hat{C}_r(s) = \frac{k_p A C_v}{s} \frac{\sqrt{st_d}}{(V_r s + Cl_r) \left\{ \sinh \sqrt{st_d} + \left[\sqrt{st_d} / (st_d V_{rN} + Cl_{rN}) \right] \cosh \sqrt{st_d} \right\}} \tag{2.36}$$

At long times $t \rightarrow \infty$, C_r is defined by Equation (2.29).

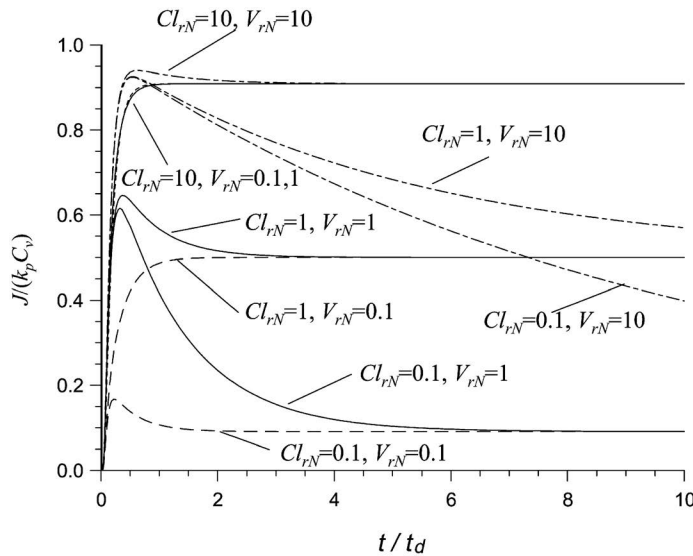


FIGURE 2.7 Normalized flux ($J/k_p C_v$) versus normalized time (t/t_d) for a finite receptor volume and limited clearance [Equation (2.32)], $Cl_{rN} = Cl_r K_r / k_p A$, $V_{rN} = V_r K_r / V_m K_m$.