

interest such as patch removal, sunscreen, and other products and toxins being washed and removed from the skin, when the assumption can be made that there has not been a significant (>10%) depletion in the concentration of solute at the surface. The amount absorbed into a systemic circulation across the skin from the time the dosage form is removed is given by:

$$Q(t) = M_\infty \left\{ 1 - \frac{4}{\pi^3} \sum_{n=0}^{\infty} \frac{(-1)^n}{[n + (1/2)]^3} \exp \left[ -\frac{t}{t_d} \pi^2 \left( n + \frac{1}{2} \right)^2 \right] \right\} \quad (2.23)$$

where  $M_\infty = K_m C_v A h_m / 2 = k_p A C_v t_d / 2$  is the amount of solute present in the skin before removal of the vehicle. The Laplace domain equivalent of this expression is:

$$\hat{Q}(s) = \frac{M_\infty}{s^2} \frac{2}{t_d} \left( 1 - \frac{1}{\cosh \sqrt{st_d}} \right) \quad (2.24)$$

The corresponding equations for flux are:

$$J_s(t) = k_p C_v \frac{2}{\pi} \sum_{n=0}^{\infty} \frac{(-1)^n}{n + (1/2)} \exp \left[ -\frac{t}{t_d} \pi^2 \left( n + \frac{1}{2} \right)^2 \right] \quad (2.25)$$

$$\hat{J}_s(s) = \frac{k_p C_v}{s} \left( 1 - \frac{1}{\cosh \sqrt{st_d}} \right) \quad (2.26)$$

Figure 2.6 shows the amount and flux–time profiles associated with donor phase removal.

The mean time for absorption of solute from the skin in this case is given by:

$$\text{MAT} = \frac{\int_0^{\infty} J_s(t) dt}{\int_0^{\infty} J_s(t) dt} = -\lim_{s \rightarrow 0} \frac{d}{ds} \ln(\hat{J}_s) = \frac{5h_m^2}{12D_m} = \frac{5}{12} t_d \quad (2.27)$$

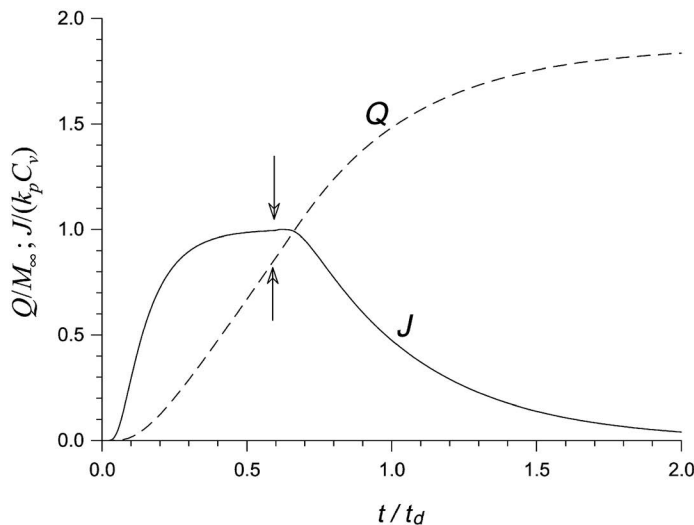


FIGURE 2.6 Changes in normalized cumulative amounts penetrating ( $Q/M_\infty$ ) and flux ( $J/k_p C_v$ ) when the vehicle is removed (as indicated by arrows) at a specific normalized time ( $t/t_d$ ) after application.