



Kinetic Modeling

A precise function was used to estimate the level of pyruvate over time. The function is of form:

$$f(t) = \begin{cases} \frac{r_p}{k_p} (1 - e^{-k_p(t - t_s)}) & t_s \le t < t_e \\ f(t_e)e^{-k_p(t - t_e)} & t \ge t_e \end{cases}$$

where r_p , k_p , t_s , and t_e are estimated by minimizing the error sum of squares.

After estimation of these parameters, we estimate the alanine-to-pyruvate and lactate-to-pyruvate ratios over time. The fitted functions are of the form:

$$g(t) = \begin{cases} 0 & t < t_{s} \\ \frac{k_{px}r_{p}}{k_{p} - k_{x}} \left(\frac{1 - e^{-k_{x}(t - t_{s})}}{k_{x}} - \frac{1 - e^{-k_{p}(t - t_{s})}}{k_{p}}\right) & t_{s} \le t < t_{e} \\ \frac{f(t_{e})k_{px}}{k_{p} - k_{x}} \left(e^{-k_{x}(t - t_{e})} - e^{-k_{p}(t - t_{e})}\right) + g(t_{e})e^{-k_{x}(t - t_{e})} & t \ge t_{e} \end{cases}$$

where x represent alanine and lactate, and k_{px} , k_x are estimated by minimizing the error sum of squares. The fitted plots are shown in the article Figure 3 C and the parameters are listed in Table 1



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