

Supplemental Digital Content 1

Table 1. Gradients Applied in the Determination of Fentanyl in Human Plasma Samples by Liquid Chromatography–Mass Spectrometry

Time [min]	Solvent A [%]	Solvent B [%]
0.1	95	5
1.00	95	5
3.50	5	95
4.00	5	95
4.10	95	5
5.50	95	5

Starting conditions were 95% A and 5% B.

Visual Predictive Checks of the Fentanyl Pharmacokinetic and Pharmacodynamic Model Outcomes

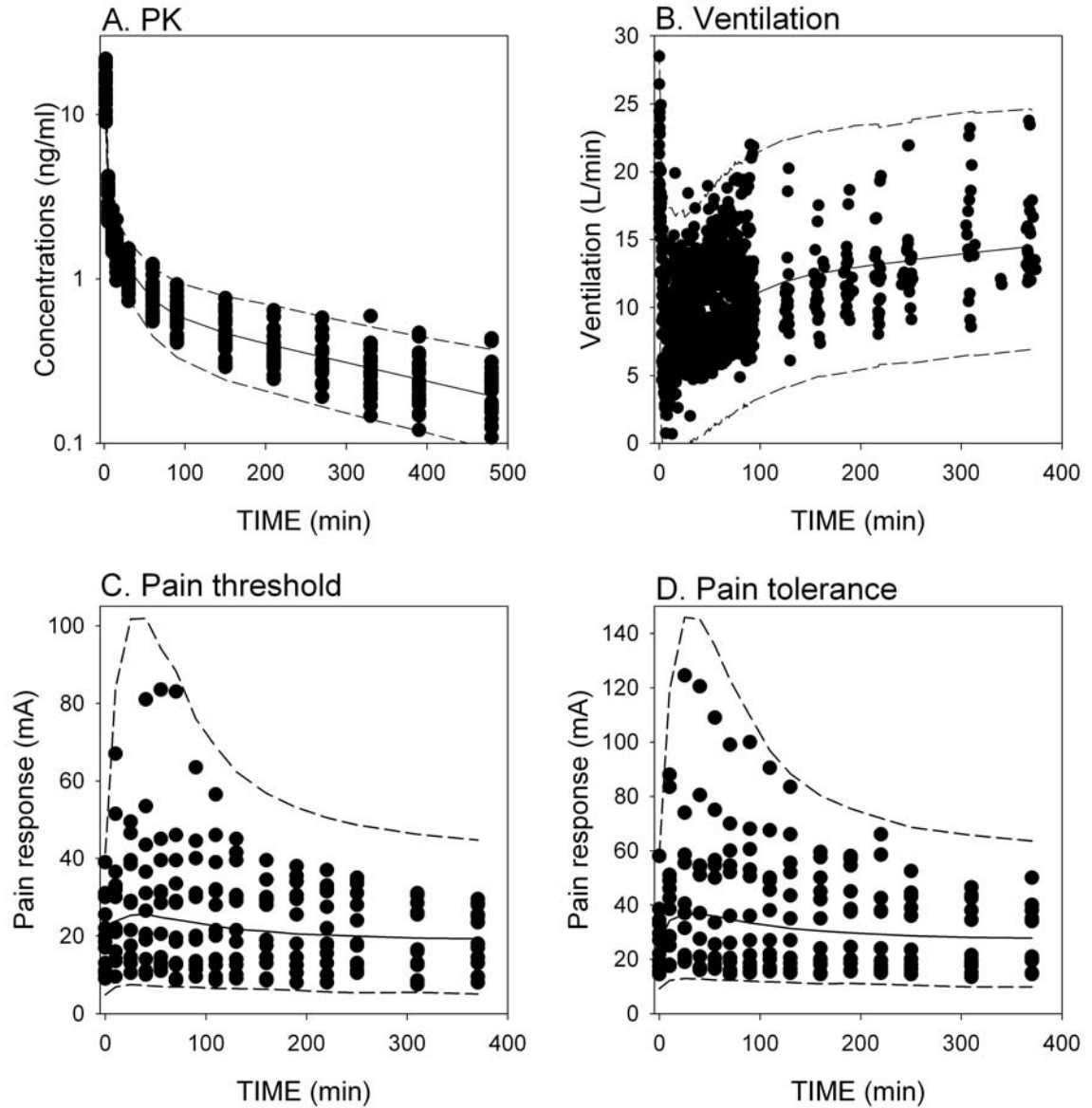


Fig. 1. Simulations performed with input function similar to the current study. (A) Fentanyl concentration, (B) Ventilation, (C) Pupil diameter, (D) Pain tolerance, (E) Pain threshold. Continuous lines: the median data fit; broken lines $\pm 95\%$ confidence intervals.