

*Drug Dosage Calculations:*

For fentanyl and midazolam, the dose calculations were as follows:<sup>17-19</sup>

Bolus dose over one minute (mcg/kg) = desired plasma level (ng/ml) x Vd (L/kg),

where Vd = volume of distribution

Infusion rate (mcg·kg<sup>-1</sup>·hr<sup>-1</sup>) = desired plasma level (ng/ml) x CL (L·kg<sup>-1</sup>·min<sup>-1</sup>) x 60,

where CL = drug clearance

Fentanyl:

<u>Age</u>	<u>Vd (L/kg)</u>	<u>CL (L·kg<sup>-1</sup>·min<sup>-1</sup>)</u>	<u>Bolus</u>	<u>Infusion</u>
1 month-1 year	4.45	0.018	18 mcg/kg	4.3 mcg·kg <sup>-1</sup> ·hr <sup>-1</sup>
1-5 years	3.06	0.012	12 mcg/kg	2.8 mcg·kg <sup>-1</sup> ·hr <sup>-1</sup>
6-9 years	2.54	0.010	10 mcg/kg	2.3 mcg·kg <sup>-1</sup> ·hr <sup>-1</sup>
10-13 years	1.92	0.007	8 mcg/kg	1.7 mcg·kg <sup>-1</sup> ·hr <sup>-1</sup>

The second bolus was 50% of the first dose, and the maintenance infusion rates then increased by 50% for the second measurement.

Midazolam:

Because of the reported variability in pharmacokinetic data of midazolam in children, the same initial and maintenance infusion was chosen for all ages.<sup>20-22</sup> Calculations were based on a CL of 0.009 L·kg<sup>-1</sup>·min<sup>-1</sup>, and a Vd of 1.9 L/kg. A dose of 0.29 mg/kg was followed by an infusion of 139 mcg·kg<sup>-1</sup>·hr<sup>-1</sup>. The second dose was 0.15 mg/kg followed by an infusion of 208 mcg·kg<sup>-1</sup>·hr<sup>-1</sup>.

*Echocardiographic Calculations:*

$$\%SF = \frac{LVEDD - LVESD}{LVEDD} \times 100$$

SF = shortening fraction; LVEDD = left ventricular end-diastolic dimension; LVESD= left ventricular end-systolic dimension

$$\%EF = \frac{LVEDV - LVESV}{LVEDV} \times 100$$

EF = ejection fraction; LVEDV = left ventricular end-diastolic volume; LVESV = left ventricular end-systolic volume

$$\text{CO} = D_a^2 \times \text{VTI}_a \times \text{HR}$$

CO = systemic cardiac output;  $D_a$  = aortic diameter;  $\text{VTI}_a$  = aortic velocity time integral.

$$\text{SV} = \frac{\text{CO}}{\text{HR}}$$

SV = stroke volume, HR = heart rate

$$\text{SVR} = \frac{\text{MAP} - \text{CVP}}{\text{CO}}$$

SVR = systemic vascular resistance, MAP = mean arterial pressure, CVP = central venous pressure.

SV, LVEDV, and CO were divided by the patient's body surface area to calculate an index to account for the different patient sizes. The SVR was multiplied by the body surface area to calculate an SVR index.