

<b>Alert</b>	High-alert medication: High risk of causing significant patient harm when used in error. <sup>1</sup> This drug should be administered in the presence of personnel trained in advanced airway management. Reversal agents should be immediately available (see Special Comments). Suggest regular cessation of infusion for a few to several hours, possibly every 24 hours (commonly referred to as a 'drug holiday') to assess the need for continued paralysis and adequacy of sedation or analgesia. Following cessation, the line should be adequately flushed to avoid unintended paralysis during later use of the line. Eye lubricant should be used whilst patient is receiving vecuronium.												
<b>Indication</b>	1. Skeletal muscle relaxation or paralysis in mechanically ventilated infants. 2. For elective endotracheal intubation.												
<b>Action</b>	Nondepolarizing neuromuscular blockers are competitive acetylcholine (ACh) antagonists that bind directly to nicotinic receptors on the postsynaptic membrane, thus blocking the binding of ACh so the motor endplate cannot depolarize. Onset of action is 1-2 minutes; duration of action is 30-40 minutes.												
<b>Drug Type</b>	Non-depolarising neuromuscular blocking agent.												
<b>Trade Name</b>	Vecure Powder for injection, Vecure Sun Powder for injection												
<b>Presentation</b>	10 mg of vecuronium bromide in glass vial (powder for reconstitution)												
<b>Dose</b>	<p><b>Intubation</b> IV bolus – 100 microgram/kg</p> <p><b>Muscle relaxation**</b></p> <p><b>Intermittent IV bolus</b> 100 microgram/kg (30-150 microgram/kg) IV push every 1-2 hours as required.</p> <p><b>Continuous IV infusion (with or without loading dose)</b> 100 microgram/kg/hour (60-200 microgram/kg/hour). Start 20 minutes post bolus recovery. Titrate in 10% dose increments until desired neuromuscular blockade is achieved.</p> <p>* Provide eye protection and instil lubricating eye drops every 2 hours. # Sensation remains intact; additional sedation &amp; analgesia should be used for painful procedures.</p>												
<b>Dose adjustment</b>	Therapeutic hypothermia – No information. ECMO – No information. Renal impairment – No specific dose adjustment. However, duration of action may be prolonged. Hepatic impairment – No specific dose adjustment. However, hepatic impairment decreases clearance resulting in prolonged duration of action.												
<b>Route</b>	IV												
<b>Maximum Dose</b>	IV bolus: 200 microgram/kg IV infusion: 200 microgram/kg/hour.												
<b>Preparation</b>	<p><b>IV infusion:</b> Note: Refer to <a href="#">Appendix</a> for tables to assist with concentration selection.</p> <p><b>Weight suggestions for infusion concentrations below are a guide only. Clinicians may choose infusion concentration different to the suggested based on expected dose and the corresponding 24 hour fluid volumes.</b></p> <table border="1"> <thead> <tr> <th>Infant weight</th> <th>≤1 kg</th> <th>&gt;1 kg</th> </tr> </thead> <tbody> <tr> <td><b>Suggested vecuronium concentration</b></td> <td><b>400 microgram/mL</b></td> <td><b>1000 microgram/mL</b></td> </tr> <tr> <td><b>100 microgram/kg/hour is equal to</b></td> <td>0.25 mL/kg/hour</td> <td>0.1 mL/kg/hour</td> </tr> <tr> <td><b>IV bolus of 100 microgram/kg is equal to</b></td> <td>0.25 mL/kg</td> <td>0.1 mL/kg</td> </tr> </tbody> </table>	Infant weight	≤1 kg	>1 kg	<b>Suggested vecuronium concentration</b>	<b>400 microgram/mL</b>	<b>1000 microgram/mL</b>	<b>100 microgram/kg/hour is equal to</b>	0.25 mL/kg/hour	0.1 mL/kg/hour	<b>IV bolus of 100 microgram/kg is equal to</b>	0.25 mL/kg	0.1 mL/kg
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## **20 mL Syringe**

**Step 1:** Add 5 mL water for injection to 10 mg vecuronium powder for reconstitution vial to make a **2 mg/mL solution**.

**Step 2:** Draw up reconstituted vecuronium and add compatible fluid\* as per table below to make a final volume of 20 mL:

Vecuronium concentration	400 microgram/mL	1000 microgram/mL
Volume of vecuronium (2 mg/mL solution)	4 mL (=8 mg)	10 mL (=20 mg)
Volume of compatible fluid*	16 mL	10 mL
Total volume	20 mL	20 mL

\*Compatible fluid: glucose 5%, or sodium chloride 0.9%

## **50 mL Syringe**

**Step 1:** Add 5 mL water for injection to 10 mg vecuronium powder for reconstitution vial to make a **2 mg/mL solution**.

**Step 2:** Draw up reconstituted vecuronium and add compatible fluid\* as per table below to make a final volume of 50 mL:

Vecuronium concentration	400 microgram/mL	1000 microgram/mL
Volume of vecuronium (2 mg/mL solution)	10 mL (=20 mg)	25 mL (=50 mg)
Volume of compatible fluid*	40 mL	25 mL
Total volume	50 mL	50 mL

\*Compatible fluid: glucose 5%, or sodium chloride 0.9%

### **IV bolus:**

Add 5 mL water for injection to 10 mg of vecuronium powder for reconstitution vial to make a 2 mg/mL solution.

**Further dilute:** From this vial, draw up 2 mL (4 mg) and add to 2 mL of sodium chloride 0.9% to make a final volume of 4 mL with a concentration of 1 mg/mL [1000 microgram/mL].

<b>Administration</b>	IV bolus: Administer over several seconds. IV infusion via syringe pump. Flush line adequately after each dose with sodium chloride 0.9% to avoid unintended paralysis and/or incompatibility with other medications during later use of the line.
<b>Monitoring</b>	Continuous cardio-respiratory and pulse oximetry monitoring. Close monitoring of neuromuscular function, sedation, and blood pressure (invasive or non-invasive). Monitor electrolytes and renal function. Monitor injection site for signs of extravasation.
<b>Contraindications</b>	Hypersensitivity to vecuronium or any component of the formulation. Cross-sensitivity with other neuromuscular-blocking agents may occur; use with extreme caution in patients with previous anaphylactic reactions. Severe electrolyte abnormalities.
<b>Precautions</b>	Avoid prolonged usage. <b>Factors which can increase duration of neuromuscular blockade:</b> Acidosis, hypothermia, neuromuscular disease, hepatic disease, hypokalaemia, hypermagnesaemia, renal failure, and younger age. Vecuronium is lipid soluble and is predominantly excreted via the liver so poor liver function can cause prolonged effects. <b>Factors which can decrease duration of neuromuscular blockade:</b> Alkalosis and hyperkalaemia.  Use cautiously in neonates with hepatic or renal impairment and in neonates with fluid and electrolyte imbalance.

	<p>Suggest regular cessation of infusion, possibly every 24 hours (commonly referred to as 'drug holiday') to assess the need for continued paralysis and adequacy of sedation or analgesia. Monitoring of fluid balance is essential due to risk of fluid retention.</p> <p>Aminoglycosides &amp; general anaesthetics can increase (potentiate) duration of neuromuscular blockade.</p> <p><b>Corticosteroids:</b> Concomitant use with corticosteroids has been reported to be associated with development of acute quadriplegic myopathy syndrome (AQMS) in adults.<sup>3</sup> However, Recent trials provided no evidence for increased risk of neuromyopathy in patients with sepsis or acute respiratory distress syndrome (ARDS) with the use of corticosteroids or neuromuscular blockers.<sup>17</sup></p> <p>Adrenaline (epinephrine) can reduce (antagonise) duration of neuromuscular blockade.</p>
<b>Drug Interactions</b>	<p>Antimicrobials like aminoglycosides, tetracyclines, polymyxins, and clindamycin can potentiate neuromuscular blockade.<sup>3</sup></p> <p>Inhaled anaesthetics can potentiate neuromuscular blockade.<sup>3</sup></p> <p>Anti-epileptics can make patients resistant to vecuronium.<sup>3</sup></p> <p>Local anaesthetics can potentiate neuromuscular blockade.<sup>3</sup></p> <p>Aminoglycosides &amp; general anaesthetics can increase duration of neuromuscular blockade.</p> <p><b>Corticosteroids:</b> Concomitant use with corticosteroids has been reported to be associated with development of acute quadriplegic myopathy syndrome (AQMS) in adults.<sup>3</sup> However, Recent trials provided no evidence for increased risk of neuromyopathy in patients with sepsis or acute respiratory distress syndrome (ARDS) with the use of corticosteroids or neuromuscular blockers.<sup>17</sup></p> <p>Dexamethasone and hydrocortisone may result in decreased vecuronium effectiveness, prolonged muscle weakness, and myopathy.<sup>3</sup></p> <p>Adrenaline (epinephrine) can reduce (antagonise) duration of neuromuscular blockade.</p>
<b>Adverse Reactions</b>	<p>Hypoxaemia may occur because of inadequate ventilation and deterioration in pulmonary mechanics.</p> <p>Hypotension and bradycardia, particularly when used in combination with opioids.</p> <p>Prolonged paralysis after long-term use.</p> <p>Rare: Anaphylactic reaction and tachycardia.</p>
<b>Overdose</b>	<p>Supportive measure: Ventilatory support and sedation.</p> <p>Reversal of neuromuscular blockade can be achieved by neostigmine (refer to special comments).</p> <p>AUSTRALIA</p> <p>Contact the Poisons Information Centre on <b>13 11 26</b> for information on the management of overdose</p> <p>NEW ZEALAND</p> <p>Contact the National Poisons Centre on <b>0800 764 766</b> for information on the management of overdose</p>
<b>Compatibility</b>	<p>Fluids:<sup>3</sup> glucose 5%, sodium chloride 0.9%.</p> <p><b>PN at Y site:</b> Amino acid solution. No information on lipid emulsions.</p> <p>Y-site:<sup>3</sup> glucose/amino acid solutions, adrenaline (epinephrine), alprostadil, amikacin sulfate, aminophylline, amiodarone, ampicillin, atenolol, azithromycin, aztreonam, caffeine citrate, calcium chloride, calcium gluconate, cefazolin, ceftazidime, ceftriaxone, ciprofloxacin, clindamycin, dexamethasone sodium phosphate, dexmedetomidine, digoxin, diltiazem, dobutamine, dopamine, enalaprilat, adrenaline (epinephrine), erythromycin lactobionate, esmolol, fentanyl, fluconazole, fluorouracil, fosphenytoin, gentamicin, glycopyrrolate, heparin, hydralazine, hydrocortisone sodium succinate, insulin (regular), isoprenaline, labetalol, lidocaine, linezolid, lorazepam, magnesium sulfate, meropenem, metoprolol, metronidazole, midazolam, milrinone, morphine, naloxone, nicardipine, nitroglycerin, norepinephrine, octreotide, ondansetron, pamidronate, pentoxifylline, phenobarbital, phenylephrine, potassium acetate, potassium chloride, propofol at vecuronium concentrations of <math>\leq 1</math> mg/mL, propranolol hydrochloride, ranitidine hydrochloride, remifentanil, sodium acetate, sodium bicarbonate, sodium nitroprusside, sodium phosphate, streptozocin, succinylcholine, tacrolimus, theophylline, ticarcillin disodium/clavulanate potassium, tigecycline, tobramycin,</p>

	sulfamethoxazole/trimethoprim, and vancomycin hydrochloride, vasopressin, verapamil, voriconazole, zidovudine, and zoledronic acid.
<b>Incompatibility</b>	Fluids: No information. No information on lipid emulsions. Y site: <sup>3</sup> Aciclovir, amphotericin B (all compounds), cefepime, cefotaxime, diazepam, furosemide, ganciclovir, ibuprofen lysine, imipenem/cilastatin sodium, methylprednisolone sodium succinate, micafungin sodium, pantoprazole, phenytoin, piperacillin sodium, piperacillin-tazobactam, propofol at vecuronium concentrations >1 mg/mL, sulbactam/durlobactam, and thiopental sodium.
<b>Stability</b>	Diluted solution stable for up to 24 hours. Discard any unused solution.
<b>Storage</b>	Store below 25°C. Protect from light. Store in accordance with local policies.
<b>Excipients</b>	Citric acid, dibasic sodium phosphate, sodium hydroxide and/or phosphoric acid, and mannitol.
<b>Special Comments</b>	Muscle relaxation is reversed by neostigmine (50 microgram/kg) and atropine (20 microgram/kg). Sugammadex is being increasingly used with extrapolated information from other populations. Sensation remains intact: sedation & analgesia should be used for painful procedures. Provide eye protection and instil lubricating eye drops every 2 hours. Vecuronium produces less tachycardia and hypotension when compared with pancuronium. <sup>15,16</sup> The neuromuscular blockade of vecuronium is of shorter duration than that of pancuronium. <sup>15,16</sup> Prior administration of suxamethonium shortens onset and may increase depth of blockade; reduce dose and give vecuronium only after recovery from suxamethonium-induced neuromuscular blockade.
<b>Evidence</b>	<b>Background</b> Nondepolarizing neuromuscular blocking agents (NMBA) can be classified into 2 classes: steroid (rocuronium, vecuronium, pancuronium) or benzylisoquinoline (mivacurium, atracurium, cisatracurium). Nondepolarizing neuromuscular blockers are competitive acetylcholine (ACh) antagonists that bind directly to nicotinic receptors on the postsynaptic membrane, thus blocking the binding of ACh so the motor endplate cannot depolarize. This leads to skeletal muscle paralysis. <sup>2</sup> Paralysis occurs sequentially because of the differing sensitivity of muscles to NMAs as well as blood flow to the area. Generally, paralysis begins with smaller, fast twitch muscles such as the eyes and larynx, then affects the limbs, neck, trunk, and upper airway, and eventually progresses to the intercostals and diaphragm until respiration terminates. Recovery from paralysis occurs in the reverse order with function of the diaphragm returning first. <sup>4</sup> <b>Vecuronium:</b> vecuronium is structurally related to pancuronium. It has a greater potency, shorter duration of action, lack significant cardiovascular effects (tachycardia), and less cumulative properties. Vecuronium produces less tachycardia and hypotension when compared with pancuronium. <sup>15,16</sup> The neuromuscular blockade of vecuronium is of shorter duration than that of pancuronium. <sup>15,16</sup> Time of onset of action is 90-120 seconds after IV bolus, with a duration of effect that lasts only 30-40 minutes. Intermittent bolus dosing would need to be so frequent (i.e., every 30 to 60 minutes) that continuous IV infusion is preferred over intermittent boluses to maintain paralysis in ventilated infants. <sup>5</sup> In comparison, rocuronium is an analogue of vecuronium with a more rapid onset of action (20-100 seconds), but less potent than vecuronium, and hence larger doses (example, 600 microgram/kg of rocuronium, compared to 100 microgram/kg of vecuronium). Tachycardia is more frequent with rocuronium, while vecuronium lacks this effect in regular doses. <sup>4</sup> The neuromuscular blockade effect of vecuronium is stronger and lasts longer in neonate than infant or adult. <sup>6</sup> The 95% effective dose (ED <sub>95</sub> ) for NMAs specifies the dose that produces 95% twitch depression in 50% of individuals. ED <sub>95</sub> of vecuronium in neonates (47 microgram/kg) is 40% less than in children (81 microgram/kg) meaning less dose is needed in neonates compared to children. <sup>7</sup> <b>Efficacy</b> <u>Muscle relaxation</u> Two prospective studies by Meretoja et al in 1988 and 1989 determined the dose responses with vecuronium bolus and continuous infusion in paediatric population. <sup>7,8</sup> The bolus dose required to achieve effective neuromuscular blockade in neonates was 40% less than in children. The median maintenance dose of 0.1 mg/kg is required in neonates to maintain 1 hour of neuromuscular blockade, in comparison to 0.217 mg/kg/hour in children 3-10 years old.

	<p>Fitzpatrick et al studied vecuronium to facilitate paralysis in mechanically ventilated paediatric population (4 neonates, and 11 infants and children). A loading dose of 0.1 mg/kg was followed by an infusion of 0.1 mg/kg/hour. The titration rate was adjusted to maintain a neuromuscular block of approximately 90% as assessed by the presence of one response to a train of four (TOF) stimulation. The duration of the infusions varied from 9.5 to 179 hours. Mean recovery times after stopping the infusion were 51.7 (<math>\pm 17.6</math>) and 46.8 (<math>\pm 16.5</math>) minutes for the children and neonates respectively. No adverse cardiovascular or toxic effects were noted.<sup>9</sup> Fisher et al determined the recovery period (time from injection to 90% recovery) after a bolus of vecuronium. Recovery was longest for infants (73<math>\pm</math>27 minutes), compared to children (35<math>\pm</math>6 minutes).<sup>10</sup> The longer recovery period in neonates is thought to be due higher volume of distribution.</p> <p>Hodges et al evaluated the appropriate vecuronium infusion rates in 12 neonates/infants and 18 children using train of four (TOF) monitoring. Neonates and infants required 45% less vecuronium (mean infusion rate 0.54 mg/kg/hour) than older children (0.99 mg/kg/hour) and had faster recovery (45 min vs 65 min), with no evidence of prolonged weakness.<sup>11</sup></p> <p><b>Safety</b></p> <p>Adults with hepatic and renal failure have been shown to experience prolonged neuromuscular blockade.<sup>12,13</sup></p> <p><b>Pharmacokinetics</b></p> <p>Hepatobiliary clearance is the primary route of elimination, accounting for approximately 50% of the dose. Vecuronium is metabolised rapidly in the liver to 3-desacetyl-vecuronium, which is 50–70% as potent as the parent compound. This metabolite is cleared primarily by renal elimination. Approximately 20–30% of vecuronium is excreted unchanged in urine.<sup>9,11,12</sup></p> <p>Time of onset of action is 90-120 seconds after IV bolus, with a duration of effect that lasts only 30-40 minutes. (prolonged with higher doses and in preterm infants).<sup>5,14</sup></p>
<b>Practice points</b>	Eye lubrication should be applied to all patients.
<b>References</b>	<ol style="list-style-type: none"> <li>1. Clinical Excellence Commission. Neuromuscular blocking agents. <a href="https://www.cec.health.nsw.gov.au/keep-patients-safe/medication-safety/high-risk-medicines/neuromuscular-blocking-agents">https://www.cec.health.nsw.gov.au/keep-patients-safe/medication-safety/high-risk-medicines/neuromuscular-blocking-agents</a>. Accessed on 7 October 2024.</li> <li>2. Cook D, Simons DJ. Neuromuscular Blockade. [Updated 2023 Nov 13]. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2024 Jan-. Available from: <a href="https://www.ncbi.nlm.nih.gov/books/NBK538301/">https://www.ncbi.nlm.nih.gov/books/NBK538301/</a>.</li> <li>3. MerativeTM Micromedex® Complete IV Compatibility (electronic version). Merative, Ann Arbor, Michigan, USA. Available at: <a href="https://www.micromedexsolutions.com/">https://www.micromedexsolutions.com/</a> (cited: Oct/8/2024).</li> <li>4. Warr J, Thiboutot Z, Rose L, Mehta S, Burry LD. Current Therapeutic Uses, Pharmacology, and Clinical Considerations of Neuromuscular Blocking Agents for Critically Ill Adults. Annals of Pharmacotherapy. 2011;45(9):1116-26.</li> <li>5. Kandasamy J, Carlo WA. Pharmacologic therapies IV. In Assisted ventilation of the neonate. 2017. Elsevier Inc.</li> <li>6. Meakin GH. Neuromuscular blocking drugs in infants and children. Continuing Education in Anaesthesia Critical Care &amp; Pain. 2007;7(5):143-7.</li> <li>7. Meretoja OA, Wirtanen K, Neuvonen PJ. Age-dependence of the dose-response curve of vecuronium in pediatric patients during balanced anesthesia. Anesthesia &amp; Analgesia. 1988;67(1):21-6.</li> <li>8. Meretoja O. Is vecuronium a long-acting neuromuscular blocking agent in neonates and infants? British Journal of Anaesthesia. 1989;62(2):184-7.</li> <li>9. Fitzpatrick KT, Black GW, Crean PM, Mirakhur RK. Continuous vecuronium infusion for prolonged muscle relaxation in children. Can J Anaesth. 1991;38:169-74.</li> <li>10. Fisher DM, Miller RD. Neuromuscular effects of vecuronium (ORG NC45) in infants and children during N<sub>2</sub>O, halothane anesthesia. Anesthesiology. 1983;58(6):519-23.</li> <li>11. Hodges U. Vecuronium infusion requirements in paediatric patients in intensive care units: the use of accelerometry. British journal of anaesthesia. 1996;76(1):23-8.</li> </ol>

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15. Basta SJ, Savarese JJ, Ali HH et al. Vecuronium does not alter serum histamine within the clinical dose range. *Anesthesiology* 1983;59:A273.
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## Appendix

### Infusion tables to assist concentration selection

**Table 1:** Infusion rates when using vecuronium concentration **400 microgram/mL** (suggested weight  $\leq 1\text{kg}$ )

Rate (mL/hr)	0.1	0.2	0.3	0.4	0.5	0.6	0.7	0.8	0.9	1
Weight (kg)	Approximate microgram/kg/hour									
0.5	80	160	240	320	400	480	560	640	720	800
1	40	80	120	160	200	240	280	320	360	400
1.5	27	53	80	107	133	160	187	213	240	267
2	20	40	60	80	100	120	140	160	180	200
2.5	16	32	48	64	80	96	112	128	144	160
3	13	27	40	53	67	80	93	107	120	133
3.5	11	23	34	46	57	69	80	91	103	114
4	10	20	30	40	50	60	70	80	90	100
4.5	9	18	27	36	44	53	62	71	80	89
5	8	16	24	32	40	48	56	64	72	80

**Table 2:** Infusion rates when using vecuronium concentration 1000 microgram/mL (suggested weight >1 kg)

Rate (mL/hr)	0.1	0.2	0.3	0.4	0.5	0.6	0.7	0.8	0.9	1
Weight (kg)	Approximate microgram/kg/hour									
0.5	200	400	600	800	1000	1200	1400	1600	1800	2000
1	100	200	300	400	500	600	700	800	900	1000
1.5	67	133	200	267	333	400	467	533	600	667
2	50	100	150	200	250	300	350	400	450	500
2.5	40	80	120	160	200	240	280	320	360	400
3	33	67	100	133	167	200	233	267	300	333
3.5	29	57	86	114	143	171	200	229	257	286
4	25	50	75	100	125	150	175	200	225	250
4.5	22	44	67	89	111	133	156	178	200	222
5	20	40	60	80	100	120	140	160	180	200

$$\text{Rate (mL/hr)} = \frac{\text{Dose (microgram/kg/hour)} \times \text{Weight (kg)}}{\text{Concentration (microgram/mL)}}$$

$$\text{Dose (microgram/kg/hour)} = \frac{\text{Rate (mL/hr)} \times \text{Concentration (microgram/mL)}}{\text{Weight (kg)}}$$

VERSION/NUMBER	DATE
Original 1.0	17/07/2025
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This standard concentration formulary has been developed by the ANMF standard concentration working group.

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