

**NORADRENALINE** **This drug must be guardrailed**

<b>Trade Name</b>	Levophed <sup>®</sup> 1:1000 Concentrate for IV Injection (Hospira) Noradrenalin- BNM <sup>®</sup> 4mg /4 mL (=1:1000)
<b>Class</b>	Vasoconstrictor sympathomimetic
<b>Mechanism of Action</b>	Increases blood pressure by stimulating alpha receptors in vascular smooth muscle resulting in peripheral vasoconstriction. Also stimulates cardiac beta receptors providing some inotropic effects including increased heart rate and vasodilatation of coronary arteries.
<b>Indications</b>	Refractory hypotension in the setting of pulmonary hypertension or septic shock with no response to fluid resuscitation, or high dose dopamine
<b>Contraindications</b>	Uncorrected hypovolaemia (absolute contraindication). History of hypersensitivity to noradrenaline or sulphites. Hypertension. Mesenteric or peripheral vascular thrombosis. Hypoxia or hypercapnia (may cause noradrenaline-induced cardiac arrhythmias) No central line access
<b>Supplied As</b>	4mg/4mL
<b>Dilution</b>	<b>See Noradrenaline infusion sheet:</b> <b>Babies greater than 2.5 kg</b> Take 1x wt (kg) in mL of noradrenaline 1mg/mL and make up to 50mL with 5% dextrose without heparin <b>0.1 microgram/kg/min = 0.3 mL/hr</b> <b>Max concentration = 100 microgram/mL</b>  <b>Babies less than or equal to 2.5kg</b> Take 2x wt (kg) in mL of noradrenaline 1mg/mL and make up to 50mL with 5% dextrose without heparin <b>0.2 microgram/kg/min = 0.3 mL/hr</b> <b>Max concentration = 100 microgram/mL</b>  Must be infused through a central line
<b>Dosage</b>  <b>*Must chart guardrail and use Alaris pump*</b>	0.05 – 0.5 microgram/kg/minute Suggest starting at 0.1 microgram/kg/minute and titrate up or down every 30 minutes until control achieved Higher doses up to 1-2 microgram/kg/minute may be required to control blood pressure and are at the Consultant's discretion

<b>Guardrail</b>	<p>Conc: Min – 4 microgram/mL      Max – 100 microgram/mL</p> <p>Soft Min: 0.05 microgram/kg/min      Hard Max: 2 microgram/kg/min</p> <p>Soft Max: 0.5 microgram/kg/min      Default: 0.1 microgram/kg/min</p>
<b>Interval</b>	Continuous iv infusion
<b>Administration</b>	<p>Continuous IV infusion via a central venous line.</p> <p><b>Not</b> to be given subcutaneously or intramuscularly due to risk of severe, rapid vasoconstriction, this may result in gangrene.</p> <p>Avoid extravasation as this will cause tissue necrosis.</p> <p>Antidote for noradrenaline extravasation is phentolamine. (Infiltrate affected area with 1- 5 mg diluted in 5 mL sodium chloride 0.9%)<sup>6</sup></p>
<b>Compatible With</b>	<p><b>Solutions:</b> Glucose 5% or glucose/ saline is preferred because glucose protects noradrenaline from oxidative degradation. Infusion in solutions other than glucose is not recommended by the manufacturer however independent reference sources eg Neofax, ANMF, Micromedex site compatibility with sodium chloride 0.9% and lactated Ringer's.</p> <p><b>Terminal Y-site:</b> adrenaline, amikacin, amiodarone, atropine, benzylpenicillin, calcium chloride, calcium gluconate, caspofungin, cefazolin, cefotaxime, cefoxitin, caftazidime, cefuroxime, dexamethasone, dexmedetomidine, digoxin, dobutamine, dopamine, erythromycin, fentanyl, fluconazole, furosemide, gentamicin, glycopyrrolate, heparin, hydrocortisone, imipenem, insulin**, lidocaine, magnesium, meropenem, midazolam, milrinone, morphine, piperacillin, potassium chloride, ranitidine, ticarcillin, tobramycin, TPN, (no information re compatibility with lipid), vancomycin, vasopressin, voriconazole.</p> <p>* Variable reports on compatibility with insulin use separate line if possible</p>
<b>Incompatible With</b>	<p>Alkaline solutions, chlorpheniramine, chlorothiazide, diazepam, diazoxide, indomethacin, iron salts, nitrofurantoin, phenobarbital, phenytoin, sodium bicarbonate, streptomycin, sulfadiazine trimethoprim/sulfamethoxazole.</p> <p>No information to confirm that prostaglandin (alprostadil) is compatible, use a separate line.</p>
<b>Interactions</b>	<p>Concurrent treatment with betablockers, doxapram or monoamine oxidase inhibitors may cause hypertension.</p> <p>Concurrent treatment with halogenated anaesthetics or digoxin may precipitate arrhythmias</p>
<b>Monitoring</b>	Blood pressure, heart rate, urine output, peripheral perfusion.
<b>Stability</b>	<p>Readily oxidised do not use if solution is brown.</p> <p>Ampoules contain no preservative and are single use only.</p> <p>Discard any remaining contents immediately after use.</p> <p>Change iv infusion every 24 hours</p>

<b>Storage</b>	Protect from light. Store below 25°C
<b>Adverse Reactions</b>	Bradycardia, arrhythmias, breathing difficulties, headache, extravasation necrosis at injection site.
<b>Metabolism</b>	Rapidly and extensively metabolised. Plasma half life is approximately 2 minutes. Steady state concentrations are reached within 10 -15 minutes of commencing the infusion. Clearance is not influenced by renal function.
<b>Comments</b>	Not extensively studied in newborns but studies have shown improved systemic blood pressure, pulmonary blood flow and cardiac output in the setting of PPHN and septic shock
<b>References</b>	<ol style="list-style-type: none"> <li>1. <a href="http://www.medsafe.govt.nz">www.medsafe.govt.nz</a></li> <li>2. <a href="http://www.adhb.govt.nz/newborn/DrugProtocols/NoradrenalinePharmacology.htm">www.adhb.govt.nz/newborn/DrugProtocols/NoradrenalinePharmacology.htm</a></li> <li>3. Medicines for Children RCPCH 1999.BNF for Children 2011-2012</li> <li>4. Paediatric and Neonatal Dosage Handbook Taketomo et al. 19<sup>th</sup> Edition 2012.</li> <li>5. Trissell Handbook on Injectable Drugs in <a href="http://www.micromedexsolutions.com">www.micromedexsolutions.com</a></li> <li>6. Neofax in <a href="http://www.micromedexsolutions.com">www.micromedexsolutions.com</a></li> </ol>
<b>Updated By</b>	<p>A Lynn, B Robertshawe, N Austin Feb 2013</p> <p>A Lynn, B Robertshawe April 2017 (added TPN compatibility)</p> <p>A Lynn B Robertshawe October 2021 (update compatibility section and max concentration.</p> <p>A Lynn, B Robertshawe Sept 2023 (allow higher concentration option)</p>