

**ACICLOVIR****This drug must be guardrailed**

<b>Trade Name</b>	Aciclovir-Clariv <sup>®</sup> (Clariv) solution for IV Infusion (AFT Pharmaceuticals Ltd) Aciclovir- Baxter <sup>®</sup> solution for IV infusion (Baxter Pharmaceuticals) Aciclovir Univir <sup>®</sup>									
<b>Class</b>	Antiviral agent									
<b>Mechanism of Action</b>	Inhibits DNA synthesis and viral replication. Active metabolite is aciclovir triphosphate.									
<b>Indications</b>	Treatment of neonatal herpes simplex infection <b>Indication 1:</b> Skin eyes and mouth (SEM) disease <b>Indication 2:</b> CNS disease <b>Indication 3:</b> Disseminated disease <b>Indication 4:</b> Empirical treatment in infants born to mothers with evidence of <b>primary</b> herpes infection with vaginal delivery, or ruptured membranes >4 hrs or instrumentation during labour									
<b>Contraindications</b>	Hypersensitivity to aciclovir. Caution in patients with pre-existing renal disease or with concurrent use of other nephrotoxic drugs									
<b>Supplied As</b>	<b>IV:</b> 250mg/10mL vial, or 250mg dry powder for reconstitution (Univir brand)* <b>Oral:</b> 400mg dispersible tablets									
<b>Dilution*</b>  <b>*ONE dilution step required</b>	<b>CAUTION: check which brand is currently available as directions for dilution differ</b>  <b>IV: Clariv and Baxter brands* - 1 step dilution process</b>  <b>Further dilute the 25mg/mL solution in the vial as per the table below:</b>  <table border="1"> <thead> <tr> <th rowspan="2">Drug</th> <th>Add Diluent</th> <th rowspan="2">Total Volume</th> <th rowspan="2"><b>FINAL CONCENTRATION</b></th> </tr> <tr> <th>0.9% sodium chloride or 5% dextrose</th> </tr> </thead> <tbody> <tr> <td>100mg = 4mL</td> <td>16 mL</td> <td><b>20 mL</b></td> <td><b>5 mg / mL</b></td> </tr> </tbody> </table>	Drug	Add Diluent	Total Volume	<b>FINAL CONCENTRATION</b>	0.9% sodium chloride or 5% dextrose	100mg = 4mL	16 mL	<b>20 mL</b>	<b>5 mg / mL</b>
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<p><b>**TWO dilution steps required</b></p>	<p><b>IV: Univir brand** - 2 step dilution process</b></p> <p><b>Step 1. Reconstitute the vial</b></p> <table border="1" data-bbox="512 309 1453 584"> <thead> <tr> <th rowspan="2">Drug</th> <th>Add Diluent</th> <th rowspan="2">Total Volume</th> <th rowspan="2">Concentration</th> </tr> <tr> <th>Water for injection</th> </tr> </thead> <tbody> <tr> <td>250mg (dry powder)</td> <td>10 mL</td> <td>10 mL</td> <td>25 mg / mL</td> </tr> </tbody> </table> <p><b>Step 2. Further dilute the 25 mg/mL solution in step 1</b></p> <table border="1" data-bbox="512 707 1453 1028"> <thead> <tr> <th rowspan="2">Drug</th> <th>Add Diluent</th> <th rowspan="2">Total Volume</th> <th rowspan="2"><b>FINAL CONCENTRATION</b></th> </tr> <tr> <th>0.9% sodium chloride or 5% dextrose</th> </tr> </thead> <tbody> <tr> <td>100mg = 4mL</td> <td>16 mL</td> <td><b>20 mL</b></td> <td><b>5 mg / mL</b></td> </tr> </tbody> </table> <p><b>Oral:</b></p> <p>If acyclovir is prescribed orally then a suspension using a crushed tablet will be required to be freshly prepared as follows</p> <table border="1" data-bbox="512 1279 1453 1480"> <thead> <tr> <th rowspan="2">Tablet</th> <th>Add Diluent</th> <th rowspan="2"><b>FINAL CONCENTRATION</b></th> </tr> <tr> <th>Water for injection</th> </tr> </thead> <tbody> <tr> <td>400 mg</td> <td>20 mL</td> <td><b>20 mg / mL</b></td> </tr> </tbody> </table>	Drug	Add Diluent	Total Volume	Concentration	Water for injection	250mg (dry powder)	10 mL	10 mL	25 mg / mL	Drug	Add Diluent	Total Volume	<b>FINAL CONCENTRATION</b>	0.9% sodium chloride or 5% dextrose	100mg = 4mL	16 mL	<b>20 mL</b>	<b>5 mg / mL</b>	Tablet	Add Diluent	<b>FINAL CONCENTRATION</b>	Water for injection	400 mg	20 mL	<b>20 mg / mL</b>
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<p><b>Dosage</b></p>	<p><b>Depending on the dose, aciclovir can either be infused via the Alaris pump with the guardrail in place or for smaller volumes (&lt;10mL) via the T34 pump.</b></p> <p><b>Indication 1:</b> 20mg/kg/dose for 14 days  <b>Indication 2/3:</b> 20mg/kg/dose for 21 days  <b>Indication 4:</b> 20mg/kg/dose until PCR and viral culture results are negative or for up to 5 days.</p>																									
<p><b>Guardrails</b></p>	<p>Concentration: 5mg/mL  Soft Alert Min: 10mg/kg/hr    Hard Alert Max: 25mg/kg/hr  Soft Alert Max: 20mg/kg/hr    Default Setting: 20mg/kg/hr</p>																									
<p><b>Interval</b></p>	<p>8 hourly  12 hourly if &lt;34 weeks gestation, or renal impairment</p>																									

<b>Administration</b>	IV infusion over 1 hour Can give oral doses after initial iv treatment at 20mg/kg/dose 6 hourly in some clinical scenarios but this must be an SMO decision.
<b>Compatible With</b>	Solutions: Glucose 4% and Sodium Chloride 0.18%, Glucose 2.5% and Sodium Chloride 0.45%, Sodium Chloride 0.9% and 0.45%, Compound Sodium Lactate (Hartmann's)  Y site: Amphotericin B liposomal, benzylpenicillin, calcium gluconate, cefazolin, cefotaxime, ceftazidime, cefuroxime, dexamethasone, dexmedetomidine, digoxin, erythromycin lactobionate, fluconazole, furosemide, heparin sodium, hydrocortisone sodium succinate, imipenem+cilastin, insulin, metronidazole, milrinone, pancuronium bromide, phenobarbital sodium, potassium chloride, propranolol, ranitidine, sodium bicarbonate, sulfamethoxazole + trimethoprim, vancomycin, voriconazole, zidovudine
<b>Incompatible With</b>	TPN Amiodarone, caffeine citrate, ciprofloxacin, dobutamine, dopamine, gentamicin, hydralazine, meropenem, midazolam, morphine, naloxone, ondansetron, paracetamol, phenytoin, piperacillin + tazobactam, potassium dihydrogen phosphate  There is no information re compatibility with Smoflipid® and therefore infusion in the same line as aciclovir cannot be recommended.
<b>Monitoring</b>	Ensure adequate hydration and monitor renal function (consider impact of concomitant nephrotoxic drugs) Maintain good urine output for 2hrs post infusion.
<b>Stability</b>	Contains no antimicrobial preservative, dilute immediately before use and discard any unused diluted solution. Use a new vial for each dose.
<b>Storage</b>	Store at room temperature, DO NOT refrigerate – precipitate may form. Protect from light.
<b>Adverse Reactions</b>	Phlebitis may occur at IV site (due to alkaline pH of 10). If present dilution should be increased. Risk of transient renal impairment and crystalluria. Reversible neurological reactions have been reported (lethargy, seizures, agitation) Nausea and vomiting
<b>Metabolism</b>	Minimal metabolism in the liver. Primary route of excretion is the kidney. Protein binding <30%
<b>Comments</b>	Infusion solution should be <7mg/ml to minimise phlebitis Use of long courses of oral aciclovir for prevention of recurrent infection can cause transient neutropenia - routine use not yet recommended. Sodium content is approximately 2.67mg/mL.

<b>References</b>	<ol style="list-style-type: none"> <li>1. Trissell Handbook on Injectable Drugs 10<sup>th</sup> Edition</li> <li>2. NZHPA notes on Injectable Drugs 5<sup>th</sup> Edition</li> <li>3. Micromedex, Neofax, 2007</li> <li>4. NEJM 1991; 324:444-449</li> <li>5. Am J Obst Gyn 1991; 164: 569-76</li> <li>6. Pediatr Inf Dis J 1996; 15(3): 247-54</li> <li>7. Herpes. 11 Suppl 2:65A-76A, 2004 Jun</li> <li>8. Guidelines for the management of genital herpes in NZ; 8<sup>th</sup> Ed: 2007</li> <li>9. Aciclovir-Clarix data sheet. AFT Pharmaceuticals Ltd. Prepared 27<sup>th</sup> March 2014</li> </ol>																				
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