

<b>Alert</b>	High risk medicine. The Antimicrobial Stewardship Team recommends this drug is listed under the following category: Restricted.																							
<b>Indication</b>	As part of therapy for suspected meningitis. Treatment of proven meningitis and sepsis caused by susceptible organisms (e.g., <i>E.coli</i> , <i>H. influenzae</i> , <i>Klebsiella</i> spp.).																							
<b>Action</b>	Bactericidal agent which inhibits cell wall synthesis in susceptible bacteria. Broad spectrum against gram positive and many gram negative organisms but not <i>Pseudomonas</i> species.																							
<b>Drug type</b>	Cephalosporin antibiotic.																							
<b>Trade name</b>	Cefotaxime Sandoz, DBL Cefotaxime Sodium																							
<b>Presentation</b>	500 mg and 1 g vial																							
<b>Dose</b>	<p><b>50 mg/kg/dose.</b></p> <table border="1" style="width: 100%; border-collapse: collapse;"> <thead> <tr> <th style="text-align: center;">Corrected Gestational Age/Postmenstrual Age</th> <th style="text-align: center;">Postnatal Age</th> <th style="text-align: center;">Interval</th> </tr> </thead> <tbody> <tr> <td style="text-align: center;">&lt; 30<sup>+0</sup> weeks</td> <td style="text-align: center;">0–28 days</td> <td style="text-align: center;">12 hourly</td> </tr> <tr> <td style="text-align: center;">&lt; 30<sup>+0</sup> weeks</td> <td style="text-align: center;">≥29 days</td> <td style="text-align: center;">8 hourly</td> </tr> <tr> <td style="text-align: center;">30<sup>+0</sup>–36<sup>+6</sup> weeks</td> <td style="text-align: center;">0–14 days</td> <td style="text-align: center;">12 hourly</td> </tr> <tr> <td style="text-align: center;">30<sup>+0</sup>–36<sup>+6</sup> weeks</td> <td style="text-align: center;">≥15 days</td> <td style="text-align: center;">8 hourly</td> </tr> <tr style="background-color: #ffff00;"> <td style="text-align: center;">≥ 37<sup>+0</sup> weeks</td> <td style="text-align: center;">0–7 days</td> <td style="text-align: center;">8 hourly</td> </tr> <tr style="background-color: #ffff00;"> <td style="text-align: center;">≥ 37<sup>+0</sup> weeks</td> <td style="text-align: center;">≥8 days</td> <td style="text-align: center;">6 hourly</td> </tr> </tbody> </table>			Corrected Gestational Age/Postmenstrual Age	Postnatal Age	Interval	< 30 <sup>+0</sup> weeks	0–28 days	12 hourly	< 30 <sup>+0</sup> weeks	≥29 days	8 hourly	30 <sup>+0</sup> –36 <sup>+6</sup> weeks	0–14 days	12 hourly	30 <sup>+0</sup> –36 <sup>+6</sup> weeks	≥15 days	8 hourly	≥ 37 <sup>+0</sup> weeks	0–7 days	8 hourly	≥ 37 <sup>+0</sup> weeks	≥8 days	6 hourly
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<b>Route</b>	IV IM																							
<b>Preparation</b>	<p><b>IV</b> Add 9.8 mL of water for injection to the <b>500 mg powder</b> to make a 50 mg/mL solution OR Add 9.6 mL of water for injection to the <b>1 g powder</b> to make a 100 mg/mL solution.</p> <p><b>IM injection</b> Add 2 mL of water for injection to the <b>500 mg powder</b> to make a 230 mg/mL solution OR Add 3 mL of water for injection to the <b>1 g powder</b> to make a 300 mg/mL solution.</p>																							
<b>Administration</b>	<p><b>IV bolus:</b> over 3–5 minutes. <b>IV infusion:</b> over 15–30 minutes <b>IM injection:</b> Inject deep into the large muscle.</p>																							
<b>Monitoring</b>	Cefotaxime has a high therapeutic index. Consider monitoring renal function, blood count and electrolytes if therapy is prolonged.																							
<b>Contraindications</b>	Hypersensitivity to cefotaxime or other cephalosporins or previous history of major allergic response to a penicillin.																							
<b>Precautions</b>	Liver and renal disease. Sodium restriction – cefotaxime contains 48.2 mg/g (2.1 mmol/g) sodium.																							
<b>Drug interactions</b>	May potentiate the renal toxicity of nephrotoxic drugs. Should not be combined with bacteriostatic antibiotics (e.g., tetracycline, erythromycin or chloramphenicol) since there may be a potential antagonistic effect.																							
<b>Adverse reactions</b>	Leucopaenia, granulocytopaenia, agranulocytosis. Moderate and transient rise in liver enzymes and/or bilirubin. Hypersensitivity reactions. Arrhythmias have occurred in patients who received rapid IV administration through a central venous catheter. Fungal sepsis. Bacterial resistance.																							
<b>Compatibility</b>	Fluids: Glucose 5%, glucose 10%, Hartmann's, sodium chloride 0.9%																							

	Y site: Amino acid solutions, aciclovir, amifostine, aztreonam, bivalirudin, dexmedetomidine, granisetron, hydromorphone, magnesium sulfate, midazolam, morphine sulfate, pethidine, remifentanyl, tigecycline.
<b>Incompatibility</b>	Fluids: Alkaline solutions e.g., containing sodium bicarbonate.  Y site: Aminoglycosides – amikacin, gentamicin, tobramycin; azathioprine, azithromycin, caspofungin, chloramphenicol, chlorpromazine, dobutamine, dolasetron, filgrastim, fluconazole, ganciclovir, haloperidol lactate, hydralazine, labetalol, methylprednisolone sodium succinate, mycophenolate mofetil, pentamidine, phenobarbitone, phentolamine, promethazine, protamine, sodium bicarbonate, vecuronium.
<b>Stability</b>	Reconstituted solution is stable for 24 hours at 2 to 8 °C. Protect from light. Do not use if powder or solutions have darkened in colour.
<b>Storage</b>	Store below 25°C Protect from light.
<b>Excipients</b>	
<b>Special comments</b>	The main metabolite of cefotaxime is desacetylcefotaxime. This metabolite is active and is thought to enhance activity against Gram negative organisms. It has a longer half-life than cefotaxime. The major route of clearance of both cefotaxime and desacetylcefotaxime is renal.
<b>Evidence</b>	To be updated.
<b>Practice points</b>	
<b>References</b>	<ol style="list-style-type: none"> <li>1. Aujard Y, Brion F, Jacqz-Aigrain E, et al: Pharmacokinetics of cefotaxime and desacetylcefotaxime in the newborn. <i>Diagn Microbial Infect Dis</i> 1989;12:87–91.</li> <li>2. Jacobs R, Kearns G: Cefotaxime and deacetylcefotaxime in neonates and children: a review of microbiologic, pharmacokinetic and clinical experience. <i>Diagn Microbial Infect Dis</i> 1989;12:93–99.</li> <li>3. Kafetzis D, Brater D, Kapiki A, et al: Treatment of severe neonatal infections with cefotaxime. Efficacy and pharmacokinetics. <i>The Journal of Pediatrics</i> 1982;100:483–489.</li> <li>4. Kearns G, Young R: Pharmacokinetics of cefotaxime and deacetylcefotaxime in the young. <i>Diagn Microbial Infect Dis</i> 1995;22:97–104.</li> <li>5. Kearns G, Jacobbs R, Thomas B, et al: Cefotaxime and desacetylcefotaxime pharmacokinetics in very low birth weight neonates. <i>The Journal of Pediatrics</i> 1989;114:461–7.</li> <li>6. Odio C: Cefotaxime for treatment of neonatal sepsis and meningitis. <i>Diagn Microbial Infect Dis</i> 1995;22:111–117.</li> <li>7. Sivanandan S, Soraisham A, Swarnam K: Choice and duration of antimicrobial therapy for neonatal sepsis and meningitis. <i>International Journal of Pediatrics</i>, 2011;712150. doi: 10.1155/2011/71215.</li> <li>8. Craig W: Interrelationship between pharmacokinetics and pharmacodynamics in determining dosage regimens for broad spectrum cephalosporins. <i>Diagn Microbial Infect Dis</i> 1995;22:89–96</li> <li>9. Pacifici G: Pharmacokinetics of cephalosporins in the neonate: a review. <i>Clinics</i> 2011;66(7):1267–1274.</li> <li>10. Young T, Mangum B <i>Neofax</i> 23rd edition, Thomson Reuters 2010.</li> <li>11. <i>Australian Injectable Drugs Handbook</i>. 5th Edition. The Society of Hospital Pharmacists of Australia. 2011.</li> <li>12. MIMS online via CIAP accessed 7<sup>th</sup> July 2015.</li> <li>13. Cotten CM, McDonald S, Stoll B, Goldberg RN, Poole K, Benjamin DK Jr, National Institute for Child Health and Human Development Neonatal Research Network. The association of third-generation cephalosporin use and invasive candidiasis in extremely low birth-weight infants. <i>Pediatrics</i> 2006;118(2):717–22.</li> <li>14. Calil R, Marba ST, von Nowakowski A, Tresoldi AT. Reduction in colonization and nosocomial infection by multiresistant bacteria in a neonatal unit after institution of educational measures and restriction in the use of cephalosporins. <i>Am J Infect Control</i> 2001;29(3):133–8.</li> <li>15. Dellagrammaticas HD, Christodoulou C, Megaloyanni E, Papadimitriou M, Kapetanakis J, Kourakis G. Treatment of gram-negative bacterial meningitis in term neonates with third generation cephalosporins plus amikacin. <i>Biol Neonate</i> 2000;77(3):139–46.</li> <li>16. Harvey D, Holt DE, Bedford H. Bacterial meningitis in the newborn: a prospective study of mortality and morbidity. <i>Semin Perinatol</i> 1999;23(3):218–25.</li> <li>17. <i>Neofax</i> accessed on <a href="http://www.neofax.micromedex.solutions.com">www.neofax.micromedex.solutions.com</a> on 29<sup>th</sup> July 2015.</li> </ol>

<b>VERSION/NUMBER</b>	<b>DATE</b>
Original 1.1	08/08/2015
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