# Louth: Antimicrobial Guidelines - Louth Hospitals: Antimicrobial Guidelines: Neonatal - ■Vancomycin IV

### Vancomycin Hydrochloride

Vancomycin is a bactericidal antibiotic with activity against some aerobic and anaerobic bacteria including multi-resistant staphylococci.

# MEDICATION SAFETY ISSUES

- Incompatible with Cephalosporins, Phenobarbital and Piperacillin-tazobactam.[1]
- Red man syndrome: flushing and hypotension if administered too quickly.
- Vancomycin exposure may be associated with hearing impairment in neonates.[2] Concomitant use of other ototoxic medication, e.g. furosemide and loop diuretics, should be avoided if possible.

## USES

Drug of choice for serious infections caused by methicillin-resistant staphylococci and penicillin-resistant pneumococci.[1]

## PRESENTATION

Vancomycin Hydrochloride 500mg Powder for concentrate for solution for infusion.[3]

# DOSAGE [4-6 ] :

DOSE: 15mg/kg for all ages without renal impairment. Preterm (<37/40 weeks gestation) Weight ≤7 days of age >7 days of age <1.2kg 18 HOURLY 12 HOURLY 1.2-2kg >2kg 12 HOURLY 8 HOURLY Term ( > 37/40 weeks gestation) ⊳7 days of age<u><</u> 28 days of age 27 days of age 12 HOURLY 3 HOURLY Term >28 days 6 HOURLY

If renal impairment, contact Consultant Microbiologist or Pharmacy for advice.

### RECONSTITUTION

### Warning- reconstitution is a two step process

Step 1 The displacement value for Vancomycin Mylan brand 500mg (stocked in OLOL) is 0.3ml. Add 9.7ml of Water for Injection to a 500mg vial and shake gently to dissolve. The resulting solution contains 50mg/ml vancomycin.

Step 2 Further dilute 1ml of the reconstituted (50mg/ml) vancomycin solution with 9ml of compatible fluid (e.g. Sodium chloride 0.9%, Glucose 5%). Ensure adequate mixing by inverting the syringe at least 10 times. The resulting solution contains **5mg/ml of vancomycin**.

# ADMINISTRATION

Use immediately once reconstituted and diluted. Infuse the prescribed dose over at least 60 minutes.

## STORAGE

Do not store above 25 ° C. Use immediately once reconstituted.

#### MONITORING

Record vancomycin levels and response on the Vancomycin Monitoring Form (see Appendix 1).

Sub-therapeutic levels can result in treatment failure or the emergence of drug resistance.

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Toxic or high levels of vancomycin can result in nephrotoxicity and/or ototoxicity. Monitor creatinine and urea at least twice weekly for the duration of treatment. If the patient is on additional nephrotoxic medication (e.g. NSAID's, aciclovir, aminoglycosides, diuretics, omeprazole), monitor renal function more frequently.

Indication	Target Trough Concentration
Uncomplicated infections	10 to 20 mg/L
•	5
Complicated infections:	15 to 20 mg/L
(e.g. Bacteraemia, endocarditis, osteomyelitis, meningitis, necrotising	
fasciitis and empyema)	
Dosing Frequency	When to take a trough level
18 HOURLY	Up to ONE HOUR before the 2 <sup>nd</sup> dose
	When levels are therapeutic, repeat every 2 days
12 HOURLY	Up to ONE HOUR before the 3rd or 4th dose
	When levels are therapeutic, repeat every 2 days
8 HOURLY	
BHOUKLI	Up to ONE HOUR before 4th or 5th dose
	When levels are therepoulting report evenu 2 deve
	When levels are therapeutic, repeat every 2 days
6 HOURLY	Up to ONE HOUR before 4 <sup>th</sup> , 5th or 6th dose
	When levels are therapeutic, repeat every 3 days
In patients with normal renal function, <u>DO NOT withhold</u> the next dose v	while awaiting the result of the trough level – this may result in the
patient being under dosed.	

If patient has renal impairment, contact Consultant Microbiologist or Pharmacy for advice.

## Vancomycin Targets and Timing of Levels:

Trough Level	Management	Timing of Next Level
Less than 5mg/L	Preterm & term < 1 month: Increase frequency	Complete 24 hours of new regime before checking
	18hours■12hours■8hours>6 hours. For example	next level. Following two consecutive levels within
	18 hours then increase to 12 hourly.	therapeutic range repeat every 2-3 days and
	Term >1 month : Increase by 20%.	monitor renal function.
5-9mg/L	Increase by 10%	*Complicated infections: Severe infection, reduced
10-20mg/L	No change (unless target level is 15-20mg/L for complicated infections* contact micro/ID)	sensitivities, bacteraemia, endocarditis, osteomyelitis, meningitis, necrotising fasciitis and
21-24mg/L	Decrease by 10% & Contact Micro	empyema
≥ 25mg/L	Contact microbiology/ ID or Pharmacy for advice	1

## SIDE EFFECTS

- Nephrotoxicity and ototoxicity.
- Rash and hypotension (red man syndrome) if given too quickly.
- Phlebitis which can be minimized by slow infusion and dilution of the drug.

#### SAMPLE CALCULATION

0.86kg neonate 28 weeks corrected gestational age.

Dose: 15mg/kg = 15mg x 0.86kg = 12.9mg

Reconstitute and dilute one 500mg vial of vancomycin following the directions above.

The resulting solution contains 5mg/ml of vancomycin.

5mg in 1ml is equivalent to 1mg in 0.2ml

12.9mg in 2.58ml

Infuse a total volume of 2.58ml at a rate of 2.58ml/hr over 60 minutes.

## REFERENCES

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- 1. <u>Reuters, T., Neofax A manual of drugs used in neonatal care</u>. Vol. 24. 2011.
- 2. Vella-Brincat, J.W., et al., Are gentamicin and/or vancomycin associated with ototoxicity in the neonate? A retrospective audit. Neonatology, 2011. 100 (2): p. 186-93.
- 3. Limited, H.U. Vancomycin Hydrochloride 500 mg and 1 g Powder for Concentrate for Infusion . 2009; Available from: http://www.medicines.org.uk/emc/medicine/20059/SPC .
- 4. <u>British Medical Association, et al., BNF for Children. Accessed via www.medicinescomplete.com</u> November 13th 2018. 2018, BMJ Group and Pharmaceutical Press: London.
- 5. Pickering, L.K., 2009 Red Book: Report of the Committee on Infectious Diseases . 28th ed ed. 2009: American Academy of Paediatrics.
- 6. <u>uptodate.com</u>. *Vancomycin: Pediatric Drug Information* . 2018.

#### Appendix 1: Neonatal Vancomycin Monitoring Form - see hardcopy in NICU folder

### Summary of Changes from Previous Versions

Date	Change
Jul 2019	Revision 2 of the Rotunda Neonatal Vancomycin Monograph received with
	permission (approved in the Rotunda in Dec 2018). Changes from previous
	version:
	<ul> <li>Medication Safety Issues: Risk of ototoxicity added.</li> </ul>
	<ul> <li>Dosage: Dose regimen changed based on audits undertaken in the</li> </ul>
	Rotunda Hospital, which found frequent low trough level results with the
	previous dosing regimen from BNFc. Updated dosing regimen based on
	American Academy of Paediatrics Red Book 2009.
	<ul> <li>Monitoring: New addition of Vancomycin Monitoring Form (see Appendix</li> </ul>
	1). Addition of advice to monitor urea and creatinine at least twice week
	and more frequently if patient on other nephrotoxic medications.
	<ul> <li>Table of when to take trough levels updated.</li> </ul>
	<ul> <li>Addition of a new table on the management of subsequent doses based</li> </ul>
	on trough level results.
	<ul> <li>Sample calculation - similar calculation to previous version, neonate</li> </ul>
	weight different.
	<ul> <li>Side effects - section shortened to the main side effects only. See BNFc</li> </ul>
	for full information.
	Additional information included in OLOL monograph:
	<ul> <li>Contact Micro/Pharmacy for advice on dosing regimen if renal</li> </ul>
	impairment.
	<ul> <li>Added displacement value for vancomycin to reconstitution section.</li> </ul>
	<ul> <li>Contact Micro/Pharmacy for advice on whether to hold dose pending</li> </ul>
	level result if renal impairment.
June 2017	Vancomycin Flynn brand now stocked in OLOL. Displacement value of
	500mg vial now 0.4ml. Monograph updated to reflect this.
Jan 2015	This is the first version of this guideline. It is based on the Rotunda Hospital
	Neonatal Monograph for Vancomycin, Doc. No. 1, Revision No. 0, date of
	issue 10/11/14.
	Changes in OLOL monograph compared to the Rotunda monograph:
	<ul> <li>Medication Safety: further detail on "red man" syndrome from BNFc.</li> </ul>
	<ul> <li>Dose: added to check BNFc and seek advice on dose if patient has renal</li> </ul>
	impairment.
	<ul> <li>Reconstitution: option to further dilute 2ml of 50mg/ml with 18ml of diluen</li> </ul>
	if high dose
	<ul> <li>Monitoring: added to contact Consultant Neonatologist if trough level out</li> </ul>
	of range, hold if > 20mg/L.
	<ul> <li>Adverse Effects: full list from BNF for Children included.</li> </ul>
	Adverse Effects, full list from Birl for Gillidreft filliduded.

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