

Therapeutic Drug Monitoring of Aminoglycosides

Aminoglycosides (gentamicin, tobramycin and amikacin) are bactericidal antibiotics used to treat infections caused by Gram-negative organisms. Therapeutic drug monitoring is recommended to ensure adequate dosing and prevent adverse effects such as nephrotoxicity and ototoxicity. Aminoglycosides demonstrate concentration-dependent bactericidal activity, which is optimal when the maximum concentration (C_{max}) is ≥ 10 times the minimum inhibitory concentration (MIC) of the organism. A MIC of 1 mg/L can be assumed for most organisms; however, higher MICs (e.g. ≥ 2 mg/L) are common with *Pseudomonas aeruginosa*. These peak concentrations correspond to an area under the concentration-time curve (AUC_{24}) of 70-100 mg/L.h for gentamicin and tobramycin. Prolonged high AUC_{24} and minimum concentrations (C_{min}) are more likely to cause toxicity.

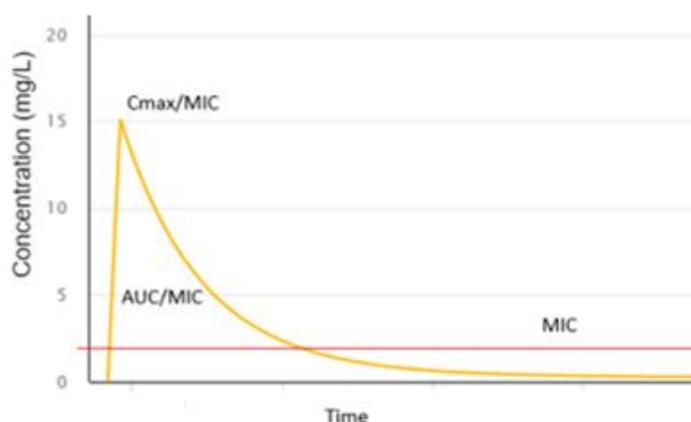
Pharmacokinetic/Pharmacodynamic Targets*

Gentamicin or tobramycin

- AUC_{24} : 70-100 mg/L.h
- C_{max} : 15-30 mg/L
- C_{min} : < 1 mg/L

Amikacin

- AUC_{24} : 160-200 mg/L.h
- C_{max} : 30-60 mg/L
- C_{min} : < 2 mg/L



*For adult patients receiving once daily dosing, and excluding those with endocarditis

- Aim for a C_{max} at the higher end of the range for infections caused by *Pseudomonas aeruginosa*, given that its MIC is likely > 1 mg/L.
- Aminoglycosides concentrate in the urine at higher concentrations than in the plasma, so an AUC_{24} at the lower end of the range (e.g. 70 mg/L.h) is usually sufficient for a urinary tract infection.
- Aim for a higher AUC_{24} (e.g. 80-100 mg/L.h) for more serious infections.
- Giving a higher dose and extending the dose interval (e.g. to 48 hours) is sometimes needed to attain both the optimal C_{max} and C_{min} targets.

For a comprehensive guide on therapeutic drug monitoring for both aminoglycosides and vancomycin see:

<http://www.medicinesinformation.co.nz/wp-content/uploads/2022/05/Aminoglycosidevancomycin-TDM-workbook-6th-edition-2022.pdf>