Adult Aminoglycoside Dosing for Gram-negative infections prior to available serum levels
(Excludes patients with cystic fibrosis, OB-GYN patients and surgical prophylaxis)

**Cr Cl ≥ 40 mL/min**
- Gentamicin and Tobramycin
  - 5 - 7 mg/kg INT-Q24H or INT-Q36H
  - (use 3 - 5 mg/kg for cystitis)
  - Obtain a random level 10 - 12 hr after the start of infusion for the 1st dose
  - Adjust frequency by Nomogram (see pg. 2 - 4)

**Cr Cl < 40 mL/min or CVVHD or Age > 65**
- Gentamicin and Tobramycin
  - 3 - 5 mg/kg x1 dose
  - Obtain 2 random levels:
    - (1) 3 hrs after dose
    - (2) 16 hrs after dose
  - Call or Text PK Service to follow patient (631-487-6728)

**HD (assume 3x/wk HD)**
- Gentamicin and Tobramycin
  - 1st dose 2 - 3 mg/kg x1
  - (1) Obtain Pre-HD level before next HD session
  - (2) If pre-HD level is less than 3 mcg/mL, give 1 - 1.5 mg/kg after HD session is completed
  - (3) Call or Text PK Service to follow patient (631-487-6728)

- Amikacin
  - 1st dose 10 mg/kg x1
  - (1) Obtain Pre-HD level before next HD session
  - (2) If pre-HD level is less than 10 mcg/mL, give 5 - 7.5 mg/kg after HD session is completed
  - (3) Call or Text PK Service to follow patient (631-487-6728)

**Dosing weight for aminoglycoside weight-based dosing methods:**
- Use Ideal body weight unless Actual body weight is less than Ideal body weight (IBW)
- If Actual Body Weight is greater than 120% of Ideal body weight, use Adjusted Dosing Weight
  
  \[ \text{Adjusted Dosing Weight} = \text{Ideal body weight} + [0.4 \times (\text{Actual body weight} - \text{Ideal body weight})] \]
Monitoring for High-Dose Extended Interval Dosing Method

For gentamicin or tobramycin dosing at 7mg per kg, use Hartford Hospital Dosing Nomogram to determine dosing frequency

How to use Hartford Hospital Nomogram when gentamicin or tobramycin is dosed at 7mg/kg:

* Obtain a random level 10 - 12 hours after the first dose
* Plot the measured gentamicin or tobramycin levels on the Hartford Nomogram corresponding to the number of hours from when the infusion was started
  * If the level falls within “Q24h” area, the dosing frequency is INT-Q24h.
  * If the level falls within “Q36h” area, the dosing frequency is INT-Q36h. Start new regimen 36 hours from the last dose and repeat a 10 - 12 hours post-dose random level.
  * If the level falls within “Q48h” area, the dosing frequency is INT-Q48h. Start new regimen 48 hours from the last dose and repeat a 10-12 hours post-dose random level.
* If the plotted level falls on or close to a division line, use the dosing frequency with the longer dosing interval
* If the plotted level falls on or above the upper line of q48h, discontinue current regimen and order for a random level 24 hours from the last dose. Call or text PK Service for assistance. (631-487-6728)

Monitoring after dosing frequency is confirmed:

* Monitor renal function
* If renal function is unchanged, recheck a random level 10-12 hours after dose every 5-7 days
* More frequent monitoring may be warranted in patients with higher risk for nephrotoxicity or unstable renal function
* When decline in renal function is detected, reassess dosing frequency by a random level 10-12 hours after the dose
SBUH Aminoglycoside Dosing Protocol

Antimicrobial Stewardship Program

Monitoring for High-Dose Extended Interval Dosing Method

For gentamicin or tobramycin dosing at 5 mg per kg, use Barnes-Jewish Hospital Dosing Nomogram to determine dosing frequency

How to use Barnes-Jewish Nomogram when gentamicin or tobramycin is dosed at 5 mg/kg

* Obtain a random level 10 - 12 hours after the first dose
* Plot the measured gentamicin or tobramycin level on the Barnes-Jewish Hospital Nomogram corresponding to the number of hours from the end of the infusion
* If the level falls within “Q24h” area, the dosing frequency is INT-Q24h.
* If the level falls within “Q36h” area, the dosing frequency is INT-Q36h. Start new regimen 36 hours from the last dose and repeat a 10-12 hours post-dose random level.
* If the level falls within “Q48h” area, the dosing frequency is INT-Q48h. Start new regimen 48 hours from the last dose and repeat a 10-12 hours post-dose random level.
* If the plotted level falls on or close to a division line, use the dosing frequency with the longer dosing interval
* If the plotted level falls on or above the upper line of q48h, discontinue current regimen and order for a random level 24 hours from the last level. Call or text PK Service for assistance. (631-487-6728)

Monitoring after dosing frequency is confirmed:

* Monitor renal function
* If renal function is unchanged, recheck a random level 10-12 hours after dose every 5-7 days
* More frequent monitoring may be warranted in patients with higher risk for nephrotoxicity or unstable renal function
* Whenever decline in renal function is detected, reassess dosing frequency by a random level 10-12 hours after the dose
Monitoring for High-Dose Extended Interval Dosing Method

For amikacin dosing at 15 mg/kg, use Barnes-Jewish Hospital Dosing Nomogram to determine dosing frequency.

How to use Barnes-Jewish Nomogram when amikacin is dosed at 15 mg/kg

- Obtain a random level 10 - 12 hours after the first dose.
- Plot the measured amikacin level on the Barnes-Jewish Hospital Nomogram corresponding to the number of hours from the end of the infusion.
- If the level falls within “Q24h” area, the dosing frequency is INT-Q24h.
- If the level falls within “Q36h” area, the dosing frequency is INT-Q36h. Start new regimen 36 hours from the last dose and repeat a 10-12 hours post-dose random level.
- If the level falls within “Q48h” area, the dosing frequency is INT-Q48h. Start new regimen 48 hours from the last dose and repeat a 10-12 hours post-dose random level.
- If the plotted level falls on or close to a division line, use the dosing frequency with the longer dosing interval.
- If the plotted level falls on or above the upper line of q48h, discontinue current regimen and order for a random level 24 hours from the last level. Call or text PK Service for assistance. (631-487-6728)

Monitoring after dosing frequency is confirmed:

- Monitor renal function.
- If renal function is unchanged, recheck a random level 10-12 hours after dose every 5-7 days.
- More frequent monitoring may be warranted in patients with higher risk for nephrotoxicity or unstable renal function.
- Whenever decline in renal function is detected, reassess dosing frequency by a random level 10-12 hours after the dose.
**Gentamicin Gram-positive Synergy Dosing**

Gentamicin should not be used as monotherapy for Gram-positive infections. A low dose of gentamicin is used in combination with a cell wall active antimicrobial agent (e.g., beta-lactam antibiotics or vancomycin) for synergy in the treatment of Gram-positive infections (e.g., endovascular infection with *Enterococcus, Staphylococcus aureus*, Penicillin-resistant *Streptococcus*, and *Listeria*).

**Target steady state peak concentration:** 3 – 4 mcg/mL

**Target steady state trough concentration:** Less than 1 mcg/mL

- **Cr Cl ≥ 60 mL/min**
  - 1 mg/kg Q8H
  - (1) Obtain peak level - 30 min after the infusion of the 4th dose ends
  - (2) Obtain trough level - within 30 min prior to the 5th dose
  - (3) Adjust dosing based on target peak and trough concentration or call Pk service to follow patient (631-487-6728)

- **Cr Cl 30 - 59 mL/min or Age > 65**
  - 1 mg/kg Q12H
  - (1) Obtain peak level - 30 min after the infusion of the 4th dose ends
  - (2) Obtain trough level - within 30 min prior to the 5th dose
  - (3) Call PK Services to follow patient (631-487-6728)

- **Cr Cl less than 30 mL/min or CVVHD**
  - 1 mg/kg q24h
  - (1) Obtain peak level - 30 min after infusion of the first dose ends
  - (2) Obtain a random level - 18 hours after the first dose
  - (3) Call PK Service to follow patient (631-487-6728)

- **Intermittent HD (assume 3x/week HD)**
  - 1 mg/kg x1 dose
  - (1) Obtain pre-HD level prior the next HD session
  - (2) If pre-HD level is ≤ 1 mcg/mL, give 1 mg/kg after HD
  - (3) Call PK Service to follow patient (631-487-6728)

- If Actual Body Weight is greater than 120% of Ideal body weight, use Adjusted Dosing Weight
  - Adjusted Dosing Weight = Ideal body weight + [0.4 x (Actual body weight – Ideal body weight)]
Target Levels

I. Aminoglycoside High Dose Extended Interval Method (Also known as Once-Daily Dosing)

Gentamicin and Tobramycin
When dosed at 5 to 7 mg per kg with extended dosing interval (e.g., Q24H, Q36H, or Q48H), the usual peak concentration is in the range of 15 to 25 mcg/mL. The goal for the trough concentration is less than 0.3 mcg/mL.

Amikacin
When dosed at 15 to 20 mg per kg with extended dosing interval (e.g., Q24H, Q36H, or Q48H), the usual peak concentration is in the range of 35 to 50 mcg/mL. The goal for the trough concentration is less than 4 mcg/mL.

II. Conventional Dosing
This dosing method uses patient’s estimated pharmacokinetic parameters derived from measured serum concentrations to determine dose and frequency to achieve target peak and trough concentrations. This dosing method is used when High Dose Extended Interval Dosing Method cannot be used for the patients who have decreased renal function.

Conventional dosing interval of q24h should not be confused with patients receiving High Dose Extended Interval Method (Once-daily). The dose used in Conventional Dosing Method is reduced in order to attain the optimal PK/PD target in patients with decreased renal function.

Target Peak and Trough at steady state for Conventional dosing

<table>
<thead>
<tr>
<th>Gentamicin and Tobramycin</th>
<th>Target Peak at steady state (mcg/ml)</th>
<th>Target Trough at steady state (mcg/ml)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Gram-negative Pneumonia</td>
<td>8-10</td>
<td>&lt; 2</td>
</tr>
<tr>
<td>Severe Gram negative Infections</td>
<td>6-8</td>
<td>&lt; 2</td>
</tr>
<tr>
<td>Urinary Tract Infections</td>
<td>4-6</td>
<td>&lt;1</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Amikacin</th>
<th>Target Peak at steady state (mcg/ml)</th>
<th>Target Trough at steady state (mcg/ml)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Gram-negative infections</td>
<td>25 - 35</td>
<td>&lt; 8</td>
</tr>
</tbody>
</table>
References