

(This summary is designed only for the lecture; it never designed for any exam.)

CLINICAL PHARMACOKINETIC FOR VANCOMYCIN:

A-Initial dose determination.

B-Use of vancomycin concentration to change the dose.

Initial dose determination:

- 1- Pharmacokinetic dosing methods.
- 2- Moellering nomogram method.
- 3- Matzke nomogram method
- 4- Literature-based methods.

Notes:

- ✓ vancomycin dose should be rounded to nearest 100-250
- ✓ Dose interval should be rounded into clinically accepted intervals (12, 18, 24, 36, 48, 72, multiples of 24)

A-Pharmacokinetic dosing methods: You need:

1- Creatinine clearance

$$\text{CrCl male} = \frac{(140 - \text{Age}) * \text{wt}}{72 * \text{Sc}}$$

$$\text{CrCl female} = 0.85 \frac{(140 - \text{Age}) * \text{wt}}{72 * \text{Sc}}$$

(Cockcroft and Gault Eq. if the Pt within 30% of its IBW (**overweight** % = $(\text{ABW} - \text{IBW}) / \text{IBW}$) and have stable renal function, Wt is patient's weight in kg, sc is serum creatinine in mg/dL, Age in years)

$$\text{CrCl male} = \frac{(137 - \text{age})[(0.285 * \text{Wt}) + (12.1 * \text{Ht}^2)]}{(51 * \text{Sc})}$$

$$\text{CrCl female} = \frac{(146 - \text{age})\{(0.287 * \text{Wt}) + (9.74 * \text{Ht}^2)\}}{(60 * \text{Sc})}$$

(Salazar and Corcoran Eq. if the Pt above 30% of its IBW (**overweight** % = $(\text{ABW} - \text{IBW}) / \text{IBW}$) and have stable renal function, Ht is the height in meter, Wt is patient weight in kg, sc is serum creatinine in mg/dL, Age in years)

2- Clearance estimate for vancomycin in term of ml/min/kg

$$\text{Cl} = 0.695(\text{CrCl}/\text{kg}) + 0.05 \quad (\text{use the total body weight even in obese})$$

3- Volume of distribution estimate

$$0.7 \text{ mg/kg} \quad (\text{Use the ideal body weight in obese})$$

$$\text{IBW (male)} = 50 + 2.3 (\text{H} - 60) \dots\dots\dots \text{IBW (female)} = 45.5 + 2.3 (\text{H} - 60)$$

H = height in inches, 5F= 60-inch, Inch=2.54 cm

4-Elimination rate constant & t ½

$$K_{el} = \frac{Cl \text{ (ml/min kg)}}{Vd \left(\frac{L}{kg}\right)} \quad (\times 60/1000 \text{ to get in h}^{-1})$$

(Note: in obese patient, multiply CL by ABW and Vd by ideal body weight)

5- Desired steady state concentration

Condition	Peak in µg/mL	trough in µg/mL
generally,	20-40	5-15

6- Equation to compute the dose

$$\tau = \frac{\ln C_{ss \text{ max}} - \ln C_{ss \text{ min}}}{K_{el}}$$

$$MD = C_{ss \text{ max}} * Vd (1 - e^{-K_{el} \times \tau})$$

$$LD = C_{ss \text{ max}} * Vd$$

2-Moellering nomogram method:

1- Compute patient's creatinine clearance (CrCl) using Cockcroft–Gault method for normal weight or Salazar-Corcoran method for obese patients (**assuming patient have stable renal function and adult**)

2- Calculate maintenance dose according to this equation:

$$D \text{ (mg/h/kg)} = 0.626 \text{ (CrCl/kg)} + 0.05$$

3- If the patient has normal renal function use 12h and his/her weight to get the maintenance dose:

Ex. if MD is 0.85 mg/h/kg & CrCl is 130 ml/min, patient wt is 70.....

Sol: $MD = 0.85 \times 70 \times 12 = 714$

But use the standard dose 1000 mg to get the suitable dose interval for patient with compromised renal function

Ex. if MD is 0.32 mg/h/kg & CrCl is 30 ml/min, patient wt is 70.....

Sol: $0.32 \text{ mg/kg/h} \times 70 \text{ kg} = 22.4 \text{ mg/h}$ 22.4 mg/1h
 $= 1000 \text{ mg/?}$

Dose interval = 44.64 h Or Dose interval = $1000 / \text{wt} \times D$
(mg/h/kg)

4- Loading dose of 15 mg/kg should be given for patients with significant renal function impairment

3-Matzke Nomogram Method

1. Compute patient's creatinine clearance (CrCl) using Cockcroft–Gault method: $\text{CrCl} = [(140 - \text{age}) \text{ BW}] / (\text{Scr} \times 72)$. Multiply by 0.85 for females.

2. Nomogram not verified in obese individuals.

3. Dosage chart is designed to achieve peak serum concentrations of 30 µg/mL and trough concentrations of 7.5 µg/mL.

4. Compute loading dose of 25 mg/kg.

5. Compute maintenance dose of 19 mg/kg given at the dosage interval listed in the following chart for the patient's CrCl:

CrCl (mL/min)	DOSAGE INTERVAL (DAYS)
≥120	0.5
100	0.6
80	0.75
60	1.0
40	1.5
30	2.0
20	2.5
10	4.0
5	6.0
0	12.0

4-Literature-based method

WEIGHT	POSTNATAL AGE	
	< 7 DAYS	≥ 7 DAYS
<1.2 kg	15 mg/kg every 24 hours	15 mg/kg every 24 hours
1.2–2 kg	10–15 mg/kg every 12–18 hours	10–15 mg/kg every 8–12 hours
>2 kg	10–15 mg/kg every 8–12 hours	10–15 mg/kg every 6–8 hours

2- Use of vancomycin concentration to change the dose

1- Linear Pharmacokinetics Method

2- Trough only method

Linear Pharmacokinetics Method

Note: it is suitable only if the con. reached the steady state (after 3-5 half-life)

$$D_{\text{new}} = (C_{ss,\text{new}}/C_{ss,\text{old}}) D_{\text{old}}$$

(C_{ss} could be peak or trough but usually trough)

If the new calculated Dose based on C_{ss} trough, then check the peak C_{ss} to make sure that it is being safe and effective, but if it is based on peak C_{ss}, then check the trough C_{ss}.

$$C_{ss,\text{new}} = (D_{\text{new}}/D_{\text{old}}) C_{ss,\text{old}}$$

2- Trough only method

Note: it is suitable only if the con. reached the steady state (after 3-5 half-life)

$$\tau_{\text{new}} = (\text{Old trough } C_{ss} / \text{New trough } C_{ss}) \tau_{\text{old}}$$

Designed by Turath Nabeel, March-2017.