Pharmacist Meeting

March 2018

Weight based dosing (total body weight vs adjusted)

Background:

- 2012 high incidence of supra-therapeutic troughs prompted change to adjusted body weight (ABW) dosing (for Vd and CrCl calculations)
 - Some data suggests clearance may be most closely associated with ABW → Vd increases in obese patients but not to the same extent as Cl (results in higher than expected troughs)

• $K = CL/\uparrow\uparrow V = \downarrow\downarrow k = \uparrow\uparrow Cmin$

Lowered our percentage of troughs exceeding goal range from ~ 30% to < 10%</p>

Implications of using adjusted body weight for LD calculation

- Adjusted body weight used to lessen chance for elevated troughs/acute kidney injury
- Loading dose goal: rapidly achieve therapeutic concentrations in patients with sepsis or complicated infections
- Smaller loading doses may contribute to initial trough being lower than needed
- Isolated cases of low troughs prompted a review ABW for maintenance dose calculation has decreased elevated levels although higher loading doses may be needed to more rapidly attain therapeutic levels

Loading Doses & Attainment of therapeutic troughs

 Retrospective review of obese patients (BMI > 30) with indications necessitating 15-20 trough



Initial troughs

Loading Doses & Attainment of therapeutic troughs

- Average initial trough values (n=61 patients):
 (excluding hemodialysis or pulse dosed patients)
 - Initial level < 15: 12.38 mg/dL (range 6 14.6 mg/L)</p>
 - Initial level < 10 mg/L: 11.5% of patients (n=7)</p>

Significant number of patients with sub-therapeutic initial troughs

> This is not a problem unique to CHI Memorial

 Various published protocols achieved initial therapeutic concentrations in only 25-52% of cases (obese & non-obese patients)

Loading Doses & Attainment of therapeutic troughs

- Average TBW loading dose \rightarrow 16.9 mg/kg, ranging from 5.7 to 29.0 mg/kg.
- Average ABW loading dose → 22.1 mg/kg, ranging from 7.8 to 36.2 mg/kg.
- ➤ Average dose variance for all patients to attain a TBW 25 mg/kg loading dose (no dose cap) → 968 mg per patient.
- > Only 3 (4.9%) out of all 61 patients reached a supra-therapeutic level.

<u>New</u> Loading Dose Process

TOTAL body weight (TBW) to now be used for LOADING DOSE determination

- Maximum of 3 gram load → 25-30 mg/kg loading dose
- Exception: Hemodialysis patients will remain with 2 gram maximum dose

Loading doses exceeding 2 grams

- To be administered using a "divided-load"
- Example: 94 kg patient with sepsis → 2,500 mg dose (26.6 mg/kg) → 1,250 mg IV Q 6 hrs x / 2 doses
- Maintenance dose to start at next scheduled dosing interval timed from the final "divided" load

Divided Loading Doses (any LD > 2 grams – patients > 84 kg)

- Doses > 2 grams → all doses will be either 2,500 mg or 3,000 mg
 - ► 2,500 mg (1,250 mg Q 6 hours x 2)
 - 3,000 mg (1,500 mg Q 6 hours x 2)
- Doses < 2 grams \rightarrow rounded to nearest 250 mg

Vancomycin <u>New</u> Loading Dose Process

Guideline based loading doses for majority of all patients (25-30 mg/kg – TBW)

- Exception: patients > 123 kg will fall short of this range
- Calculator will advise you to administer the first maintenance dose early for these patients (examples on next slides)

Applies to all patients with desired trough 15-20 mg/L (except HD patients)

- Sepsis, meningitis, pneumonia, endocarditis, osteomyelitis
- Acute renal failure patients \rightarrow also to receive TBW loading dose
 - Subsequent doses to be based on levels

Vancomycin Dosing

Calculator Changes (comparison on next slide)

• Loading dose \rightarrow based on TBW for ALL patients

- Automatically rounds all doses to the appropriate dose (obese & non-obese)
- Rounds all LD's exceeding 2 grams to either 2.5 gm or 3 gm doses

• Maintenance dose calculation \rightarrow no change

Still uses adjusted body weight (ABW) for Vd & CrCl

Patients > 123 kg (if loading dose needed)

- 3 gm LD will <u>not</u> be b/w 25-30 mg
- Calculator will advise you to give the 1st maintenance dose early based on half-life
 - < 8 hours → 6 hours post LD; 8-12 hours → 8 hours post LD; 13-24 → 12 hours post LD,
 × 24 hours → based on levels
 </p>

Loading Dose:	3000	mg		LD:	22.2	_mg/kg [·]	TBM
Give as 1500 mg Q6	hrs x 2 dose	es.					
Begin Maintenance	Dose 8 hrs a	after 2nd loa	ding dose.				

Vancomycin Calculator Changes



Vancomycin Dosing Changes

Questions???