

Ceftazidime-Avibactam vs Meropenem-Vaborbactam

Use: Enterobacteriales family of Gram-negative bacteria

- *Escherichia coli*
- *Klebsiella pneumoniae*
- *Enterobacter* species

Mechanism of resistance: production of broad-spectrum beta lactamase that hydrolyze and inactivate beta-lactam antibiotics, including carbapenems → carbapenem-resistant Enterobacteriaceae (CRE)

	Ceftazidime-Avibactam (Avycaz)	Meropenem-Vaborbactam (Vabomere)
Mechanism of action	<u>Ceftazidime</u> : β-lactam, 3 rd generation cephalosporin <u>Avibactam</u> : β-lactamase inhibitor, extends ceftazidime spectrum of activity	<u>Meropenem</u> : β-lactam, carbapenem <u>Vaborbactam</u> : β-lactamase inhibitor, protects meropenem degradation by extended spectrum β-lactamases
FDA approved indication	<ul style="list-style-type: none"> • Complicated intra-abdominal infections (IAI) with metronidazole • Complicated urinary tract infections (UTI) • Hospital-acquired pneumonia/ ventilator-associated pneumonia 	<ul style="list-style-type: none"> • Complicated UTI
Spectrum of activity	<ul style="list-style-type: none"> • CRE 	<ul style="list-style-type: none"> • CRE
Common dosing	<ul style="list-style-type: none"> • 2.5 g (ceftazidime 2g-avibactam 0.5g) IV every 8 hours • Renal dose adjustment for CrCl < 50 ml/min 	<ul style="list-style-type: none"> • 4 g (meropenem 2g-vaborbactam 2g) IV every 8 hours • Renal dose adjustment for CrCl < 50 ml/min
AWP cost per dose	\$430.57	\$396.00
Trial data	<p>Mazuski et al:</p> <ul style="list-style-type: none"> • <u>Patient population</u>: IAI • <u>Intervention</u>: <ul style="list-style-type: none"> ◦ Ceftazidime-avibactam 2-0.5g + metronidazole 500 mg every 8 hours ◦ Meropenem 1g every 8 hours ◦ Placebo every 8 hours • <u>Outcome</u>: clinical cure at 28-35 days with ceftazidime-avibactam + metronidazole (81.6%) was noninferior versus meropenem (85.1%) <p>Carmeli et al:</p> <ul style="list-style-type: none"> • <u>Patient population</u>: complicated UTI or complicated IAI due to ceftazidime-resistant Enterobacteriaceae or <i>P. aeruginosa</i> • <u>Intervention</u>: ceftazidime-avibactam 2-0.5g every 8 hours versus best available therapy (carbapenem used in 97% of best available therapy group) • <u>Outcome</u>: clinical cure at test of cure visit was similar in ceftazidime-avibactam (91%) versus best available (91%) 	<p>TANGO-I:</p> <ul style="list-style-type: none"> • <u>Patient population</u>: complicated UTI • <u>Intervention</u>: <ul style="list-style-type: none"> ◦ Meropenem-vaborbactam 2g-2g IV every 8 hours ◦ Piperacillin-tazobactam 4g-0.5g every 8 hours • <u>Outcome</u>: clinical cure and microbial eradication with meropenem-vaborbactam (98.4%) was noninferior to piperacillin-tazobactam (94%) <p>TANGO-II:</p> <ul style="list-style-type: none"> • <u>Patient population</u>: suspected or confirmed CRE infections • <u>Intervention</u>: meropenem-vaborbactam 2g-2g every 8 hours versus best available therapy (combination therapy with carbapenem) • <u>Outcome</u>: clinical cure rate at end of therapy higher in meropenem-vaborbactam (64.3%) versus best available therapy (33.3%)

References

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