ECMO Antimicrobial and Sedation Dosing Reference

LALK (Revised February 2024)

Extracorporeal Membrane Oxygenation (ECMO) is a life support system for the heart and lungs where blood is removed from the body before being oxygenated and returned to the patient. The ECMO circuit consists of tubing, both from the patient to the machine and within the machine; pumps; and an oxygenator, which oxygenates the blood and removes CO₂.

There are several dosing considerations related to ECMO, including the following:

- Sequestration drug lost to the circuit by adherence to the tubing and/or oxygenator membranes
- Volume of Distribution increased effective drug distribution due to hemodilution or distribution into the circuit itself, which lowers the patient's drug levels
- Clearance altered drug clearance due to the ECMO circuit

Upward arrows indicate an increase in the corresponding pharmacokinetic parameter while downward arrows indicate a decrease.

Patients on ECMO are often also on CRRT, and the <u>CRRT dosing protocol</u> should also be referenced for these patients.

Vd: volume of distribution, Cl: clearance, t_{1/2}: half-life, PK: pharmacokinetics IV: intravenous, PO: oral, CF: cystic fibrosis, FN: febrile neutropenia, PNA: pneumonia, SSTI: skin and soft tissue infection, BSI: bloodstream infection, IE: infective endocarditis, IAI: intra-abdominal infection, CNS: central nervous system, UTI: urinary tract infection, Obesity: TBW/IBW ≥120%, RRT: renal replacement therapy

	Usual dose	ECMO Effects on PK				
Medication		Sequestration	Vd	CI	t _{1/2}	Drug Dosing Changes
Amphotericin B Underweight: TBW Normal weight: TBW Obese: DW **Do not automatically adjust without contacting MD first	Mucormycosis** 5 mg/kg IV q24h	<u>Deoxycholate</u> Minimal	↑	$\uparrow \downarrow$	$\uparrow \downarrow$	Reasonable to start normal dosing, discuss risk/benefit with clinical pharmacy
	Other indications** 3-4 mg/kg IV q24h	<u>Liposomal</u> Moderate				May consider deoxycholate formulation
Cefepime *Highly recommend extended 3 to 4-hour infusions	UTI, SSTI 2 g IV q12h ICU, CNS, FN, CF, PNA, IE, Pseudomonas 2 g IV q8h	Minimal	No change	↓ ↓	↑ ↑	CrCl ≥ 61: 2 g IV q8h CrCl 30-60: 1 g IV q8h CrCl 11-29: 1 g IV q12h CrCl ≤ 10: 1 g IV q24h All given over 3 hours
Cefiderocol Restricted to ID 3-hour infusion time for all doses	2 g IV q8h	Minimal	Unknown	Unknown	Unknown	Insufficient data Reasonable to start normal dosing, discuss risk/benefit with clinical pharmacy
Ceftaroline Restricted to ID	600 mg IV q12h MRSA BSI 600 mg IV q8h	Significant	↑	Unknown	Unknown	Insufficient data Reasonable to start normal dosing, discuss risk/benefit with clinical pharmacy

Medication	Usual dose	ECMO Effects on PK				
		Sequestration	Vd	CI	t _{1/2}	Drug Dosing Changes
Ceftolozane/tazobactam Restricted to ID	IAI, UTI, SSTI, other infections 1.5 g IV q8h Pneumonia 3 g IV q8h	Minimal	1	No change	No change	No adjustment necessary
Ceftriaxone	2 g IV q24h CNS or Enterococcal Endocarditis 2 g IV q12h	Minimal	No change	No change	No change	No adjustment necessary
Ciprofloxacin (IV)	400 mg IV q12h Nosocomial PNA or Intermediate susceptibility 400 mg IV q8h	Minimal	1	↓	↑	No adjustment necessary If MIC > 0.25 mg/L, use alternative agent due to decreased likelihood of reaching a therapeutic level
Fluconazole (PO/IV) Round doses UP to the nearest 200 mg	Vaginal Yeast Infection 200 mg PO/IV x1 Heme/onc prophylaxis 400 mg PO/IV q24h Esophageal candidiasis or symptomatic urinary infection 3 mg/kg PO/IV Invasive; CNS, IAI, BSI 12 mg/kg IV/PO loading dose, then 6 mg/kg IV/PO q24h Susceptible dose- dependent Nakaseomyces (Candida) glabrata 12 mg/kg IV/PO q24h	Minimal	1	No change	No change	No adjustment necessary; dose based on TBW
Gentamicin Underweight: TBW Normal weight IBW Obese: DW	Pharmacist PK dosing by drug levels: see facility's aminoglycoside dosing protocol	Minimal	1	↓	1	Therapeutic drug monitoring recommended

Medication	Usual dose	ECMO Effects on PK					
		Sequestration	Vd	CI	t _{1/2}	Drug Dosing Changes	
Linezolid	600 mg PO/IV q12h	Minimal	No change	No change	No change	No adjustment necessary	
Meropenem Highly recommend extended 3-hour infusions	500 mg IV q6h	Minimal	↑	↑	↓	Limited data, utilize 1-2g extended infusion dosing below	
	1 g IV q8h					No adjustment necessary, administer as an extended infusion over 3 hours	
	Meningitis, CF, or MIC of 4 mcg/mL 2 g IV q8h						
Micafungin Restricted to ID or Critical Care	100 mg IV q24h	Minimal to Moderate	↑	$\uparrow\downarrow$	$\downarrow \uparrow$	150 mg IV q24	
Piperacillin/tazobactam (Extended Infusion – over 4h)	4.5 g IV q8h	Minimal	↑	No change	No change	No adjustment necessary	
Piperacillin/tazobactam (Standard infusion)	4.5 g IV q6h	Minimal	Minimal	^	No change	No change	Use extended infusion
	Pseudomonas ruled out 3.375 g IV q6h		1	ino change	No change	Recommend against 30 minute infusions	
Vancomycin IV Use total body weight for dosing	Pharmacist PK dosing by drug levels	Minimal	↑	No change	No change	No adjustment necessary; see facility's vancomycin dosing protocol	
Voriconazole Underweight: TBW Normal weight: TBW Obese: DW Adjust dose based on therapeutic drug monitoring; goal steady-state (≥7 days) trough of 1-6 mg/L	Treatment 6 mg/kg PO/IV q12h x2 doses, then 4 mg/kg PO/IV q12h	- Significant	↑	$\uparrow \downarrow$	↑↓	Therapeutic drug monitoring strongly recommended – recommend pharmacy consult to assist	
	Prophylaxis 200 mg PO q12h						

Medication	Usual dose	ECMO Effects on PK				
		Sequestration	Vd	CI	t _{1/2}	Drug Dosing Changes
Dexmedetomidine	0.2 – 1.5 mcg/kg/h Titrate: 0.2 mcg/kg/h q30min to target RASS	Significant	Unknown	Unknown	Unknown	0.2 – 1.5 mcg/kg/h Titrate: 0.2 mcg/kg/h q30min to target RASS Not to exceed 1.5 mcg/kg/h due to bradycardia
Fentanyl	25 – 200 mcg/h Titrate: 25 mcg/h q15min to target RASS	Significant	1	Unknown	Unknown	50 – 400 mcg/h Titrate: 25 mcg/h q15min to target RASS
Hydromorphone	0.5 – 4 mg/h Titrate: 0.25 mg/h q60min to target RASS	Minimal	Unknown	Unknown	Unknown	1 – 8 mg/h Titrate: 0.25 mg/h q60min to target RASS
Ketamine	0.05 – 2.5 mg/kg/h Titrate: 0.1 mg/kg/h q15min to target RASS	Minimal to moderate	Unknown	Unknown	Unknown	0.5 – 2.5 mg/kg/h Titrate: 0.1 mg/kg/h q15min to target RASS
Midazolam	0.02 – 0.1 mg/kg/h Titrate: 0.02 mg/kg/h q10min to target RASS	Significant	1	Unknown	Unknown	1 – 10 mg/h Titrate: 1 mg/h q10min to target RASS
Morphine	2 – 30 mg/h Titrate: 0.5 mg/h q30min to target RASS	Minimal to moderate	1	Unknown	Unknown	2 – 30 mg/h Titrate: 0.5 mg/h q30min to target RASS
Propofol	5 – 50 mcg/kg/min Titrate: 5 mcg/kg/min q5min to target RASS	Significant	Unknown	Unknown	Unknown	10 – 50 mcg/kg/min Titrate: 5 mcg/kg/min q5min to target RASS

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