

## Urine Trouble! Renal Dysfunction and Renal Replacement Therapy in ECMO Patients

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## Objectives

- Definition of Acute Kidney Injury (AKI)
- Incidence of AKI and Renal Replacement Therapy (RRT) in patients requiring ECMO
- Pathophysiology of AKI during ECMO
- Indications for RRT in patients requiring ECMO
- Mechanics of RRT during ECMO
- Outcomes in patients that require RRT and ECMO

## Definitions of Acute Kidney Injury

Stage	AKIN criteria <sup>a</sup>		Stage	RIFLE criteria <sup>b</sup>	
	Creatinine	Urine output		Creatinine or GFR	Urine output
1	Increase of >0.3 mg/dL or 0.5-2 times (baseline)	<0.5 mg/kg/hr for >6 hr	Risk	sCr increase $\times 1.5$ (baseline) or GFR decrease >25%	<0.5 mg/kg/hr for >6 hr
2	2-3 times (baseline)	Anuria for 12 hr or <0.5 mg/kg/hr for >12 hr	Injury	sCr increase $\times 2.0$ (baseline) or GFR decrease >50%	<0.5 mg/kg/hr for 12 hr
3	RRT or level of 4.0 mg/dL with an acute increase of 0.5 mg/dL	<0.3 mg/kg/hr for 24 hr	Failure	sCr increase $\times 3.0$ (baseline) or GFR decrease >75% or 4.0 mg/dL with an acute increase of 0.5 mg/dL	<0.5 mg/kg/hr for 12 hr
			Loss	Persistent AKI (loss of renal function >4 weeks)	
			ESKD	End stage kidney disease >3 months	

<sup>a</sup>KDIGO criteria for AKI very similar to AKIN

<sup>b</sup>AKI-D now often in the literature as well = AKI requiring dialysis

## ECMO and Acute Kidney Injury – The Numbers

### The Incidence of Acute Kidney Injury and its Effect on Neonatal and Pediatric ECMO Outcomes: A multicenter report from the KIDMO Study Group

Geoffrey M. Fleming, MD<sup>1</sup>, Rashmi Sahay, PhD<sup>2</sup>, Michael Zappitelli, MD<sup>3</sup>, Eileen King, PhD<sup>2</sup>, David J. Askenazi, MD<sup>4</sup>, Brian C. Bridges, MD<sup>1</sup>, Matthew L. Paden, MD<sup>5</sup>, David T. Selewski, MD<sup>6</sup>, and David S. Cooper, MD<sup>7</sup>

- 832 pediatric patients on ECMO at 6 ECMO centers
- 74% had AKI using KDIGO criteria
- 52% had AKI before initiation of ECMO, 86% by 48hrs after ECMO initiation
- 48% had AKI-D and received RRT (only 3% prior to ECMO)

## ECMO and Acute Kidney Injury – The Numbers

Study	Sample size (N) Exclusions for ECMO	Type of ECMO Number of patients (N)	Number of patients dependent on RRT (%)
Berges et al. (2014)	N = 1471 N = 1484 (AKIN) N = 176 (pediatric) N = 10 (AKI) N = 40 (trauma) N = 40 (postoperative pneumonia) N = 10 (trauma) N = 20 (anemia) N = 120 (miscellaneous)	VV ECMO, N = 761 VA ECMO, N = 207 VV to VA ECMO, N = 50 Other, N = 31	N = 160 (44%)
Tan et al. (2015)	N = 127 N = 41 (CRRT) and/or renal replacement N = 8 (trauma) N = 10 (postoperative) N = 10 (trauma) N = 10 (trauma) N = 10 (trauma) N = 10 (trauma) N = 10 (trauma)	No data on the type of ECMO provided	N = 10 (44%)
Wu et al. (2016)	N = 100 N = 100 (trauma) N = 100 (trauma) N = 100 (trauma) N = 100 (trauma) N = 100 (trauma) N = 100 (trauma) N = 100 (trauma)	Only VA ECMO	N = 10 (44%)
Lee et al. (2016)	N = 100 N = 100 (trauma) N = 100 (trauma) N = 100 (trauma) N = 100 (trauma) N = 100 (trauma) N = 100 (trauma) N = 100 (trauma)	No data on the type of ECMO provided	N = 10 (44%)
Lee et al. (2016)	N = 100 N = 100 (trauma) N = 100 (trauma) N = 100 (trauma) N = 100 (trauma) N = 100 (trauma) N = 100 (trauma) N = 100 (trauma)	VA ECMO, N = 107 VV ECMO, N = 48	N = 70 (44%)
Klein et al. (2016)	N = 100 N = 100 (trauma) N = 100 (trauma) N = 100 (trauma) N = 100 (trauma) N = 100 (trauma) N = 100 (trauma) N = 100 (trauma)	VA ECMO, N = 101 VV ECMO, N = 51 VA ECMO, N = 101	N = 10 (44%) N = 10 (44%) N = 10 (44%)

- Incidence of RRT during ECMO is around 50% for adults as well

### Abbott 2016-2018

- ECMO = 249 patients
- Survival after ECMO = 142 patients (57.0%)
- CRRT = 85 patients (31.8%)
- Survival CRRT/ECMO = 30 patients (35.3%)

## ECMO and Acute Kidney Injury – Why?

### Non-ECMO Factors

- All the usual suspects! Hemodynamics, hypo-perfusion, cardiorenal, SIRS/sepsis/cytokines, IV contrast, antibiotics, etc.

### ECMO Factors

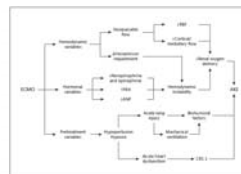


Fig. 1. Major hemodynamic factors related to the development of AKI in patients treated with ECMO (2016-2018).

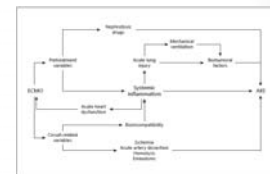


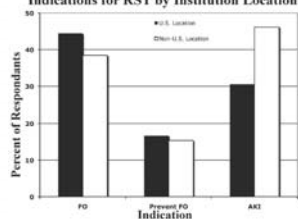
Fig. 2. Major inflammatory factors related to the development of AKI in patients treated with ECMO.

- Bottom line: under-oxygenated/perfused + overworked + poisoned

## Indications for RRT in ECMO patients

- Same indications as everyone else
- But with different proportions for the indication to initiate RRT

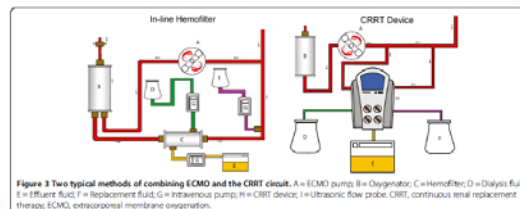
Indications for RRT by Institution Location



- Indications:
  - 59% fluid overload
  - 35% AKI
  - 4% electrolyte imbalance
  - 2% other
- My experience
  - Often more than one indication
  - Fluid overload is usually the primary indication

## Mechanics of RRT in ECMO

- We rarely use a dialysis catheter during ECMO



## Outcomes in Pediatric Patients Requiring ECMO/RRT

### Recovery of renal function and survival after continuous renal replacement therapy during extracorporeal membrane oxygenation

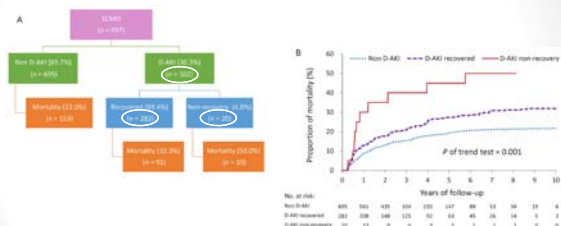
Matthew L. Paden, MD, FAAP<sup>1</sup>, Barry L. Warshaw, MD<sup>2</sup>, Michael L. Heard, RN<sup>3</sup>, and James D. Fortenberry, MD, FAAP, FCCM<sup>4</sup>

- 154 **pediatric** patients requiring RRT during ECMO
- 44 survived to discharge (28.6%) vs. 81/244 non-RRT patients (33.2%)
- 18 required RRT after ECMO was discontinued
- Renal recovery occurred in 96% before discharge
- Only one patient without pre-existing kidney disease did not recover renal function

## Outcomes in Adult Patients Requiring ECMO/RRT

- Three largest studies of adults
  - Germany tertiary care center, 200 ECMO patients from 2005-2010
    - 60% required RRT
    - 3 month survival 53% in ECMO vs. 17% in ECMO/RRT
  - University of AZ, 40 ECMO + RRT patients from 2007-2012
    - 1 month survival 20%
    - 5 survivors came off RRT, 3 remained on RRT after 30 days
  - Tawain National Database, 3,251 ECMO patients from 2003-2013
    - 54% required RRT
    - Survival to hospital discharge 26.2% in RRT/ECMO vs. 57.8% in ECMO
    - Long term survival in patients surviving at least 90 days after hospital discharge is on next slide

## Outcomes in Adult Patients Requiring ECMO/RRT



## Summary

- AKI is common in patients requiring ECMO with about half requiring RRT
- AKI is a marker of disease severity, but is also an independent predictor of mortality
- The mortality is high for patients requiring ECMO and CRRT, but survival to hospital discharge is >20% in most cohorts
- Most patients that require RRT during ECMO will not be dependent on RRT long term

## References

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