

# A Risk Stratification Guide for Bridging Patients on ECMO to Heart Transplantation

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# BACKGROUND

- VA-ECMO is a critical therapy in the management of patients in refractory cardiogenic sock, and is often employed as a bridge to recovery, decisionmaking, durable VAD implantation, or heart transplantation (HT).
- The 2018 heart allocation policy resulted in a rapid increase in the use of VA-ECMO as a bridge to transplant (ECMO-BTT).<sup>1</sup>
- Patients bridged to HT with VA-ECMO face significantly elevated risk of mortality and have been shown to have worse post-HT outcomes.<sup>2</sup>
- Specific risk factors for post-transplant mortality in patients bridged to HT from VA-ECMO remain inadequately characterized.
- Therefore, this study sought to identify predictors of post-transplant mortality among ECMO-BTT patients to develop a risk stratification framework to inform clinical decision-making.

# **METHOD**

- Data Source
  - UNOS Registry
  - Included: all adult ECMO-BTT patients between January 2000 and June 2024
  - Excluded: patients <18, multiorgan transplants
- Outcomes of interest
  - Primary Outcome: 1-year mortality
  - Secondary Outcomes: post-transplant dialysis, stroke, pacemaker implant, acute rejection, length of stay
- Determination of Risk Factors
  - Univariable and multivariable cox regression models were performed using baseline cohort characteristics to determine potential risk factors for 1-year mortality.
  - Significant variables from the multivariable model were then re-analyzed in the same multivariable cox regression model to determine final hazard ratios associated with each variable.
- Risk Group Stratification

The cohort was then stratified by the presence of 0 risk factors, 1 risk factor, 2 risk factors, and  $\geq$  3 risk factors.

Kaplan-Meier analysis was then performed to determine 1-year mortality by risk factor group.

# RESULTS

**Table 1. Recipient risk factors for 1-year mortality.** Significant risk factors included age > 61, ventilator-dependence, total bilirubin > 3.1, ICM, and "other etiologies" of HF.

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Univariable	P-value	Multivariable	P-value	Final Model	P-value
1.12 (1.02-1.22)	0.012	1.13 (1.00-1.27)	0.042	1.16 (1.04-1.30)	0.01
1.30 (1.04-1.62)	0.02	1.21 (0.91-1.62)	0.184		
1.47 (1.11-1.94)	0.007	1.16 (0.82-1.65)	0.4		
2.15 (1.67-2.76)	<0.001	1.84 (1.33-2.53)	<0.001	2.21 (1.63-2.99)	<0.001
1.39 (1.08-1.80)	0.011	1.16 (0.85-1.60)	0.356		
2.83 (2.11-3.79)	<0.001	1.38 (0.75-2.57)	0.31		
1.33 (1.24-1.43)	<0.001	1.17 (0.97-1.40)	0.099		
1.08 (1.05-1.11 )	<0.001	1.07 (1.04-1.11)	<0.001	1.09 (1.05-1.12)	<0.001
Reference		Reference		Reference	
1.54 (1.13-2.08)	0.005	1.62 (1.10-2.39)	0.015	1.64 (1.12-2.38)	0.01
1.42 (0.72-2.81)	0.313	1.66 (0.76-3.63)	0.204	1.85 (0.87-3.99)	0.111
0.66 (0.31-1.42)	0.287	0.85 (0.34-2.13)	0.731	0.77 (0.31-1.92)	0.576
2.68 (1.95-3.70)	<0.001	2.16 (1.43-3.27)	<0.001	2.65 (1.80-3.88)	<0.001
	1.12 (1.02-1.22) 1.30 (1.04-1.62) 1.47 (1.11-1.94) 2.15 (1.67-2.76) 1.39 (1.08-1.80) 2.83 (2.11-3.79) 1.33 (1.24-1.43) 1.08 (1.05-1.11)  Reference 1.54 (1.13-2.08) 1.42 (0.72-2.81) 0.66 (0.31-1.42)	1.12 (1.02-1.22) 0.012 1.30 (1.04-1.62) 0.02 1.47 (1.11-1.94) 0.007 2.15 (1.67-2.76) <0.001 1.39 (1.08-1.80) 0.011 2.83 (2.11-3.79) <0.001 1.33 (1.24-1.43) <0.001 1.08 (1.05-1.11) <0.001  Reference 1.54 (1.13-2.08) 0.005 1.42 (0.72-2.81) 0.313 0.66 (0.31-1.42) 0.287	1.12 (1.02-1.22)       0.012       1.13 (1.00-1.27)         1.30 (1.04-1.62)       0.02       1.21 (0.91-1.62)         1.47 (1.11-1.94)       0.007       1.16 (0.82-1.65)         2.15 (1.67-2.76)       <0.001       1.84 (1.33-2.53)         1.39 (1.08-1.80)       0.011       1.16 (0.85-1.60)         2.83 (2.11-3.79)       <0.001       1.38 (0.75-2.57)         1.33 (1.24-1.43)       <0.001       1.17 (0.97-1.40)         1.08 (1.05-1.11)       <0.001       1.07 (1.04-1.11)         Reference         1.54 (1.13-2.08)       0.005       1.62 (1.10-2.39)         1.42 (0.72-2.81)       0.313       1.66 (0.76-3.63)         0.66 (0.31-1.42)       0.287       0.85 (0.34-2.13)	1.12 (1.02-1.22)       0.012       1.13 (1.00-1.27)       0.042         1.30 (1.04-1.62)       0.02       1.21 (0.91-1.62)       0.184         1.47 (1.11-1.94)       0.007       1.16 (0.82-1.65)       0.4         2.15 (1.67-2.76)       <0.001       1.84 (1.33-2.53)       <0.001         1.39 (1.08-1.80)       0.011       1.16 (0.85-1.60)       0.356         2.83 (2.11-3.79)       <0.001       1.38 (0.75-2.57)       0.31         1.33 (1.24-1.43)       <0.001       1.17 (0.97-1.40)       0.099         1.08 (1.05-1.11)       <0.001       1.07 (1.04-1.11)       <0.001         Reference         1.54 (1.13-2.08)       0.005       1.62 (1.10-2.39)       0.015         1.42 (0.72-2.81)       0.313       1.66 (0.76-3.63)       0.204         0.66 (0.31-1.42)       0.287       0.85 (0.34-2.13)       0.731	1.12 (1.02-1.22)       0.012       1.13 (1.00-1.27)       0.042       1.16 (1.04-1.30)         1.30 (1.04-1.62)       0.02       1.21 (0.91-1.62)       0.184         1.47 (1.11-1.94)       0.007       1.16 (0.82-1.65)       0.4         2.15 (1.67-2.76)       <0.001       1.84 (1.33-2.53)       <0.001       2.21 (1.63-2.99)         1.39 (1.08-1.80)       0.011       1.16 (0.85-1.60)       0.356         2.83 (2.11-3.79)       <0.001       1.38 (0.75-2.57)       0.31         1.33 (1.24-1.43)       <0.001       1.17 (0.97-1.40)       0.099         1.08 (1.05-1.11)       <0.001       1.07 (1.04-1.11)       <0.001       1.09 (1.05-1.12)         Reference         1.54 (1.13-2.08)       0.005       1.62 (1.10-2.39)       0.015       1.64 (1.12-2.38)         1.42 (0.72-2.81)       0.313       1.66 (0.76-3.63)       0.204       1.85 (0.87-3.99)         0.66 (0.31-1.42)       0.287       0.85 (0.34-2.13)       0.731       0.77 (0.31-1.92)

Table 2. Donor risk factors for 1-year mortality. Significant risk factors included female gender and ischemic times  $\geq$ 4.47 hours.

	Variable	Univariable	P-value	Multivariable	P-value	Final Model	P-value
	Gender						
	Male	Reference		Reference		reference	
	Female	1.60 (1.22-2.11)	0.001	1.59 (1.11-2.27)	0.011	1.61 (1.16-2.23)	0.004
	Mechanism of Death						
	Trauma	reference		Reference			
	Drug overdose	0.71 (0.49-1.01)	0.055	0.712 (0.46-1.11)	0.139		
	Stroke	1.13 (0.79-1.64)	0.493	0.88 (0.56-1.39)	0.583		
J	Anoxia	1.14 (0.711.83)	0.586	1.33 (0.79-2.25)	0.288		
	Cardiac	0.74 (0.45-1.19)	0.213	0.81 (0.47-1.38)	0.43		
	Other	1.00 (0.37-2.71)	0.995	0.91 (0.28-2.93)	0.875		
_	ВМІ	1.03 (0.83-1.28)	0.796				
	Ischemia Time	1.08 (0.96-1.22)	0.176	1.15 (1.00-1.31)	0.049	1.14 (1.00-1.30)	0.048

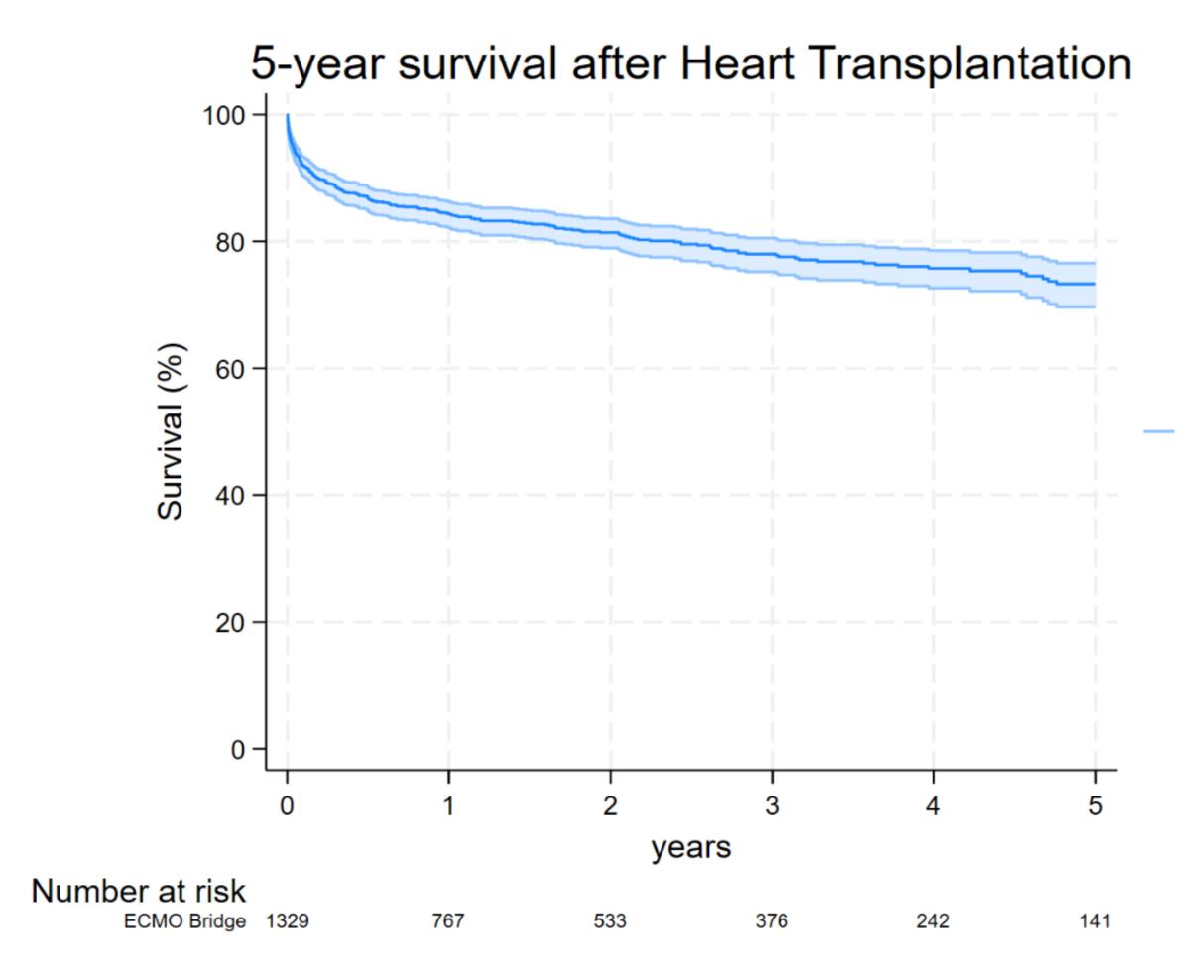


Figure 1. Overall post-transplant survival. Unadjusted survival for all ECMO-BTT patients at 30 days, 90 days, 1-year, and 5-years was 92.73%, 89.26%, 84.33%, and 73.30%, respectively.

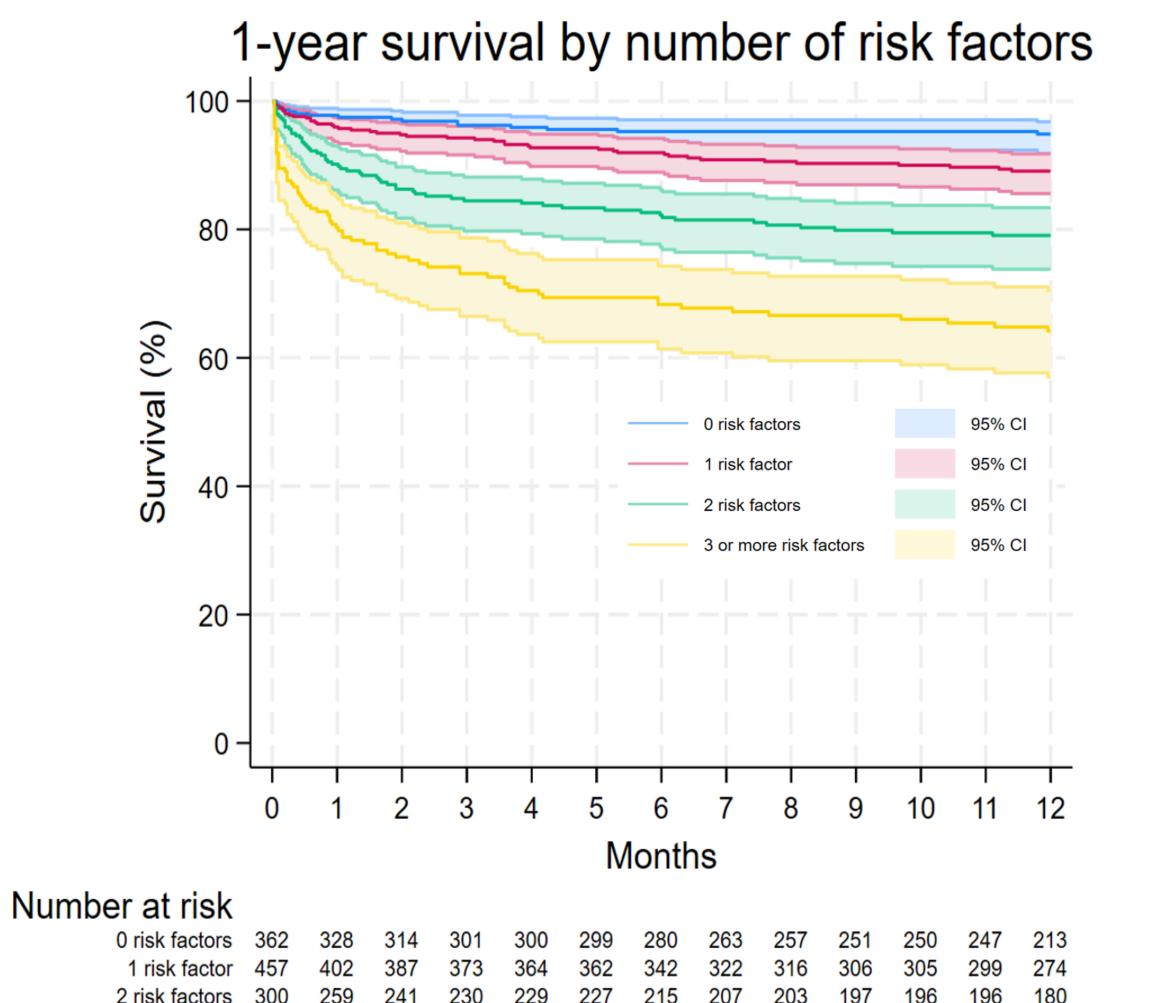


Figure 2. Post-transplant mortality by number of risk factors. 1-year post-HT survival for patients with 0, 1, 2, or ≥3 risk factors was 94.86%, 89.09%, 79.07%, and 64.17%, respectively. The hazard ratio for 1-year mortality for patients with 1, 2, or ≥3 risk factors was 2.09 (1.2-3.6), 4.39 (2.6-7.5), and 8.48 (5.0-14.4), respectively.

# CONCLUSIONS

- Patients bridged to transplant with VA-ECMO represent a complex patient population with elevated post-transplant mortality.
- We identified the following risk factors for 1-year mortality in ECMO-BTT patients:
  - Age ≥ 61 years
  - Ventilator dependence prior to transplant
  - Bilirubin ≥ 3.1
  - Ischemic and unspecified/mixed cardiomyopathies
  - Donor organ ischemic time ≥ 4.47 hours
  - Female donors
- Patients with 2 or more of the above risk factors should proceed cautiously as this is associated with less than 80% 1-year survival.

### REFERENCES

- 1. DeFilippis EM, Clerkin K, Truby LK, et al. ECMO as a Bridge to Left Ventricular Assist Device or Heart Transplantation. *JACC Heart Fail.* 2021;9(4):281-289.
- 2. Moonsamy P, Axtell AL, Ibrahim NE, et al. Survival After Heart Transplantation in Patients Bridged With Mechanical Circulatory Support. *J Am Coll Cardiol*. 2020;75(23):2892-2905