

Blood-based versus crystalloid cardiopulmonary bypass priming in left ventricular assist device surgery: impact on end-organ function and survival

Sophie Bonni ¹, Rashad Zayat ¹, Natasja W. M. Ramnath ¹, Nima Hatam ¹, Lachmandath Tewarie ¹, Ajay Moza ¹, Marjolijn C. Sales ¹

¹Department of Cardiac Surgery, University Hospital RWTH Aachen, Aachen, Germany

Corresponding author: Sophie Bonni, Department of Cardiac Surgery, University Hospital RWTH Aachen, Pauwelsstraße 30, 52074 Aachen, Germany, +492418036780, sbonni@ukaachen.de.

Part of this manuscript was presented at the DGTHG Annual Meeting (German Society for Cardiothoracic and Vascular Surgery) in Cologne on February 21, 2026.

Word count 4500.

Clinical registration number: Ethics Committee of the Medical Faculty of RWTH Aachen (EK 25-154).

ABSTRACT

Objectives

Cardiopulmonary bypass (CPB) priming may influence hemodilution, bleeding and end-organ injury during HeartMate 3 (HM3) implantation, but evidence in contemporary left ventricular assist device (LVAD) surgery is limited.

Methods

We retrospectively analysed 92 HM3 implantations (2015–2025) by priming strategy (blood-based [FFP+RBC], n=36; crystalloid, n=56). Primary endpoint was postoperative right heart failure (RHF; INTERMACS). Multivariable regression and propensity score-based inverse probability of treatment weighting (IPTW) sensitivity analyses were performed.

Results

Baseline characteristics were balanced between groups, with comparable age (blood-based: 58 [49.5–66.0]; crystalloid: 62 [55.8–68.2]; p=0.069) and female sex (blood-based: n=3/36 [8.3%]; crystalloid: n=8/56 [14.3%]; p=0.518). RHF was lower with blood-based priming (blood-based: n=3/36 [8.3%]; crystalloid: n=17/56 [30.4%]; p=0.012; adjusted OR 0.23 [95% CI 0.06–0.86]; p=0.030). Dialysis was lower unadjusted (blood-based: n=4/36 [11.1%]; crystalloid: n=18/56 [32.1%]; p=0.021), with a directionally lower adjusted estimate (OR 0.30 [95% CI 0.09–1.01]; p=0.053). Delirium was lower with blood-based priming (blood-based: n=3/36 [8.3%]; crystalloid: n=16/56 [28.6%]; p=0.019; adjusted OR 0.22 [95% CI 0.06–0.84]; p=0.027). Survival did not differ significantly (log-rank p=0.055; adjusted HR 0.59 [95% CI 0.27–1.32]; p=0.201). IPTW analyses showed lower delirium (OR 0.23; [95% CI 0.06–0.84]) and attenuated RHF association (OR 0.35; [95% CI 0.10–1.23]).

Conclusions

Blood-based CPB priming was associated with lower RHF and delirium, and directionally lower dialysis rates, after HM3 implantation. Survival estimates were not statistically significant. Findings require prospective validation.

Keywords blood-based priming; fresh frozen plasma; red blood cells; cardiopulmonary bypass; HeartMate 3; left ventricular assist device.

INTRODUCTION

Left ventricular assist device (LVAD) implantation is an established therapy in end-stage heart failure (1, 2). Despite improved technologies and perioperative management, complications, including bleeding, thromboembolic events, infections, and end-organ injury, remain high (3-5).

Acute right heart failure (RHF) affects 20–40% of patients after LVAD implantation (6, 7). RHF post-LVAD surgery involves multifactorial mechanisms (8), with right ventricular function being highly sensitive to loading changes (9).

Cardiopulmonary bypass (CPB) priming affects physicochemical and homeostasis balance (10, 11). While crystalloids cause dilution, blood-based priming with fresh frozen plasma (FFP) and red blood cells (RBC) aims to preserve oncotic pressure, reduce hemodilution and may decrease bleeding and postoperative organ dysfunction. However, evidence supporting blood-based priming in contemporary LVAD surgery remains limited (12, 13).

We hypothesized that blood-based CPB priming [FFP+RBC] is associated with lower rates of clinically significant postoperative RHF and end-organ dysfunction compared with crystalloid priming. Early postoperative outcomes were defined as in-hospital complications and early mortality including 30-day mortality.

MATERIAL AND METHODS

Study design

We performed a single-center retrospective observational study of 92 adult patients who underwent LVAD implantation (HeartMate 3, Abbott Laboratories, Pleasanton,

CA, USA) from October 2015 to April 2025. Clinical, laboratory, echocardiographic, and hemodynamic data were collected from electronic and paper records into an electronic case report file. Patients were retrospectively categorized according to the CPB priming strategy used at surgery (blood-based [FFP + RBC] versus crystalloid); no assignment to treatment groups occurred. Outcomes and adverse events were defined in accordance with the Interagency Registry for Mechanically Assisted Circulatory Support (INTERMACS) definitions (14).

The primary endpoint was postoperative RHF, defined as elevated central venous pressure >16 mmHg or echocardiographic/clinical equivalents and manifestations of venous congestion. RHF severity was further classified by the need for temporary right ventricular mechanical support or requirement for continuous inotropic support extending beyond 14 days postoperatively. Secondary endpoints included renal failure (new onset dialysis), delirium (clinical diagnosis by positive Confusion Assessment Method for the Intensive Care Unit score), major bleeding (any bleeding event requiring transfusion of ≥ 2 units of RBC involving a critical organ or necessitating surgical re-exploration), transfusions, and survival (time from LVAD implantation to death or last follow-up)

Ethics

Only routinely collected clinical data were used; no biological material, research biobank, or database for indefinite future use was established. Therefore, the WMA Declaration of Taipei did not impose additional requirements. The Ethics Committee of RWTH Aachen University approved the study (EK 25-154, May 15, 2025) and waived informed consent.

CPB, surgery and postoperative management

Priming strategy (blood-based versus crystalloid) was determined by surgeon preference. Crystalloid priming comprised balanced crystalloid (1150 mL), mannitol (100 mL), and 5000 IU unfractionated heparin; blood-based priming comprised four FFP units, two RBC units, and 5000 IU heparin. During CPB, unfractionated heparin (500 IU/kg) targeted ACT >500 seconds and was reversed 1:1 with protamine. CPB discontinuation required absence of relevant surgical bleeding. Cell-saver use was non-mandatory and reserved for high preoperative bleeding risk after interdisciplinary discussion. ROTEM-guided transfusion and vasoactive management followed institutional and guideline-based protocols (15).

Statistical analysis

Analyses were performed in R (v4.2.3). Continuous variables are shown as median (IQR) and compared using Wilcoxon–Mann–Whitney tests; categorical variables as n (%) and compared using χ^2 or Fisher's exact tests. We report RR/OR with 95% CI. Logistic models were unadjusted and adjusted for age, sex, and INTERMACS profile. Survival was assessed by Kaplan–Meier/log-rank and Cox models. We performed propensity score–based inverse probability of treatment weighting (IPTW) sensitivity analysis using stabilized weights truncated at the 1st/99th percentiles and robust standard errors; covariate balance was assessed by standardized mean differences (SMD) (target |SMD|<0.10, acceptable <0.15). Covariates, balance, and weighted estimates are reported in Supplementary Tables S1A–C; the STROBE flow diagram is shown in Supplementary Figure S1.

RESULTS

Baseline preoperative characteristics

Blood-based priming was used in 36 patients [39.1%] and crystalloid priming in 56 [60.9%]. Most baseline characteristics were comparable between both groups (Table 1). Age (blood-based: 58 [49.5–66.0] years; crystalloid: 62 [55.8–68.2] years; $p=0.069$), female sex (blood-based: $n=3/36$ [8.3%]; crystalloid: $n=8/56$ [14.3%]; $p=0.518$), and INTERMACS profile distribution (blood-based: [25,0%]; crystalloid: [25,0%]; $p=0.509$) did not differ significantly. Notably, prior PCI was more frequent in the crystalloid group (blood-based: $n=8/36$ [22.2%]; crystalloid: $n=32/56$ [57.1%]; $p<0.001$).

Risk scores, comorbidities, diagnostic findings, and key preoperative laboratory parameters are summarized in Table 1, with extended panels in Supplementary Table S2. Missingness is summarized in Supplementary Table S3.

Intraoperative course

CPB duration, transfusions, catecholamines, and inhaled pulmonary vasodilator use were similar between the groups (Table 2). Hemoglobin (g/dL) at ICU admission was higher in the blood-based priming group (blood-based: 10.3 [9.2–10.8]; crystalloid: 9.2 [8.9–10.0]; $p=0.002$), whereas hemoglobin at 24 hours (blood-based: 9.5 [9.1–10.1]; crystalloid: 9.3 [8.8–9.8]; $p=0.070$) was similar between groups (Table 2). Fibrinogen administration was comparable; however, prothrombin complex concentrate (IU) use was higher in the crystalloid priming group (blood-based: 1500 [0–2000]; crystalloid: 2000 [0–3000]; $p=0.049$) (Table 2).

Postoperative outcomes

Postoperative outcomes are summarized in Table 3. The incidences of RHF (blood-based: $n=3/36$ [8.3%]; crystalloid: $n=17/56$ [30.4%]; $p=0.012$), dialysis (blood-based: $n=4/36$ [11.1%]; crystalloid: $n=18/56$ [32.1%]; $p=0.021$), and delirium (blood-based:

163 n=3/36 [8.3%]; crystalloid: n=16/56 [28.6%]; p=0.019) were lower after blood-based
164 priming.

165 Major bleeding was numerically lower after blood-based priming (blood-based:
166 n=10/36 [27.8%]; crystalloid: n=26/56 [46.4%]; p=0.074). Chest drain output at 24
167 hours and postoperative transfusion requirements within 24 hours (excluding fixed
168 priming units) were comparable between groups (Table 3). Need for liver
169 replacement therapy was numerically lower after blood-based priming (blood-based:
170 n=2/36 [5.6%]; crystalloid: n=11/56 [19.6%]; p=0.058).

171 Duration of mechanical ventilation (h) was similar between both groups (blood-
172 based: 18 [2–40]; crystalloid: 20 [1–39]; p=0.953).

173 In multivariable logistic regression adjusted for age, sex, and INTERMACS profile
174 (Table 4), blood-based priming was associated with lower odds of RHF (OR 0.23
175 [95% CI 0.06–0.86]; p=0.030) and delirium (OR 0.22 [95% CI 0.06–0.84]; p=0.027).
176 Dialysis was directionally lower but imprecise (OR 0.30 [95% CI 0.09–1.01];
177 p=0.053). For major bleeding, the results showed (OR 0.47 [95% CI 0.19–1.18];
178 p=0.109).

179 **Survival**

180 Survival analysis (median follow-up 24 months) showed no statistically significant
181 difference between groups (Figure 1) with a log-rank p=0.055. In multivariable Cox
182 regression adjusted for age, sex, and INTERMACS profile (Table 5), blood-based
183 priming was associated with a non-significant lower hazard of mortality during follow-
184 up (HR 0.59 [95% CI 0.27–1.32]; p=0.201). Age remained associated with mortality
185 (HR 1.05 per year [95% CI 1.01–1.10]; p=0.007), and a higher INTERMACS profile
186 was protective (HR 0.52 per 1-level increase [95% CI 0.38–0.70]; p<0.001). In

propensity-weighted sensitivity analyses, effect estimates were directionally consistent for postoperative endpoints (Supplementary Table S1B) and survival (Supplementary Table S1C), although precision remained limited.

DISCUSSION

In this study, blood-based CPB priming during LVAD implantation was associated with lower incidence of postoperative RHF and delirium, and with directionally lower dialysis rates, compared with crystalloid priming. Baseline characteristics were mostly balanced between the groups, although the non-randomized priming strategy precluded exclusion of residual confounding.

The main finding was the association between blood-based CPB priming and lower postoperative RHF. RHF is a major post-LVAD complication (16). In our cohort, RHF was threefold higher with crystalloid priming (blood-based: $n=3/36$ [8.3%]; crystalloid: $n=17/56$ [30.4%]; $p=0.012$; adjusted OR 0.23 [95% CI 0.06–0.86]; $p=0.030$). In IPTW analysis, this association attenuated (OR 0.35 [95% CI 0.10–1.23]), consistent with limited precision and potential residual confounding, while the direction remained concordant. The right ventricle is load-dependent and vulnerable to postoperative geometric changes (8, 9, 17, 18). Crystalloid priming and transfusions may cause hemodilution and edema, impairing RV contractility and increasing RHF risk (19, 20). By preserving oncotic pressure, FFP may limit interstitial fluid accumulation and help explain lower RHF rates (21, 22).

The lower unadjusted rate of postoperative dialysis in the blood-based priming group may suggest renal protection. Acute kidney injury requiring dialysis after LVAD implantation worsens outcomes (23). In our cohort, dialysis occurred more frequently

211 in the crystalloid group (blood-based: n=4/36 [11.1%]; crystalloid: n=18/56 [32.1%];
212 p=0.021), although the adjusted estimate was imprecise (OR 0.30 [95% CI 0.09–
213 1.01]; p=0.053). Blood-based priming may maintain intravascular volume better,
214 supporting renal perfusion during CPB (24). In addition, FFP provides coagulation
215 factors and other plasma proteins that may attenuate inflammatory and
216 coagulopathic cascades, potentially limiting kidney injury (25, 26).

217 Furthermore, our data showed that despite similar overall transfusion rates and
218 comparable 24-hour hemoglobin levels, blood-based priming maintained higher
219 hemoglobin levels at ICU admission. This suggests that prevention of marked
220 intraoperative hemodilution may contribute to lower volume-overload complications
221 such as RHF and dialysis.

222 Delirium was also lower with blood-based priming (blood-based: n=3/36 [8.3%];
223 crystalloid: n=16/56 [28.6%]; p=0.019; adjusted OR 0.22 [95% CI 0.06–0.84];
224 p=0.027). Post-cardiac surgery delirium reflects cerebral hypoperfusion,
225 inflammation, microemboli, and metabolic disturbances; risk factors include
226 prolonged CPB, transfusion, age, and cognitive impairment (27, 28). The association
227 may reflect improved hemodynamic stability, microcirculatory flow, and less systemic
228 inflammation (29, 30). Without neurologic monitoring or biomarker data, causality
229 cannot be inferred, but concordant signals across RHF, renal dysfunction, and
230 delirium support a potential end-organ perfusion benefit (26, 28, 29).

231 Major bleeding commonly complicates complex cardiac surgery (15). FFP has been
232 proposed as prophylactic priming to reduce bleeding. Bianchi et al. showed that FFP
233 priming in pediatric cardiac surgery reduces bleeding and increases fibrinogen levels
234 (31). Other studies showed improved thromboelastometric parameters and reduced

transfusions (12). However, some randomized studies and meta-analyses reported inconsistent effects of blood-based priming on postoperative bleeding and transfusions (13, 32). In our cohort, bleeding events were numerically fewer in the blood-based priming group (blood-based: n=10/36 [27.8%]; crystalloid: n=26/56 [46.4%]; p=0.074; adjusted OR 0.47 [95% CI 0.19–1.18]; p=0.109).

Survival estimates favored blood-based priming but were not statistically significant (log-rank p=0.055; adjusted HR 0.59 [95% CI 0.27–1.32]; p=0.201). Given the wide CI and few deaths, these data should not be interpreted as evidence of survival benefit.

This study is among the first evaluating the association between blood-based priming and early postoperative outcomes in HM3 implantation, extending evidence from pediatric and non-LVAD cardiac surgery. Multicenter and randomized studies are needed to validate results.

Limitations

The retrospective single-centre design precludes causal inference, and clinician-selected priming introduces potential residual confounding. The small sample size, especially in the blood-based group, limited power for infrequent outcomes such as major bleeding and mortality. Few deaths and wide confidence intervals further limit mortality inference; survival findings should be considered exploratory. Laboratory markers of hemodilution, endothelial injury, and inflammation were not systematically analyzed. Findings may not generalize to centers with different perfusion or perioperative protocols.

CONCLUSION

Our study suggests that blood-based CPB priming is associated with lower rates of postoperative RHF and delirium, and with directionally lower dialysis rates, in HM3 implantation. Survival estimates favored blood-based priming but were not statistically significant. These findings are hypothesis-generating and should be confirmed in prospective multicentre studies, ideally a randomized controlled trial.

ACKNOWLEDGMENTS

Not applicable.

FUNDING

No external funding.

CONFLICTS OF INTEREST

No conflicts of interest.

DATA AVAILABILITY

The data underlying this article will be shared on reasonable request to the corresponding author.

AUTHOR CONTRIBUTIONS

280 S.B. conceived the study, collected data, performed analyses and drafted the
281 manuscript. R.Z. contributed to analyses and interpretation. N.R., N.H., L.T., A.M.
282 and M.S. contributed to study design/interpretation and critically revised the
283 manuscript.

284 N.H. additionally collected the missing clinical data, performed the updated statistical
285 analysis, and generated the new supplementary tables. All authors approved the
286 final version.

287 **FIGURE LEGENDS**

288 Graphical Abstract. The scheme outlines the impact on end-organ function and
289 survival of blood-based priming (fresh frozen plasma and red blood cells, n=36)
290 versus crystalloid priming (n=56) in a retrospective study of 92 patients who
291 underwent LVAD surgery. Blood-based priming was associated with lower rates of
292 postoperative right heart failure and delirium, and with directionally lower dialysis
293 rates. Survival estimates favored blood-based priming but were not statistically
294 significant.

295 Alt-text: Graphical abstract comparing blood-based versus crystalloid CPB priming in
296 92 LVAD patients, highlighting lower right heart failure, delirium and directionally
297 lower dialysis rates after blood-based priming, with a non-significant survival
298 estimate.

299 Figure 1. Kaplan–Meier curves showing postoperative survival in HM3 patients with
300 blood-based or crystalloid priming.

301 Alt-text: Kaplan–Meier survival curves comparing blood-based and crystalloid CPB
302 priming after HM3 implantation, showing numerically higher survival in the blood-
303 based group without statistical significance.

304

Table 1. Baseline preoperative characteristics.

Variable	Crystalloid priming (n=56)	Blood-based priming (n=36)	P-value
Demographics			
Age, years	62.0 (55.8–68.2)	58.0 (49.5–66.0)	0.069
Female sex, n (%)	8 (14.3%)	3 (8.3%)	0.518
Body mass index, kg/m ²	26.8 (24.1–29.7)	27.1 (24.7–30.0)	0.640
Body surface area, m ²	2.02 (1.93–2.16)	2.04 (1.97–2.17)	0.614
Heart failure severity and operative risk			
Left ventricular ejection fraction, %	20 (15–25)	20 (15–25)	0.700
NYHA class III, n (%)	24 (42.9%)	14 (38.9%)	0.706
NYHA class IV, n (%)	30 (53.6%)	19 (52.8%)	0.941
LVAD implantation strategy, n (%)			0.052
Bridge to transplant	18 (32.1%)	16 (44.4%)	
Destination therapy	21 (37.5%)	12 (33.3%)	
Bridge to candidacy	16 (28.6%)	3 (8.3%)	
Bridge to recovery	1 (1.8%)	2 (5.6%)	
Bridge to destination	0 (0.0%)	2 (5.6%)	
Unknown	0 (0.0%)	1 (2.8%)	
INTERMACS profile, n (%)			0.509
1	10 (17.9%)	5 (13.9%)	
2	8 (14.3%)	2 (5.6%)	
3	17 (30.4%)	14 (38.9%)	
4	21 (37.5%)	15 (41.7%)	
EuroSCORE II, %	5.16 (3.01–15.29)	4.12 (3.34–8.49)	0.592
EUROMACS score	2.0 (0.0–4.8)	2.0 (0.0–2.0)	0.200
Comorbidities and preoperative support			
Coronary artery disease, n (%)	7 (12.5%)	4 (11.1%)	1
Peripheral arterial disease, n (%)	4 (7.1%)	3 (8.3%)	1
Insulin dependent diabetes mellitus, n (%)	6 (10.7%)	4 (11.1%)	1
Non-insulin dependent diabetes mellitus, n (%)	16 (28.6%)	7 (19.4%)	0.324
Chronic obstructive pulmonary disease, n (%)	21 (37.5%)	11 (30.6%)	0.495
Restrictive lung disease, n (%)	26 (46.4%)	17 (47.2%)	0.941
Pulmonary hypertension, n (%)	30 (53.6%)	23 (63.9%)	0.328
Chronic kidney disease, n (%)	21 (37.5%)	12 (33.3%)	0.684
Prior cardiac surgery, n (%)	10 (17.9%)	6 (16.7%)	0.883
Prior percutaneous coronary intervention, n (%)	32 (57.1%)	8 (22.2%)	<0.001
Dialysis, n (%)	5 (8.9%)	2 (5.6%)	0.701
Mechanical ventilation, n (%)	14 (25.0%)	4 (11.1%)	0.101
Intra-aortic balloon pump, n (%)	0 (0.0%)	0 (0.0%)	1
Percutaneous microaxial pump, n (%)	10 (17.9%)	4 (11.1%)	0.379
Extracorporeal membrane oxygenation, n (%)	10 (17.9%)	2 (5.6%)	0.117
Right ventricular function			
Tricuspid annular plane systolic excursion, mm	16.0 (12.0–19.0)	15.0 (12.5–17.5)	0.808
Pulmonary artery systolic pressure, mmHg	44 (33–55)	37 (30–40)	0.527
Right atrial to pulmonary capillary wedge pressure ratio	0.42 (0.23–0.62)	0.37 (0.14–0.50)	0.425
Preoperative laboratory values			

Leukocyte count, /nL	7.3 (6.3–10.0)	9.2 (7.8–11.1)	0.013
Hemoglobin, g/dL	11.9 (9.4–13.4)	12.8 (10.3–14.1)	0.117
Platelet count, /nL	209 (141–250)	217 (174–298)	0.151
International Normalized Ratio	1.10 (1.03–1.26)	1.07 (1.02–1.19)	0.537
Fibrinogen, g/L	386.00 (298.00–494.00)	469.00 (409.00–640.50)	0.180
Creatinine, mg/dL	1.18 (0.91–1.60)	1.08 (0.91–1.44)	0.697
Glomerular filtration rate, mL/min/1.73 m ²	72.1 (49.4–86.2)	75.9 (49.3–96.0)	0.453
Bilirubin, mg/dL	0.54 (0.35–0.83)	0.62 (0.42–0.90)	0.590
Lactate, mmol/L	0.85 (0.70–1.27)	0.90 (0.62–1.40)	0.857
N-terminal-pro-B-type natriuretic peptide, ng/L	2046 (1355–4150)	3128 (1866–6588)	0.253
<i>Euroscore II, European System for Cardiac Operative Risk Evaluation; INTERMACS, Interagency Registry for Mechanically Assisted Circulatory Support; LVAD, left ventricular assist device; NYHA, New York Heart Association.</i>			

306

307

Table 2. Intraoperative course.

Variable	Crystalloid priming (n=56)	Blood-based priming (n=36)	P-value
Sternotomy, n (%)	46 (85.2%)	29 (80.6%)	0.775
Left lateral thoracotomy, n (%)	8 (14.8%)	7 (19.4%)	0.775
Cytokine adsorption, n (%)	3 (5.4%)	1 (2.8%)	1
Hemofiltration, n (%)	6 (10.7%)	1 (2.8%)	0.240
Cell saver use, n (%)	22 (39.3%)	19 (52.8%)	0.204
Total heparin dose, IU	40000 (35000–50000)	45000 (40000–50000)	0.281
Tranexamic acid, g	2.0 (2.0–2.0)	2.0 (2.0–4.0)	<0.001
Bypass time, min	104 (80–136)	98 (82–135)	0.839
Last intraoperative lactate, mmol/L	3.30 (2.25–5.07)	2.50 (1.90–5.15)	0.338
Hemoglobin at ICU admission, g/dL	9.2 (8.9–10.0)	10.3 (9.2–10.8)	0.002
Hemoglobin 24h postoperative, g/dL	9.3 (8.8–9.8)	9.5 (9.1–10.1)	0.070
Protamine, IU	35000 (30000–42750)	40000 (35000–45000)	0.213
Prothrombin complex concentrate, IU	2000 (0–3000)	1500 (0–2000)	0.049
Fibrinogen, g	4.0 (0.0–4.0)	2.0 (0.0–4.0)	0.456
Red blood cells, units	4 (1–7)	3 (2–5)	0.168
Fresh frozen plasma, units	4 (0–7)	4 (4–6)	0.575
Platelets, units	2 (2–4)	2 (2–3)	0.511
Inhaled pulmonary vasodilators, n (%)	35 (62.5%)	18 (50.0%)	0.236

All products listed above were administered as adjuncts to the priming solution and were not included in the prime itself.

Table 3. Postoperative outcomes.

Outcome	Crystalloid priming (n=56)	Blood-based priming (n=36)	RR (95% CI)	P-value
Right heart failure, n (%)	17 (30.4%)	3 (8.3%)	0.27 (0.09–0.87)	0.012
Dialysis, n (%)	18 (32.1%)	4 (11.1%)	0.35 (0.13–0.94)	0.021
Delirium, n (%)	16 (28.6%)	3 (8.3%)	0.29 (0.09–0.93)	0.019
Acute kidney injury, n (%)	3 (5.4%)	2 (5.6%)	1.04 (0.18–5.91)	1
Need for liver replacement therapy, n (%)	11 (19.6%)	2 (5.6%)	0.28 (0.07–1.20)	0.058
Any major bleeding, n (%)	26 (46.4%)	10 (27.8%)	0.60 (0.33–1.09)	0.074
Re-thoracotomy, n (%)	24 (42.9%)	9 (25.0%)	0.58 (0.31–1.11)	0.081
Ischemic stroke, n (%)	5 (8.9%)	3 (8.3%)	0.93 (0.24–3.67)	1
Hemorrhagic stroke, n (%)	2 (3.6%)	0 (0.0%)	0.31 (0.02–6.24)	0.518
Pneumonia, n (%)	24 (42.9%)	19 (52.8%)	1.23 (0.80–1.90)	0.352
Sepsis, n (%)	24 (42.9%)	15 (41.7%)	0.97 (0.60–1.59)	0.910
30-day mortality, n (%)	9 (16.1%)	1 (2.8%)	0.17 (0.02–1.31)	0.082
Right ventricular assist device, n (%)	4 (7.1%)	2 (5.6%)	0.78 (0.15–4.03)	1
Extracorporeal membrane oxygenation, n (%)	4 (7.1%)	3 (8.3%)	1.17 (0.28–4.91)	1
Inotropes >14 days, n (%)	16 (28.6%)	6 (16.7%)	0.58 (0.25–1.35)	0.191
Vasopressors >14 days, n (%)	20 (35.7%)	8 (22.2%)	0.62 (0.31–1.26)	0.170
Inhaled pulmonary vasodilators >14 days, n (%)	21 (37.5%)	14 (38.9%)	1.04 (0.61–1.76)	0.893
Total duration of inhaled nitric oxide, hours	12 (0–40)	0 (0–28)		0.272
Chest drain output 24 h, mL	820 (578–1152)	840 (485–1080)		0.451
Postoperative transfusions within 24 h (excluding priming), units				
Red blood cell units (24 h)	1 (0–2)	1 (0–2)		0.849
Fresh frozen plasma units (24 h)	0 (0–2)	0 (0–0)		0.459
Platelets (24 h)	0 (0–0)	0 (0–0)		0.548
Mechanical ventilation, hours	20 (1–39)	18 (2–40)		0.953
Intensive care unit stay, days	39.0 (20.8–49.8)	30.5 (19.0–51.0)		0.565

312

Table 4. Adjusted association between blood-based priming and key postoperative outcomes.

Outcome	Adjusted OR (95% CI)	P-value
Right heart failure	0.23 (0.06–0.86)	0.030
Dialysis	0.30 (0.09–1.01)	0.053
Delirium	0.22 (0.06–0.84)	0.027
Any major bleeding	0.47 (0.19–1.18)	0.109
<i>Models adjusted for age, sex, and INTERMACS profile.</i>		

313

314

315

Table 5. Multivariable Cox regression for all-cause mortality.

Predictor	Adjusted HR (95% CI)	P-value
Blood-based priming vs. crystalloid	0.59 (0.27–1.32)	0.201
Age, years	1.05 (1.01–1.10)	0.007
Female sex	1.47 (0.55–3.96)	0.445
INTERMACS profile (per 1-level increase)	0.52 (0.38–0.70)	<0.001
<i>Models adjusted for age, sex, and INTERMACS profile.</i>		

316

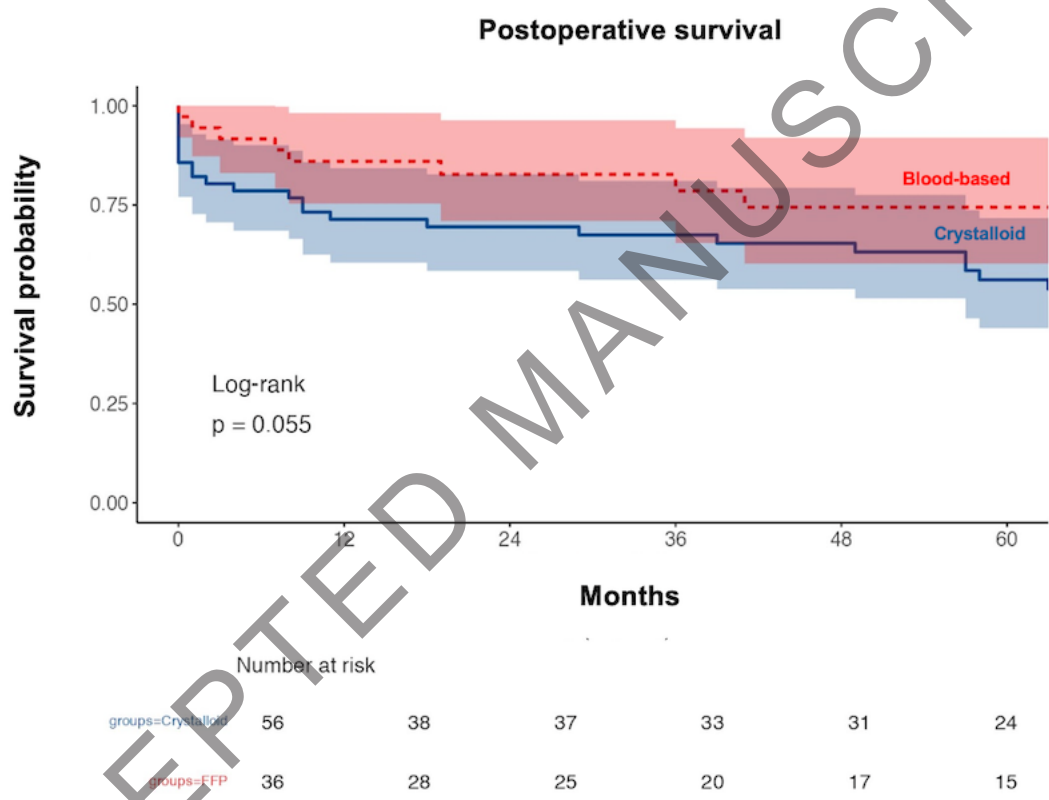
317

318 REFERENCES

- 319 (1) T. A. McDonagh, M. Metra, M. Adamo et al. 2021 esc guidelines for the
320 diagnosis and treatment of acute and chronic heart failure. *Eur Heart J*
321 2021;42:3599-726.
- 322 (2) E. V. Potapov, C. Antonides, M. G. Crespo-Leiro et al. 2019 eacts expert
323 consensus on long-term mechanical circulatory support. *Eur J Cardiothorac Surg*
324 2019;56:230-70.
- 325 (3) J. S. Hanke, G. Dogan, A. Zoch et al. One-year outcomes with the heartmate
326 3 left ventricular assist device. *J Thorac Cardiovasc Surg* 2018;156:662-69.
- 327 (4) I. M. Hariri, T. Dardas, M. Kanwar et al. Long-term survival on lvad support:
328 Device complications and end-organ dysfunction limit long-term success. *J Heart*
329 *Lung Transplant* 2022;41:161-70.
- 330 (5) U. P. Jorde, O. Saeed, D. Koehl et al. The society of thoracic surgeons
331 intermacs 2023 annual report: Focus on magnetically levitated devices. *Ann Thorac*
332 *Surg* 2024;117:33-44.
- 333 (6) D. Bellavia, A. Iacovoni, C. Scardulla et al. Prediction of right ventricular
334 failure after ventricular assist device implant: Systematic review and meta-analysis of
335 observational studies. *Eur J Heart Fail* 2017;19:926-46.
- 336 (7) B. C. Lampert and J. J. Teuteberg. Right ventricular failure after left ventricular
337 assist devices. *J Heart Lung Transplant* 2015;34:1123-30.
- 338 (8) S. Adamopoulos, M. Bonios, T. Ben Gal et al. Right heart failure with left
339 ventricular assist devices: Preoperative, perioperative and postoperative
340 management strategies. A clinical consensus statement of the heart failure
341 association (hfa) of the esc. *Eur J Heart Fail* 2024;26:2304-22.
- 342 (9) M. Dandel, E. Potapov, T. Krabatsch et al. Load dependency of right
343 ventricular performance is a major factor to be considered in decision making before
344 ventricular assist device implantation. *Circulation* 2013;128:S14-23.
- 345 (10) A. M. Beukers, J. V. Hugo, R. G. Haumann et al. Changes in colloid oncotic
346 pressure during cardiac surgery with different prime fluid strategies. *Perfusion*
347 2024;39:1371-79.
- 348 (11) C. Y. Xian-Yu, J. B. Xu, Y. T. Ma et al. Management of priming fluids in
349 cardiopulmonary bypass for adult cardiac surgery: Network meta-analysis. *Ann Med*
350 2023;55:2246996.
- 351 (12) M. Abedzadeh, N. Kachoueian, A. Fazli et al. The impact of using fresh frozen
352 plasma in cardiopulmonary bypass preparation on thromboelastometric parameters
353 and receiving blood products among pediatric patients undergoing cardiac surgery. *J*
354 *Cardiovasc Thorac Res* 2023;15:9-13.
- 355 (13) A. Dieu, M. Rosal Martins, S. Eeckhoudt et al. Fresh frozen plasma versus
356 crystalloid priming of cardiopulmonary bypass circuit in pediatric surgery: A
357 randomized clinical trial. *Anesthesiology* 2020;132:95-106.
- 358 (14) R. L. Kormos, C. F. J. Antonides, D. J. Goldstein et al. Updated definitions of
359 adverse events for trials and registries of mechanical circulatory support: A
360 consensus statement of the mechanical circulatory support academic research
361 consortium. *J Heart Lung Transplant* 2020;39:735-50.
- 362 (15) R. Salenger, R. C. Arora, A. Bracey et al. Cardiac surgical bleeding,
363 transfusion, and quality metrics: Joint consensus statement by the enhanced
364 recovery after surgery cardiac society and society for the advancement of patient
365 blood management. *Ann Thorac Surg* 2025;119:280-95.

- 366 (16) C. J. Kapelios, L. H. Lund, O. Wever-Pinzon et al. Right heart failure following
367 left ventricular device implantation: Natural history, risk factors, and outcomes: An
368 analysis of the sts intermacs database. *Circ Heart Fail* 2022;15:e008706.
- 369 (17) B. A. Houston, K. B. Shah, M. R. Mehra and R. J. Tedford. A new "twist" on
370 right heart failure with left ventricular assist systems. *J Heart Lung Transplant*
371 2017;36:701-07.
- 372 (18) C. Sciacaluga, M. C. Procopio, L. Potena et al. Right ventricular dysfunction
373 in left ventricular assist device candidates: Is it time to change our prospective?
374 *Heart Fail Rev* 2024;29:559-69.
- 375 (19) W. L. Miller. Fluid volume overload and congestion in heart failure. *Circulation:
376 Heart Failure* 2016;9:e002922.
- 377 (20) N. Bouabdallaoui, W. Beaubien-Souligny, A. Y. Denault and J. L. Rouleau.
378 Impacts of right ventricular function and venous congestion on renal response during
379 depletion in acute heart failure. *ESC Heart Fail* 2020;7:1723-34.
- 380 (21) M. M. McCall, M. M. Blackwell, J. T. Smyre et al. Fresh frozen plasma in the
381 pediatric pump prime: A prospective, randomized trial. *Ann Thorac Surg*
382 2004;77:983-7; discussion 87.
- 383 (22) S. Woods Cuneo, B. A. Abi-Nader, S. J. Blasczynski and M. Chigerwe. Effects
384 of plasma and hetastarch administration on colloid oncotic pressure and coagulation
385 variables in dairy calves and goats. *J Vet Intern Med* 2025;39:e70079.
- 386 (23) J. Udzik, J. Pacholewicz, A. Biskupski et al. Alterations to kidney physiology
387 during cardiopulmonary bypass-a narrative review of the literature and practical
388 remarks. *J Clin Med* 2023;12.
- 389 (24) J. Chen, Z. Ma, K. Peng, F. Ji and N. K. Shirakawa. Intraoperative colloid use
390 on post-operative renal function. *Curr Anesthesiol Rep* 2024;14:306-11.
- 391 (25) D. Knežević, B. Ćurko-Cofek, T. Batinac et al. Endothelial dysfunction in
392 patients undergoing cardiac surgery: A narrative review and clinical implications.
393 *Journal of Cardiovascular Development and Disease* 2023;10:213.
- 394 (26) M. Straat, M. C. Müller, J. C. Meijers et al. Effect of transfusion of fresh frozen
395 plasma on parameters of endothelial condition and inflammatory status in non-
396 bleeding critically ill patients: A prospective substudy of a randomized trial. *Crit Care*
397 2015;19:163.
- 398 (27) D. Greaves, P. J. Psaltis, D. H. J. Davis et al. Risk factors for delirium and
399 cognitive decline following coronary artery bypass grafting surgery: A systematic
400 review and meta-analysis. *J Am Heart Assoc* 2020;9:e017275.
- 401 (28) Y. Wang and B. Wang. Risk factors of delirium after cardiac surgery: A
402 systematic review and meta-analysis. *J Cardiothorac Surg* 2024;19:675.
- 403 (29) D. P. van den Brink, D. J. B. Kleinveld, C. A. Polet et al. Plasma attenuates
404 endothelial injury compared to crystalloids in a ventilated rat pneumosepsis model.
405 *PLoS One* 2025;20:e0319272.
- 406 (30) J. L. Mow, A. Gandhi and D. Fulford. Imaging the "social brain" in
407 schizophrenia: A systematic review of neuroimaging studies of social reward and
408 punishment. *Neuroscience & Biobehavioral Reviews* 2020;118:704-22.
- 409 (31) P. Bianchi, M. Cotza, C. Beccaris et al. Early or late fresh frozen plasma
410 administration in newborns and small infants undergoing cardiac surgery: The
411 appear randomized trial. *Br J Anaesth* 2017;118:788-96.
- 412 (32) A. C. Casbard, L. M. Williamson, M. F. Murphy, K. Rege and T. Johnson. The
413 role of prophylactic fresh frozen plasma in decreasing blood loss and correcting
414 coagulopathy in cardiac surgery. A systematic review. *Anaesthesia* 2004;59:550-8.

ACCEPTED MANUSCRIPT



Blood-based versus crystalloid cardiopulmonary bypass priming in left ventricular assist device surgery: impact on end-organ function and survival

Summary

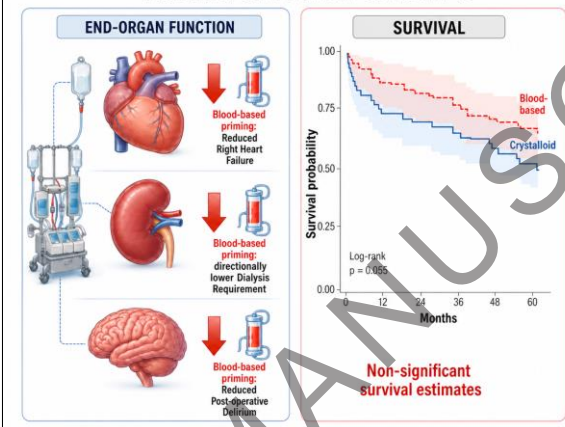
Retrospective, single-centre study of 92 patients with LVAD implantation.
CPB priming: blood-based (FFP/RBC, n=36) versus crystalloid (n=56).

Endpoints: right heart failure, dialysis, delirium, major bleeding, transfusions, and survival.

Blood-based priming showed:

- lower right heart failure and delirium
- directionally lower dialysis requirement
- non-significant survival estimates

IMPACT OF BLOOD-BASED VERSUS CRYSTALLOID PRIMING ON PATIENT OUTCOMES



Abbreviations: CPB, cardiopulmonary bypass; FFP, fresh frozen plasma; RBC, red blood cells; LVAD, left ventricular assist device.