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## Low-Dose Donor Dopamine is Associated with a Decreased Risk of Right Heart Failure in Pediatric Heart Transplant Recipients

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### Abstract

**Background**—Previous studies in adults have suggested that donor dopamine treatment may improve recipient outcomes in organ transplantation; in this analysis, we aimed to determine if donor dopamine reduces the incidence of post-operative right heart failure in pediatric heart transplant recipients.

**Methods**—Data for recipients aged 1–18 transplanted at our institution between 1/1/2000–6/15/2011 and their respective donors were obtained. The presence of postoperative right heart failure was assessed for in all subjects. Donor dopamine dose was stratified into 3 groups: none, low-dose ( $\leq 5$  mcg/kg/min), and high-dose ( $>5$  mcg/kg/min). Logistic regression was used to assess the relationship between donor dopamine dose and recipient right heart failure.

**Results**—Of 192 recipients, 34 (18%) experienced postoperative right heart failure. There was no difference in baseline demographics between recipients with and without right heart failure. When controlling for pulmonary vascular resistance index, graft ischemic time, and cardiopulmonary bypass time, donor low-dose dopamine was independently associated with a decreased risk of

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### DISCLOSURES

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right heart failure (OR=0.16 [0.04–0.70]; p=0.02); however high-dose DA was neither associated with, nor protective of, RHF (OR: 0.31 [0.06–1.6]; p=0.16).

**Conclusions**—Despite advances in perioperative care of the recipient, right heart failure persists as a complication of pediatric heart transplantation. In this study donor pre-treatment with low-dose dopamine is associated with a decreased risk of postoperative right heart failure in pediatric heart recipients. Further studies into this association may be useful in determining the utility of empiric donor pre-treatment with low-dose dopamine.

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## INTRODUCTION

Heart transplantation is an increasingly successful treatment strategy for end stage heart failure due to a wide variety of congenital and acquired cardiac diseases in children<sup>1</sup>. However, primary graft failure, and in particular postoperative acute right heart failure (RHF), remains a significant barrier to survival<sup>2–4</sup>.

Treatment of brain-dead donors with low-dose dopamine prior to graft procurement improves adult recipient outcomes in renal transplantation; reducing post-transplant dialysis requirements<sup>5</sup>. The use of inotropic support in potential cardiac donors, including dopamine, is usually considered a negative indicator of donor quality. However, low-dose dopamine pretreatment of brain-dead donors has recently been associated with improved long-term survival in adult cardiac transplant recipients<sup>6</sup>. Nevertheless, the possible peri-operative benefits of donor dopamine in pediatric cardiac transplant recipients have yet to be examined.

In this retrospective analysis, we examined the hypothesis that pretreatment of donors with dopamine is associated with improved postoperative outcomes, specifically the development of right heart failure, in pediatric heart transplant recipients.

## METHODS

### Study Population

After Institutional Review Board approval was obtained, the study population of all pediatric heart transplant recipients less than or equal to 18 years of age transplanted from Jan 1, 2000 through June 15, 2011 at our institution was identified. Clinical data were obtained through retrospective institutional recipient chart review and review of corresponding donor medical records. Donor data collected included age, gender, mechanism of death, requirement of cardiopulmonary resuscitation, use of thyroxine replacement, echocardiographic data and cardiac enzyme levels. Donor echocardiograms were considered abnormal if the report indicated an abnormal ejection fraction, wall motion abnormalities, or more than mild valvar regurgitation. Recipient demographics and pre-transplant factors along with operative variables such as graft ischemic time and cardiopulmonary bypass time were collected as well. For analysis, recipients were divided into 3 donor dopamine pre-treatment groups, determined at the time of organ procurement: no dopamine, low-dose dopamine (dose 5 mcg/kg/min), or high-dose dopamine (dose > 5 mcg/kg/min).

## Outcome Measures

The primary outcome measure of this study was postoperative right heart failure of the recipient requiring treatment. This outcome was defined by either the need for mechanical circulatory support of the right ventricle or rising central venous pressure with right ventricular hypocontractility and progressive right ventricular dilatation on echocardiography requiring medical intervention (initiation or increase of inotropic support in conjunction with pulmonary vasodilator therapy, typically inhaled Nitric Oxide).

Secondary outcomes measures included complication frequency, time to extubation, intensive care unit (ICU) length of stay, total postoperative length of stay, and short- and long-term survival.

## Statistical Analysis

Summary statistics are presented as median (interquartile range), mean  $\pm$  standard deviation, or number (percent) as appropriate. Donor and recipient characteristics and short-term outcomes were analyzed using Chi-square, Fisher's exact, Student's t-test, Wilcoxon rank-sum test, ANOVA, or Kruskal-Wallis test, as appropriate to variable type and distribution. The primary outcomes of the relationship between donor dopamine dose and recipient postoperative RHF was assessed using logistic regression. Variables for building the multivariable logistic regression model were based on known clinical factors, as well as statistical interactions. Those variables with an unadjusted p-value of  $<0.1$  were considered candidates for the multivariable model as were variable defined *a priori* to be assumed risk factors of RHF. In deference to parsimony, variables that were neither significant, nor had interactions (defined as a minimum of a 10% change in coefficient) with other variables were eliminated from the final model. Kaplan-Meier analysis with log-rank test was used to analyze actuarial survival. The conventional p-value  $< 0.05$  was used to determine statistical significance, and all reported p-values are 2-sided. The data were analyzed with StataIC 11.0 (Stata Corp LP, College Station, TX).

## RESULTS

### Study Population

A total of 192 pediatric heart transplant recipients 18 years of age were transplanted from January 1, 2000 through June 15, 2011 at our institution. The total study population had a median age of 8.25 years (IQR: 1.7–13.8 years) and was 46.9% female. Indications for transplant were cardiomyopathy (59.4%), congenital heart disease (29.7%), retransplantation (6.8%), and other diagnoses (4.2%); including viral myocarditis and intracardiac tumor. At the time of transplant, 82% were UNOS status 1A with 13% on mechanical ventilation, 66% on high-dose or multiple inotropic support, and 19% on mechanical circulatory support. Of the 192 donors, 118 (61%) were on inotropic support (epinephrine, norepinephrine, milrinone, dobutamine or dopamine) at the time of organ procurement with 17 of those on 2 or more inotropic drips. Dopamine was the most commonly used inotrope in 77 donors (40%), followed by norepinephrine (n=30, 16%), epinephrine (n=19, 10%), dobutamine (n=9, 5%) and milrinone (n=2, 1%). With respect to donor dopamine administration, 115 (60%) recipients were included in the no dopamine group, 51 (26%) in the low-dose group,

and 26 (14%) in the high-dose group. Recipient baseline characteristics did not differ significantly between donor dopamine groups (Table 1a). Furthermore, there were no significant differences in donor characteristics across dopamine dose groups (Table 1b).

### Clinical outcomes based on donor dopamine dose

Of the 192 recipients, 34 (17.7%) experienced the primary outcome of postoperative RHF. With the exception of pulmonary vascular resistance index (PVRi), recipient baseline characteristics did not differ between those who had RHF and those who did not (Table 2). Importantly, perioperative characteristics of graft cold ischemic time and cardiopulmonary bypass time were not different between those who experienced postoperative RHF and those who did not.

Donor administration of dopamine showed significant associations with recipient outcomes (Table 3). Specifically, there was a significant reduction in the development of RHF when the donor received dopamine, particularly low-dose dopamine (none: 23.5%, low-dose: 7.8%, high-dose: 11.5%;  $p=0.03$ ). Those recipients whose donors received dopamine had a significantly shorter time to extubation and a shorter hospital length of stay compared to the no dopamine group. Of note, there was no significant difference in short-term recipient mortality (30-day and 1-year) by donor dopamine dose.

### Clinical outcomes based on RHF status post-transplant

Recipient postoperative RHF was significantly associated with poorer clinical outcomes (Table 4). Median time to extubation (4 vs 2 days,  $p=0.0001$ ) and ICU length of stay (12 vs 8 days,  $p=0.01$ ) were significantly longer in recipients who developed RHF as compared to those who did not. Furthermore, the development of RHF was associated with worse 30-day mortality (11.8% vs. 1.3%,  $p=0.01$ ). Despite the increase in 30-day mortality, RHF did not appear to be associated with poorer overall survival ( $p=0.41$ ) (Figure 1).

### Relationship between Donor Dopamine Administration and Recipient RHF

In univariable logistic regression analysis, only donor pretreatment with dopamine and recipient pretransplant PVRi were significantly associated with the development of postoperative RHF. A list of all variables considered for analysis is provided in Table 5a. Based on the univariable analysis, risk factors included in the multivariable logistic regression were donor dopamine group, recipient pre-operative PVRi (dichotomized at  $6 \text{ WU} \times \text{m}^2$ ), and the *a priori* determined risk factors of graft cold ischemic time, and total cardiopulmonary bypass time. The results of the multivariable logistic regression are shown in Table 5b. As the nonsignificant risk factors did not show interaction with the primary variable of interest (donor dopamine), a reduced model was created by removing the nonsignificant risk factors of graft cold ischemic time and total cardiopulmonary bypass time (Table 5c). Most importantly, when controlling for graft ischemic time and total cardiopulmonary bypass time; donor low-dose dopamine remained associated with a decreased risk of recipient postoperative RHF (OR: 0.16 [0.34–0.78];  $p=0.02$ ), independent of recipient pre-operative PVRi. However, high-dose dopamine was neither associated with, nor protective of recipient postoperative RHF (OR: 0.32 [0.06–1.6];  $p=0.16$ ). This relationship held true even when not controlling for ischemic and cardiopulmonary bypass

times. Despite this association there was no significant difference in long-term survival when recipients were stratified by their donor's dopamine dose ( $p=0.66$ ) (Figure 2). To examine the lack of association with high-dose dopamine, the multivariable model was rerun using low-dose dopamine as the indicator variable. In this model, no dopamine was significantly associated with an increased risk of postoperative RHF (OR: 6.15 [1.28–29.4],  $p=0.02$ ), and high-dose dopamine was again not significantly associated with the outcome (OR: 1.93 [0.235–15.84],  $p=0.54$ ).

## DISCUSSION

While heart transplantation is a largely successful therapy for children with end stage heart failure, early mortality remains a significant problem; despite the improved treatment of and decreasing mortality from early postoperative right heart failure (4). While the association of right heart failure and high recipient pulmonary vascular resistance is well described, the importance of donor factors prior to procurement is becoming increasingly clear<sup>2–4,7–9</sup>. The use of low-dose dopamine in donors prior to organ procurement has been shown to improve kidney graft outcomes, and may be linked to improved long-term outcomes in adult heart recipients<sup>5,6</sup>. The present study showed an association between the use of low-dose dopamine in donors and a decrease in the incidence of early postoperative right heart failure in pediatric heart recipients as well as benefits in duration of mechanical ventilation and hospital length of stay.

While elevated pulmonary vascular resistance is often considered the prime risk factor for right heart failure after orthotopic heart transplantation, it has been demonstrated that the cardiac graft is likely predisposed to right ventricular failure well before organ procurement. Animal studies have shown significantly decreased right and left ventricular preload recruitable stroke work in both the brain dead donor as well as the recipient of the graft<sup>8,10</sup>. Interestingly, the right ventricle appears to be more susceptible to this type of injury<sup>11</sup>. This decrease in right ventricular contractility pre-procurement is further worsened by cold ischemia; resulting in a graft that is significantly impaired in its ability to adapt to increases in right ventricular afterload<sup>8</sup>, a problem compounded by elevated recipient pulmonary vascular resistance. Besides stimulating dopamine specific receptors at low doses, recent studies have suggested that the benefit of low-dose dopamine may be secondary to antioxidant properties which can attenuate the damage myocytes experience during cold ischemia, thereby minimizing at least one of the factors contributing to postoperative right heart failure<sup>12</sup>. In this study, only low dose dopamine exhibited a protective effect on the development of right heart failure. It is possible that higher dose dopamine was an indicator of donor hemodynamic instability, cardiac dysfunction, or perhaps there may be negative effects upon the graft at higher doses due to increased myocardial oxygen consumption or other mechanisms. Additionally, as few donors were on high-dose dopamine, we may not be able to detect a true difference between high-dose dopamine and no dopamine due to inadequate statistical power. Certainly prospective studies would be needed to assess whether this protective benefit is limited to low-dose dopamine or if intermediate or higher doses could confer a similar benefit in a suitable donor.

The study population was derived from recipients at a single high-volume institution. This allowed for better ascertainment of the occurrence of right heart failure, data which is not routinely collected in current multicenter registries such as the UNOS/OPTN dataset or ISHLT registry. To that end, although the study period encompassed over a decade of pediatric heart transplants, there was uniformity of treatment throughout the study period in the ICU as the use of inhaled nitric oxide for posttransplant right heart failure began in 1995. The only difference in clinical care was the use of sildenafil instead of nifedipine to transition off of inhaled nitric oxide in recent years.<sup>4,13</sup> Furthermore, donor management occurred at the local level, and the recipient medical team did not consider type or dose of donor medications into their treatment decisions for the recipient, making such bias unlikely. To this end, it is interesting to hypothesize why some donors received dopamine infusions while others did not. Of course, the lack of clarity as to the indication for dopamine administration in the donor is a major limitation. Due to the retrospective nature of this study not only are the decision making processes of the donor team impossible to know, but even the specifics of duration of treatment and any titration of the dopamine dose are unknown. It is entirely possible that these unknown variables are responsible for the findings of this study and donor dopamine administration is merely a mediator or causal partner in the effect model. This would not negate the association seen in this study, but rather offer a different understand of why it is. However, in preliminary analysis, no obvious donor factors seemed to play a role in the decision to administer dopamine, and as literature supporting the use of donor dopamine is quite recent, it is unlikely that this played a factor either<sup>5,6</sup>. Of note, we did not notice any time dependent effect in the proportion of donors receiving dopamine with similar rates of use occurring throughout the study period (unpublished data).

The usual limitations of retrospective data collection and analysis were minimized in this study as determination of right heart failure occurred independently and prior to donor data collection, eliminating this particular bias. While specific measurements of right heart function (central venous pressure, right ventricular end-diastolic dimension, etc.) were unable to be attained for all patients, the records were sufficiently complete to determine the initiation of treatment (mechanical circulatory support, inotropes, inhaled nitric oxide) for clinical right heart failure as described above. The association with poorer clinical outcomes (increased time of mechanical ventilation, hospital length of stay and 30-day mortality) appears to validate the determination of right heart failure as described; however, we acknowledge that the lack of an objective measure of right heart failure remains a major limitation of the study and tempers the findings. Furthermore, we were able to obtain all donor records for patients transplanted at our institution during the study period, confirming either the absence or use of donor dopamine and also the specific dose, thereby minimizing any selection bias that could have occurred due to incomplete records. Our institution has a high proportion (18%) of recipients with pretransplant elevated PVRi ( $>6 \text{ WU} \times \text{m}^2$ ), however even when controlling for recipient PVRi, low-dose donor dopamine remained protective of right heart failure, suggesting that the benefit of donor dopamine persists even in the face of elevated recipient PVRi. We similarly also controlled for factors previously defined as conferring risk of right heart failure (graft cold ischemic time, total cardiopulmonary bypass time)<sup>2,3</sup>, despite no significant associations in our study population. While this study did not show a survival benefit to the use of low-dose donor dopamine, this

is likely due to improved treatment of right heart failure resulting in equivalent survival curves for patients with and without early right heart failure, as well as the low number of early deaths (6 of 192) in the study population, making this study underpowered to detect a difference for that time point.

Despite advances in perioperative care of the pediatric heart recipient, early right heart failure remains prevalent and can lead to further morbidities and mortality. Although early right heart failure may have a multifactorial etiology, donor pre-treatment with low-dose dopamine may abrogate the risk of the development of early right heart failure in pediatric heart recipients leading to decreases in duration of mechanical ventilation and overall hospital stay. Given the limitations of this study, further follow-up and larger, prospective studies are necessary to fully evaluate the effects of donor pre-treatment with low-dose dopamine on the pediatric heart recipient as well as the pediatric recipients of other solid organs.

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### FREEDOM OF INVESTIGATION

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## Abbreviations

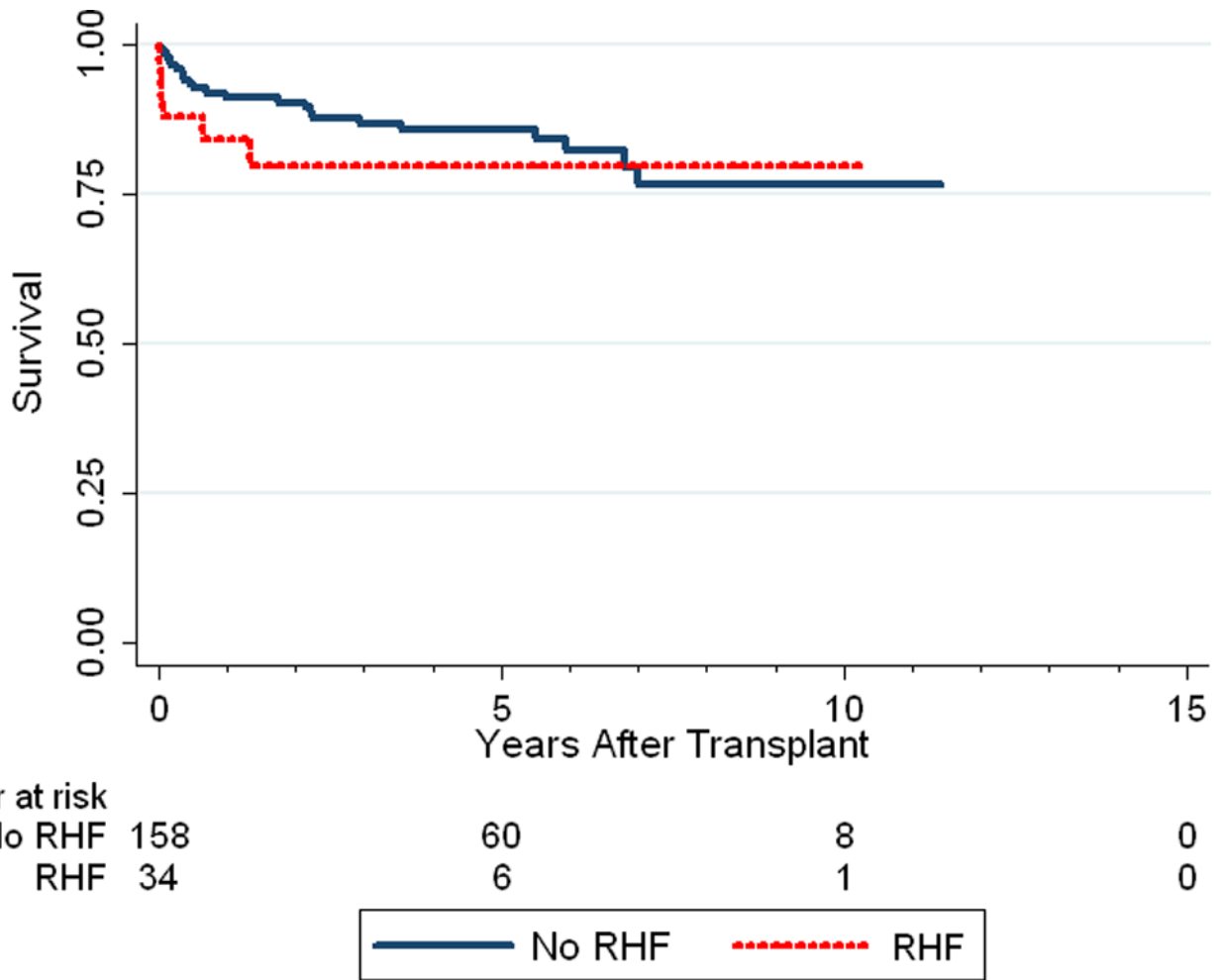
<b>RHF</b>	Right Heart Failure
<b>PVRi</b>	Pulmonary Vascular Resistance Index
<b>ICU</b>	Intensive Care Unit

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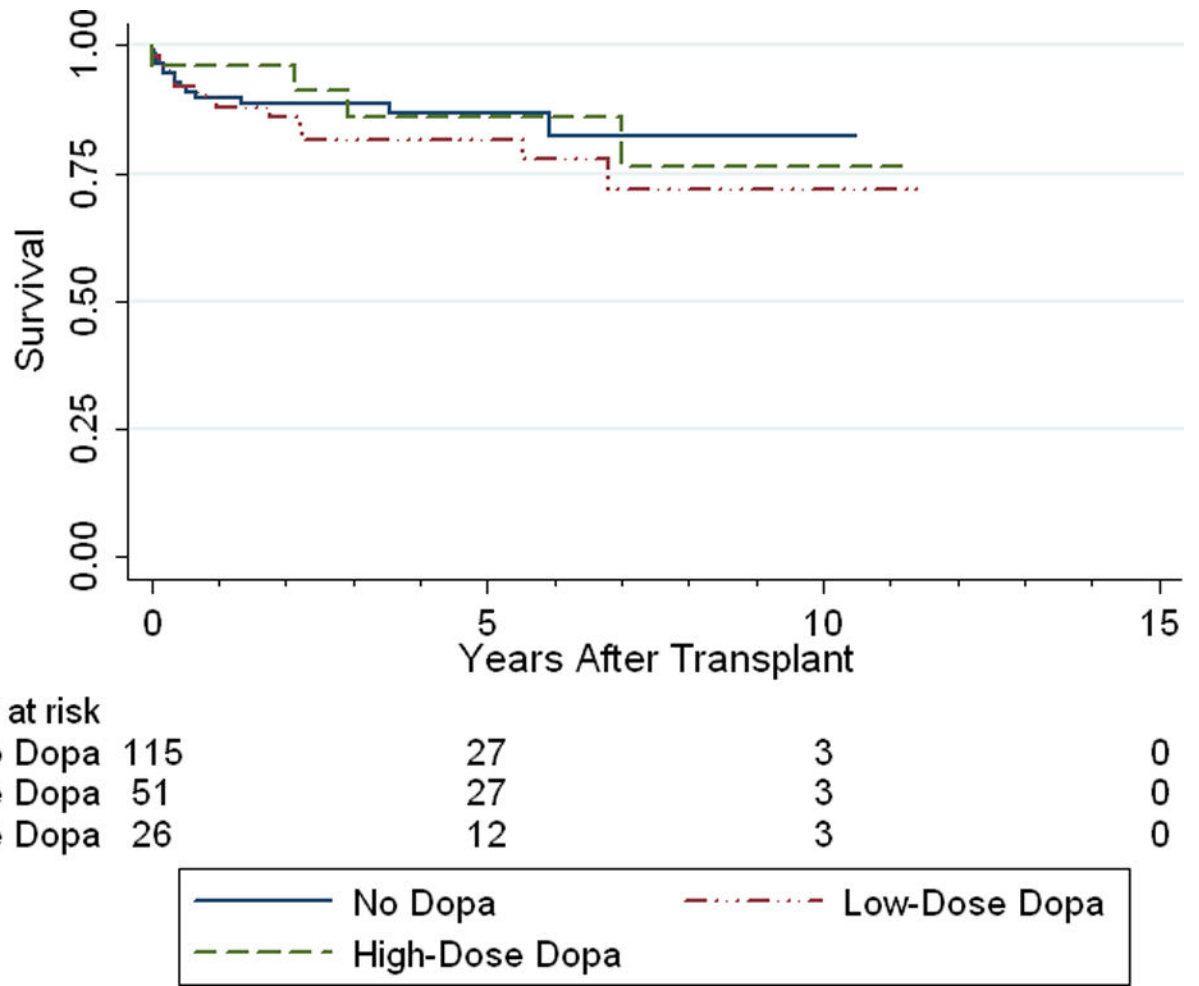
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**Figure 1.** Kaplan-Meier survival curve depicting recipient survival between those who experienced RHF and those who did not (p=0.41).



**Figure 2.**  
Kaplan-Meier recipient survival curve divided by donor dopamine dose

Table 1

a. Recipient Demographics compared among donor dopamine groups.				
	No Dopamine (n=115)	Low-Dose Dopamine (n=51)	High-Dose Dopamine (n=26)	p-value
Age (IQR) (years)	8.9 (1.7–13.9)	7.7 (1.7–15.8)	7.2 (1.6–11.8)	0.74
BSA (IQR) (m <sup>2</sup> )	0.9 (0.5–1.4)	0.9 (0.4–1.6)	0.8 (0.5–1.2)	0.64
Gender (% female)	52 (45.2%)	25 (49.0%)	13 (50.0%)	0.85
Cardiac diagnosis	-	-	-	0.38
Cardiomyopathy	65 (56.5%)	36 (70.6%)	13 (50.0%)	
Congenital	36 (31.3%)	11 (21.6%)	10 (38.5%)	
Retransplant	8 (7.0%)	2 (3.9%)	3 (11.5%)	
Ventilator support, pre-operative	12 (10.4%)	8 (15.7%)	5 (19.2%)	0.39
Inotropic support, pre-operative	82 (71.3%)	32 (62.8%)	13 (50%)	0.11
VAD, pre-operative	13 (11.3%)	9 (17.6%)	2 (7.7%)	0.42
ECMO, pre-operative	5 (4.4%)	4 (7.8%)	3 (11.5%)	0.27
PVRI, pre-operative (IQR) (WU × m <sup>2</sup> )	2.7 (1.7–5.6)	3.0 (2.1–5.1)	2.4 (1.7–3.1)	0.52
Graft cold ischemic time (min)	196 ± 55	214 ± 68	216 ± 80	0.13
Cardiopulmonary bypass time (min)	158 ± 60	150 ± 48	154 ± 61	0.73

b. Donor demographics compared among donor dopamine groups				
	No Dopamine (n=115)	Low-Dose Dopamine (n=51)	High-Dose Dopamine (n=26)	p-value
Donor Age (IQR) (years)	11.0 (2.0–18.0)	8.0 (1.6–16.6)	6.6 (1.2–15.1)	0.34
Donor BSA (IQR) (m <sup>2</sup> )	1.2 (0.6–1.7)	1.0 (0.5–1.6)	0.9 (0.5–1.5)	0.29
Donor Gender (% female)	50 (43.5%)	25 (49.0%)	14 (53.8%)	0.58
Cause of death	-	-	-	0.29
Anoxia	37 (32.2%)	17 (33.3%)	5 (19.2%)	
Cerebrovascular	16 (13.9%)	10 (19.6%)	1 (3.8%)	
Head trauma	57 (49.6%)	22 (43.1%)	18 (69.2%)	
Downtime	52 (45.2%)	23 (45.1%)	9 (34.6%)	0.60
Positive blood culture	7 (6.1%)	4 (7.8%)	1 (3.8%)	0.83
Thyroxine replacement	46 (40.0%)	17 (33.3%)	7 (23.1%)	0.24
Vasopressin administration	46 (40.0%)	19 (37.2%)	15 (57.7%)	0.20
Other inotrope administration	41 (36%)	10 (20%)	5 (20%)	0.06
Ejection fraction (%)	61.1 ± 7.7	59.4 ± 5.4	62.4 ± 1.4	0.10
Abnormal echocardiogram	7 (6.1%)	4 (7.8%)	3 (11.5%)	0.52
Troponin, at procurement (IQR) (ng/mL)	0.10 (0.04–0.66)	0.10 (0.04–0.40)	0.21 (0.04–0.80)	0.85

**Table 2**

Recipient characteristics by development of right heart failure (RHF)

	No RHF (n=158)	RHF (n=34)	p-value
Age (IQR) (years)	7.6 (1.5–13.9)	10.6 (3.2–13.6)	0.24
BSA (IQR) (m <sup>2</sup> )	0.84 (.43–1.4)	1.1 (.6–1.4)	0.25
Gender (% female)	76 (48.1%)	14 (41.2%)	0.57
Cardiac diagnosis, on admission	-	-	0.07
Cardiomyopathy	100 (63.3%)	14 (41.2%)	
Congenital	44 (27.9%)	13 (38.2%)	
Retransplant	9 (5.7%)	4 (11.8%)	
Ventilator support, pre-operative	22 (13.9%)	3 (8.8%)	0.58
High-dose or multiple inotropes	101 (63.9%)	26 (76.5%)	0.23
VAD, pre-operative	21 (13.3%)	3 (8.8%)	0.58
ECMO, pre-operative	9 (5.7%)	3 (8.8%)	0.45
PVRI, pre-operative (IQR) (WU × m <sup>2</sup> )	2.7 (1.8–4.3)	4.4 (1.8–8.6)	0.01
Graft cold ischemic time (mins)	204.8 ± 64.2	197.7 ± 54.8	0.55
Cardiopulmonary bypass time (mins)	153.7 ± 59.4	163.2 ± 45.6	0.40

**Table 3**

Clinical Outcomes by donor dopamine dose

	No Dopamine (n=115)	Low-Dose Dopamine (n=51)	High-Dose Dopamine (n=26)	p-value
<b>Right Heart Failure</b>	<b>27 (23.5%)</b>	<b>4 (7.8%)</b>	<b>3 (11.5%)</b>	<b>0.03</b>
<b>Mechanical Circulatory Support</b>	7 (6.1%)	1 (2.0%)	1 (3.8%)	0.63
<b>Time to Extubation (IQR) (days)</b>	<b>2 (1–5)</b>	<b>1 (1–3)</b>	<b>2 (1–4)</b>	<b>0.01</b>
<b>ICU Length of Stay (IQR) (days)</b>	10 (6–17)	7 (5–11)	9 (6–15)	0.07
<b>Post-op Length of Stay (IQR) (days)</b>	<b>21 (14–33)</b>	<b>16 (11–21)</b>	<b>18 (12–26)</b>	<b>0.01</b>
<b>30-day Mortality</b>	4 (3.5%)	1 (2.0%)	1 (3.8%)	0.84
<b>1-year Mortality</b>	10 (8.7%)	6 (11.7%)	1 (3.8%)	0.63

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**Table 4**

Clinical outcomes by RHF status

	No RHF (n=158)	RHF (n=34)	p-value
<b>Mechanical Circulatory Support</b>	6 (3.8%)	3 (8.8%)	0.20
<b>Time to Extubation (IQR) (days)</b>	<b>2 (1–4)</b>	<b>4 (3–7)</b>	<b>0.0001</b>
<b>ICU Length of Stay (IQR) (days)</b>	<b>8 (6–15)</b>	<b>12 (9–18)</b>	<b>0.01</b>
<b>Post-op Length of Stay (IQR) (days)</b>	18 (12–28)	24 (17–30)	0.10
<b>30-day Mortality</b>	<b>2 (1.3%)</b>	<b>4 (11.8%)</b>	<b>0.01</b>
<b>1-year Mortality</b>	12 (8.9%)	5 (19.3%)	0.16

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Table 5

a. Univariate analysis of proposed factors associated with postoperative RHF			
Variable	Unadjusted Odds Ratio	95% Confidence Interval	p-value
Donor: Weight (kg)	1.4	0.8–2.6	0.24
Donor: Cause of death	1.1	0.7–1.6	0.68
Donor: Low-dose dopamine pre-treatment	<b>0.3</b>	<b>0.1–0.8</b>	<b>0.02</b>
Donor: High-dose dopamine pre-treatment	0.4	0.1–1.4	0.16
Donor: Other inotrope pre-treatment	1.8	0.6–4.9	0.27
Donor: Thyroxine pre-treatment	0.9	0.4–2.0	0.88
Donor: Downtime (y/n)	0.8	0.4–1.9	0.74
Donor: Abnormal echocardiogram	0.7	0.2–3.6	0.73
Recipient: Admission diagnosis	1.1	0.7–1.9	0.61
Recipient: Ventilator support, pre-operative	0.6	0.2–2.1	0.43
Recipient: VAD placement, pre-operative	0.6	0.2–2.3	0.48
Recipient: ECMO, pre-operative	1.6	0.4–6.3	0.50
Recipient: High-dose/Multiple Inotropes	1.8	0.8–4.3	0.17
Recipient: Status 1A, prior to transplant	1.8	0.6–5.4	0.32
Recipient: PVRi, pre-operative	<b>1.2</b>	<b>1.1–1.4</b>	<b>0.006</b>
Graft cold ischemic time (mins)	1.0	0.9–1.1	0.55
Cardiopulmonary bypass time (mins)	1.0	0.9–1.1	0.40

b. Multivariable model of risk factors associated with postoperative RHF			
Variable	Adjusted Odds Ratio	95% Confidence Interval	p-value
Donor: Low-dose dopamine	0.16	0.04–0.78	0.02
Donor: High-dose dopamine	0.31	0.06–1.6	0.16
Recipient: Elevated Pre-operative PVRi (>6 WU × m <sup>2</sup> )	4.0	1.4–11.9	0.01
Graft cold ischemic time (mins)	0.99	0.98–1.0	0.40
Cardiopulmonary bypass time (mins)	1.0	0.99–1.01	0.61

c: Reduced multivariable model of risk factors associated with postoperative RHF			
Variable	Adjusted Odds Ratio	95% Confidence Interval	p-value
Donor: Low-dose dopamine	0.14	0.03–0.69	0.02
Donor: High-dose dopamine	0.31	0.06–1.5	0.15
Recipient: Elevated Pre-operative PVRi (>6 WU × m <sup>2</sup> )	5.1	1.8–14.5	0.002