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Combined Heart and Kidney Transplantation – Is There a Protective Effect Against Cardiac Allograft Vasculopathy by Intravascular Ultrasound?

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Abstract

Background: Combined organ transplants have been associated with less acute and chronic post-transplant rejection than isolated organ transplants, suggesting an altered immune response to the multi-organ allograft milieu. The incidence of combined heart and kidney transplantation (HKTx) has significantly increased over the last few years. Whether the addition of kidney transplantation has a protective immune effect against developing cardiac allograft vasculopathy (CAV) has not been fully investigated. We evaluated the incidence of CAV in HKTx recipients at our single center.

Methods: Between 2010-2016, we assessed 23 HKTx patients (pts) and 224 isolated heart transplant (HTx) pts in the first-year post-transplant. Pts underwent first-year intravascular ultrasound (IVUS) at 4-6 weeks (baseline) and at 12 months. The change in maximal intimal thickness (MIT) was obtained in all pts to evaluate for early post-transplant CAV. Diagnosis of CAV was based on $a \ge 0.5$ mm increase in maximal intimal thickness (MIT) from baseline in any matched site. Clinical outcomes such as 1-year survival, 1-year freedom from non-fatal major cardiac events (NF-MACE: myocardial infarction, new onset heart failure, coronary intervention, defibrillator or pacemaker implant, stroke), 1-year freedom from any-treated rejection, acute cellular rejection and antibody-mediated rejection were also examined.

Demographics

Demographics Mean Preinight Age, Vegra + SD		H	HKTX HTX			P-Value	
		() 50	n=23)	(n=224))	0 411	
Mean Donor Age, Years \pm SD		38.	8 ± 10.4	30.4 ± 13.0 24.2 ± 12.7		0.411	
Redu Mass Index Masn + SD		39. 24	7 ± 12.2	$34.2 \pm 13.$		0.061	
Body Mass Index, Mean \pm SD		24	$.3 \pm 5.1$	25.2 ± 4.4		0.349	
remaie (%)		2	26.1%	28.6%		0.801	
Previous Pregnancy in Females (%)		5	50.0%	81.0%		0.08	
Ischemic Time, Mean Mins ± SD		155	$.8 \pm 54.3$	161.7 ± 57.1		0.638	
Underlying Diagnosis of Coronary		6	5.0%	34.1%		0.006	
Artery Disease (%)				75 40/		0.442	
Status I at Transplant (%)		5	52.6%	/5.4%		0.443	
Cytomegalovirus Mismatch (%)			27.3% 28.8%			0.882	
Diabetes Mellitus (%)		2	26.1%	23.7%		0.795	
Treated Hypertension (%)		7	8.6%	47.0%		0.022	
Insertion of Mechanical Circulatory Support Device (%)		2	26.1%	22.3%		0.681	
Prior Blood Transfusion (%)		7	6.2% 41.5%			0.002	
Pre-Transplant PRA ≥ 10% (%)		3	80.4%	29.9%		0.958	
Pre-Transplant Creatinine, Mean ± SD		3.	2 ± 1.9	1.9 1.2 ± 0.4		<.001	
Outco		m	es				
Endpoints	HKTx (n=23)		HTx (n=224)		P-Value		
Mean Baseline MIT (mm) ± SD	0.4 ± 0.4		0.3 ± 0.4		-	-Value	
Mean 1-Vear MIT			0.3 :	± 0.4	-	-Value 0.255	
$(mm) \pm SD$	0.6 ± 0.4		0.3 = 0.6 =	± 0.4 ± 0.5		-Value 0.255 1.000	
$(mm) \pm SD$ $Mean \Delta MIT (mm) \pm SD$	$\begin{array}{c} 0.6\pm0.4\\ 0.2\pm0.3\end{array}$		0.3 = 0.6 = 0.3 =	± 0.4 ± 0.5 ± 0.4		-Value 0.255 1.000 0.245	
$(mm) \pm SD$ $Mean \Delta MIT (mm) \pm$ SD $\Delta MIT \geq 0.5 mm, \%$	0.6 ± 0.4 0.2 ± 0.3 13.0% (3/2)	3)	0.3 = 0.6 = 0.3 = 25.5% (± 0.4 ± 0.5 ± 0.4 (57/224)		-Value 0.255 1.000 0.245 0.305	
Initial form(mm) \pm SDMean Δ MIT (mm) \pm SD Δ MIT \geq 0.5 mm, %1-Year Survival	0.6 ± 0.4 0.2 ± 0.3 13.0% (3/2) 100.0%	3)	0.3 = 0.6 = 0.3 = 25.5% (100	± 0.4 ± 0.5 ± 0.4 (57/224) 0.0%		-Value 0.255 1.000 0.245 0.305 1.000	
Initial Formula Formula $(mm) \pm SD$ Mean Δ MIT $(mm) \pm$ SD Δ MIT \geq 0.5 mm, %1-Year Survival1-Year Freedom fromNF-MACE	0.6 ± 0.4 0.2 ± 0.3 13.0% (3/2) 100.0% 100.0%	3)	0.3 = 0.6 = 0.3 = 25.5% (100 96.	± 0.4 ± 0.5 ± 0.4 (57/224) 0.0%		-Value 0.255 1.000 0.245 0.305 1.000 0.330	
Intent I= Itean INIT(mm) ± SDMean Δ MIT (mm) ±SDΔ MIT ≥ 0.5 mm, %1-Year Survival1-Year Freedom fromNF-MACE1-Year Freedom fromAny-Treated Rejection	0.6 ± 0.4 0.2 ± 0.3 13.0% (3/2) 100.0% 82.6%	3)	0.3 = 0.6 = 0.3 = 25.5% (100 96. 88.	± 0.4 ± 0.5 ± 0.4 (57/224) 0.0% 0.0%		-Value 0.255 1.000 0.245 0.305 1.000 0.330 0.330	
<pre>Intent I⁻ Teal INIT (mm) ± SD Mean Δ MIT (mm) ± SD Δ MIT ≥ 0.5 mm, % 1-Year Survival 1-Year Freedom from NF-MACE 1-Year Freedom from Any-Treated Rejection 1-Year Freedom from Acute Cellular Rejection</pre>	0.6 ± 0.4 0.2 ± 0.3 13.0% (3/2) 100.0% 82.6% 100.0%	3)	0.3 = 0.6 = 0.3 = 25.5% (100 96. 88. 88.	± 0.4 ± 0.5 ± 0.4 (57/224) 0.0% 0.0% 0.0% 0.0% 0.0% 0.0% 0.0% 0.0% 0.0%		-Value 0.255 1.000 0.245 0.305 1.000 0.330 0.330 0.330	

<u>Results</u>: There was no significant difference in the proportion of HKTx vs HTx pts with Δ MIT ≥ 0.5 mm in the first-year post-transplant. The mean baseline MIT, 1-year MIT, and change in MIT was equivalent between the two groups. There was no significant difference in 1-year survival, 1-year freedom from NF-MACE, and 1-year freedom from rejection between the two groups.

<u>Conclusion</u>: There is no difference in CAV progression at 1-year post-transplant between isolated HTx pts and dual HKTx pts. Further investigation is warranted to confirm these results.

Background

- Combined organ transplants have been associated with less acute and chronic post-transplant rejection than isolated organ transplants, suggesting an altered immune response to the multi-organ allograft milieu.
- The incidence of combined heart and kidney transplantation (HKTx) has significantly increased over the last few years.
- Whether the addition of kidney transplantation has a protective immune effect against developing cardiac allograft vasculopathy (CAV) has not been fully investigated.

Purpose

• To assess the incidence of CAV by IVUS in HKTx recipients at our single center.

Methods

- Between 2010-2016, we assessed 23 HKTx patients (pts) and 224 isolated heart transplant (HTx) pts in the first-year post-transplant
- Pts underwent first-year intravascular ultrasound (IVUS) at 4-6 weeks (baseline) and at 12 months

Results Summary

- There was no significant difference in the proportion of HKTx vs HTx pts with Δ MIT \geq 0.5 mm in the first-year post-transplant
- The mean baseline MIT, 1-year MIT, and change in MIT was similar between the two groups
- There was no significant difference in 1-year survival, 1-year freedom from NF-MACE, and 1-year freedom from rejection
- The change in maximal intimal thickness (MIT) was obtained in all pts to evaluate for early post-transplant CAV
- Diagnosis of CAV was based on a \geq 0.5 mm increase in maximal intimal thickness (MIT) from baseline in any matched site
- Endpoints included:
 - 1-year survival
 - 1-year freedom from non-fatal major cardiac events (NF-MACE: myocardial infarction, new onset heart failure, coronary intervention, defibrillator or pacemaker implant, stroke)
 - 1-year freedom from any-treated rejection
 - 1-year freedom from acute cellular rejection
 - 1-year freedom from antibody-mediated rejection

between the two groups



• Combined heart and kidney transplant does not appear to provide a protective effect on CAV development (by IVUS).

• Further investigation is warranted to confirm these results

Author Disclosures

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