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CLINICAL RESEARCH

Repair or Replacement for Isolated Tricuspid Valve Pathology? Insights from a Surgical Analysis on Long-Term Survival

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Back Material/N	kground: Aethods:	limited. Current American Heart Association guidelin gical intervention is indicated. Our aim was to investi vival in patients undergoing isolated tricuspid valve Between 1995 and 2011, 109 consecutive patients u at our institution for varying structural pathologies.	nderwent surgical correction of tricuspid valve pathology A total of 41 (37.6%) patients underwent tricuspid annu- on, while 68 (62.3%) patients received tricuspid valve re-				
	Results:	Early survival at 30 days after surgery was 97.6% in the 89.1% in the TAP group and 87.8% in the TVR group further mortality observed after one year post surger	The TAP group and 91.1% in the TVR group. After 6 months, by were alive. In terms of long-term survival, there was no ery in both groups (Log Rank p =0.919, Breslow p =0.834, analysis. The 1-, 5-, and 8-year survival rates were 85.8%				
Con	clusions:		survival benefit when compared to replacement. Hence in patients with reasonable suspicion that regurgitation				
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Background

The indication for tricuspid valve (TV) surgery is mostly considered at the time of mitral or aortic valve surgery and most often because of tricuspid regurgitation (TR). Concomitant TV repair in this scenario does not significantly increase crossclamping time and may reduce the morbidity associated with reoperation for isolated TR, leading to an increase in operative correction procedures over the past decade [1,2]. Hence most follow-up studies are concerned with comparing the difference between tricuspid annuloplasty/repair (TAP) and tricuspid valve replacement (TVR). However, few studies have investigated the outcome in patients with the need of tricuspid surgery because of isolated TV pathology [3,4]. The etiologies here for consist mostly for annular dilation, infective endocarditis (IE), congenital (structural) pathology of the leaflets, post-interventional destruction (e.g., post-biopsy or pacemaker implantation), and rarely rheumatic disorder. Severe TR is associated with poor prognosis and surgical treatment is known to provide a superior outcome than conservative medical treatment [5]. We present our follow-up data to evaluate long-term outcomes in patients undergoing isolated TAP or TVR.

Material and Methods

Study population

Patients were selected through review of our prospectively maintained TV registry. Data analysis was handled anonymously and without additional patient contact or examinations, hence our institutional review board waived the need of informed consent. Between February 1995 and June 2011, 109 consecutive patients underwent isolated TV procedures at University Hospital of Heidelberg: 41 (37.6%) received tricuspid valve annuloplasty/repair (TAP) and 68 (62.3%) underwent tricuspid valve replacement (TVR). The majority of patients suffered from tricuspid regurgitation due to structural pathology, annular dilation, or infective endocarditis. Detailed breakdown of pathologies revealed 32 patients suffered from isolated tricuspid endocarditis while 16 patients had prior pacemaker implantation causing valve defects in form of microbial vegetation, structural damage, or stenosis, 37 patients had tricuspid insufficiency (TI) because of annular dilation caused by right ventricular dilation or DCMP, while four patients had prior aortic or mitral valve replacements and developed severe TI in the further clinical course, whereas six patients suffered TI after cardiac transplantation. Six patients had congenital anomalies such as Ebstein's malformation, RVOTO, or ASD-II. Furthermore, four patients suffered from mechanical TK-prosthesis thrombosis, four patients had primary tricuspid stenosis and one patient developed severe insufficiency after blunt trauma causing contusio cordis. Comprehensive data

such as patient demographics, cardiovascular risk factors, cardiac function assessed by two-dimensional echocardiograms, intraoperative characteristics as well as the postoperative outcomes including long-term survival were compared (Tables 1–3).

Tricuspid valve annuloplasty/repair (TAP)

In most cases the procedures were performed through median sternotomy using cardio-pulmonary bypass (CPB) with cardioplegic arrest and standard cannulation of ascending aorta and venae cava superior and inferior (bicaval cannulation). However, some cases were performed through a transversal (n=1, 2%), parasternal (n=1, 2%), or antero-lateral (n=3, 7%) access. An alternative cannulation strategy through the femoral vessels was used in three patients (7%). There were several techniques applied depending on the morphological abnormalities of the valve including standard ring annuloplasties (n=11), ring reconstructions or tightening using the DeVega technique (n=6), valvuloplasty with pericardial patching or bicuscpidalization (n=20), commissurotomy (n=2), papillary muscle or chordae plasty or combination of them. The chest was closed in routine manner.

Tricuspid valve replacement (TVR)

In most cases the procedures were performed through median sternotomy except for two cases (3%) where the anterolateral access was used. All procedures were done under CPB with cardioplegic arrest and ascending aorta cannulation as well as bicaval cannulation. An alternative cannulation strategy through the femoral vessels was used in seven patients (10%). The choice of prosthesis was primarily based on patients' age, hence patients up to the sixth decade received mechanical and patients in their sevenths decade received biological prosthesis. Exceptions were decided in respect to patients' comorbidities, such as contraindications for continued anticoagulation and prosthesis were chosen in consent with the patient.

Anticoagulation protocol

Intravenous heparin infusion was started on the first postoperative day. A target activated partial thromboplastin time of 50–70 seconds was selected, which was measured. After removal of chest drains, warfarin medication was initiated to keep the international normalized ratio (INR) between 2.5–3.5. The heparin infusion was not discontinued before the target INR was achieved. Patients receiving biological prosthesis or annuloplasty/repair discontinued warfarin three months after surgery, if no other reason for anticoagulation existed. Patients with mechanical prosthesis needed to continue lifelong anticoagulation. Table 1. Patient's demographics and preoperative baseline characteristics.

	т	AP		TVR	p-value
Demographic data					
Age (years)	50.7	±19.4	55	.7±15.9	0.163
Female	18	(43.9%)	37	(54.4%)	0.305
Height (cm)	172.9	±8.9	170	.2±9.1	0.135
Weight (kg)	73.5	±18.5	72	.4±21.2	0.773
BMI	24.5	±5.2	24	.9±6.9	0.713
Angina					
Exertional angina	4	(9.8%)	13	(19.1%)	0.283
Rest angina	0		5	(7.4%)	0.157
Unstable angina	0		2	(2.9%)	0.534
Dyspnea					
Exertional dyspnea	26	(63.4%)	58	(85.2%)	0.053
Rest dyspnea	9	(22.0%)	22	(32.4%)	0.382
Liver enlargement	9	(22.0%)	21	(30.9%)	0.505
Pulmonary congestion	6	(14.6%)	14	(20.6%)	0.613
Pulmonary edema	2	(4.9%)	8	(11.8%)	0.324
Peripheral edema	15	(36.6%)	26	(38.2%)	1.000
Previous MI	1	(2.4%)	5	(7.4%)	0.408
Previous syncope	3	(7.3%)	5	(7.4%)	0.741
Previous embolism	3	(7.3%)	3	(4.4%)	0.602
Previous decompensation	7	(17.1%)	19	(27.9%)	0.252
Previous CPR	1	(2.4%)	0		0.349
Previous CVA	0		7	(10.3%)	0.089
Mechanical ventilation	2	(4.9%)	0		0.119
IABP	0		1	(1.5%)	1.000
Cardiovascular risk factors					
Family history	11	(26.8%)	9	(13.2%)	0.178
Hyperlipidemia	10	(24.4%)	21	(30.9%)	0.559
Hyperuricemia	7	(17.1%)	8	(11.8%)	0.585
Hypertension		(51.2%)	30	(44.1%)	0.430
Smoking	19	(46.3%)	27	(39.7%)	0.548
Pulmonary hypertension	0	(19.5%)	20	(29.4%)	0.466

Table 1 continued. Patient's demographics and preoperative baseline characteristics.

		ТАР		TVR	p-value
Diabetes mellitus					0.660
Туре І	1	(2.4%)	6	(8.8%)	
Type II not insulin dependent	3	(7.3%)	5	(7.4%)	
Type II insulin dependent	3	(7.3%)	4	(5.9%)	
Medication					
Oral nitrates	2	(4.9%)	4	(5.9%)	1.000
i.v. nitrates	1	(2.4%)	2	(2.9%)	1.000
Beta-blockers	13	(31.7%)	31	(45.6%)	0.165
ACE-antagonists	19	(46.3%)	23	(33.8%)	0.226
Ca-antagonists	5	(12.2%)	11	(16.2%)	0.781
Glycosides	11	(26.8%)	28	(41.2%)	0.152
Diuretics	24	(58.5%)	43	(63.2%)	0.216
Inotropic agents	3	(7.3%)	2	(2.9%)	0.359
Antiarrhythmic agents	4	(9.8%)	5	(7.4%)	0.726
Vasodilators	3	(7.3%)	1	(1.5%)	0.149
Steroids	1	(2.4%)	0		0.354
Immunosuppressive drugs	2	(4.9%)	2	(2.9%)	0.612
Antibiotics	18	(43.9%)	20	(29.4%)	0.149
Bronchodilators	1	(2.4%)	4	(5.9%)	0.653
Anticoagulants	16	(39.0%)	33	(48.5%)	0.430
Previous surgery					
Coronary surgery	2	(4.9%)	7	(11.1%)	0.481
Aortic valve surgery	2	(4.9%)	4	(5.9%)	1.000
Mitral valve surgery	3	(7.3%)	9	(13.2%)	0.530
Tricuspid valve surgery	0		10	(14.7%)	0.013
Pulmonary valve surgery	0		1	(14.7%)	1.000
Surgery for congenital vitium	1	(2.4%)	6	(8.8%)	0.257
Cardiac transplantation	3	(7.3%)	6	(8.8%)	1.000
Infection	21	(51.2%)	22	(32.4%)	0.068
PVD	0		4	(5.9%)	0.292
i.v. drug abuser	1	(2.4%)	2	(2.9%)	1.000
Malignancy	2	(4.9%)	4	(5.9%)	1.000

TAP – tricuspid valve annuloplasty; TVR – tricuspid valve replacement; BMI – body mass index; MI – myocardial infarction; CPR – cardiopulmonary resuscitation; CVA – cerebrovascular event; IABP – intra-aortic balloon pump; ACE – angiotensin-converting enzyme; PVD – peripheral vascular disease.

Table 2. Preoperative cardiac function and disease classification.

	1	TAP	1	VR	p-value
LV ejection fraction					0.640
>55%	34	(83%)	53	(78%)	
41–55%	4	(10%)	10	(15%)	
26–40%	1	(2%)	3	(4%)	
<25%	2	(5%)	2	(3%)	
Hypertrophy	16	(39%)	31	(46%)	1.000
Dilation	14	(34%)	33	(49%)	0.401
Anterior hypokinesia	6	(15%)	9	(13%)	1.000
Posterior hypokinesia	1	(2%)	2	(3%)	1.000
Septum hypokinesia	6	(15%)	11	(16%)	1.000
Apical hypokinesia	6	(15%)	11	(16%)	1.000
Anterior akinesia	1	(2%)	0		0.333
Posterior akinesia	1	(2%)	2	(3%)	1.000
Septum akinesia	1	(2%)	0		1.000
Septum aneurysm	1	(2%)	2	(3%)	1.000
Apical aneurysm	2	(5%)	0		0.333
RV function					0.73
Mild	12	(30%)	31	(45%)	
Moderate	23	(56%)	24	(35%)	
Severe	6	(14%)	13	(20%)	
PH (mean >40 mmHg)	23	(57%)	31	(46%)	1.000
Tricuspid regurgitation					0.721
Severe	35	(86%)	31	(46%)	
Moderate	4	(10%)	12	(17%)	
Mild	2	(4%)	1	(2%)	

LV – left ventricle; RV – right ventricle; PH – pulmonary hypertension.

Statistics

All data were processed with the Statistical Package for Social Sciences (SPSS Inc., Chicago, IL, USA) and are given as continuous or categorical variables. Continuous data were shown as mean \pm standard deviation and analyzed with the Student *t*-test. Pearson's χ^2 or Fisher exact tests were utilized for categorical data. Kaplan-Meier survival estimation was used for patient's survival investigation. The groups were compared using Log rank, Breslow and Tarone-Ware tests. A value of *p*<0.05 was considered statistically significant.

Results

Preoperative characteristics are shown in Table 1. There were no statistically significant differences between the two groups regarding age (50.7±19.4 years in the TAP group vs. 55.7±15.9 years in the TVR group, p=0.163), gender distribution (43.9% vs. 54.4% female patients in the TAP and TVR groups, respectively, p=0.305), and body mass index (BMI) (24.5±5.2 in the TAP group vs. 24.9±6.9 in the TVR group, p=0.713). Moreover, there were no significant differences regarding preoperative angina or dyspnea status, liver, renal and pulmonary function, infections, hemodynamic status, and cardiovascular risk

Table 3. Intraoperative data.

	T	'AP	TVR		p-value
Urgency of procedure					0.93
Elective	26	(63%)	45	(66%)	
Urgent	9	(23%)	16	(24%)	
Emergency	5	(12%)	5	(7%)	
Salvage	1	(2%)	2	(3%)	
Cross clamp time in min	28.1	L±23.4	43	1±38.4	0.14
Type of prosthesis					
Biological			36	(53%)	
Mechanical			32	(47%)	

Urgent procedure: requiring surgical intervention within 24 h; emergency procedure: requiring immediate surgical intervention within 2 hours because of increasing hemodynamic instability, salvage procedure: manifest significant hemodynamic instability.

factors (Table 1). Furthermore, the differences between the two groups in terms of previous coronary (p=0.481), aortic valve (p=1.000), mitral valve (p=0.530), or pulmonary valve surgery (p=1.000) was not statistically significant, whereas patients in the TVR group had a significantly higher rate of previous tricuspid valve repair (p=0.013). Preoperative cardiac function assessed by transesophageal echocardiography was also similar and the severity of the disease reflected by pulmonary hypertension and right ventricular function did not differ significantly in both groups (Table 2). Additionally, there was no significant difference in the intraoperative characteristics in terms of cross clamping time or urgency of the procedure (Table 3).

Postoperatively, patients from both groups had similar left ventricular and right ventricular function assessed by two-dimensional echocardiography. There were no statistically significant differences in terms of postoperative arrhythmias, the need for intra-aortic balloon pump or ventricular assist device implantations. Both groups were comparable in terms of inotropic support requirement, postoperatively, except for noradrenaline support which was needed more frequently in the replacement group (50.0% in the TVR group *vs.* 24.4% in the TAP group, p=0.014). The distribution of postoperative complications such as renal failure requiring conservative treatment, dialysis/hemofiltration, pleural effusions requiring negative balance, or drainage was similar in both groups (Table 4).

Early survival at 30 days after surgery was 97.6% in the TAP group and 91.1% in the TVR group. After 6 months, 89.1% in the TAP group and 87.8% in the TVR group were alive. In terms of long-term survival, there was no further mortality observed after one year post-surgery in both groups (Log Rank p=0.919, Breslow p=0.834, Tarone-Ware p=0.880) in the Kaplan-Meier

survival analysis. The 1-, 5-, and 8-year survival rates were 85.8% for TAP and 87.8% for TVR group (Figure 1).

Discussion

Only a few studies have investigated the outcome of isolated tricuspid surgery. Significant tricuspid disease requiring surgical correction is a challenging pathology, demanding critical decision-making regarding reconstruction versus replacement and timing of the procedure [6,7]. The reasons here for are the reported high mortality and the fact that TR is relatively well tolerated, even if severe. Hence patients present with complex morbidity such as manifest congestive heart failure, severe cardiac dilation, pulmonary hypertension or endocarditis. The presence of these serious preoperative conditions present highly confounding factors altering postoperative outcomes significantly and probably accounting for previously reported poor outcomes. Currently the indication for surgical correction vs. replacement remains at the surgeon's discretion, although recently there is a growing tendency towards reconstructive strategies [8]. Theoretical benefits are the avoidance of inserting a rigid prosthesis into the thin-walled, low-pressure right ventricle, which can result further deterioration of right-ventricular dysfunction [9-11]. However, early and longterm outcomes after TAP show high rates of recurrent TR despite the use of annuloplasty rings [12]. In a large series reported by the Toronto group and the Cleveland Clinic group the recurrence of moderate to severe regurgitation was as high as 38% and 20% respectively [9,12]. This can account for diminished survival because deteriorating right ventricular function and redo tricuspid valve surgery in that case is associated with higher morbidity and mortality [13]. Hence outcome analysis of that patient cohort is of great interest. However, most

Table 4. Postoperative characteristics.

		TAP TVR		TVR	p-value	
Left ventricular function					0.534	
Mild	4	(9.8%)	22	(32.4%)		
Moderate	4	(9.8%)	10	(14.7%)		
Severe	0		1	(1.5%)		
Right ventricular function					0.879	
Mild	2	(4.9%)	8	(11.8%)		
Moderate	6	(14.6%)	23	(33.8%)		
Severe	0		1	(21.6%)		
Rhythm disturbance						
AV-Block	4	(9.8%)	0		0.761	
Ventr. extrasystole	1	(2.4%)	12	(17.6%)	1.000	
Supr. extrasystole	9	(22.0%)	7	(10.3%)	0.811	
Atrial fibrillation					0.820	
Paroxysmal	4	(9.8%)	9	(13.2%)		
Persistent	3	(7.3%)	5	(7.4%)		
Permanent	1	(2.4%)	4	(5.9%)		
Low output syndrome					0.402	
Conservative treatment	0		1	(1.5%)		
IABP	1	(2.4%)	4	(5.9%)		
Assist device	1	(2.4%)	0			
Inotropic drugs						
Noradrenaline	10	(24.4%)	34	(50.0%)	0.014	
Dopamine	7	(17.1%)	14	(20.6%)	0.804	
Dobutamine	24	(58.5%)	51	(75.0%)	0.086	
Adrenaline	11	(26.8%)	14	(20.6%)	0.189	
Renal failure					0.477	
Conservative treatment	7	(17.1%)	18	(26.5%)		
Dialysis	0		1	(1.5%)		
Hemofiltration	2	(4.9%)	2	(2.9%)		
Respiratory insufficiency					0.233	
Forced respirat. therapy	10	(23.3%)	26	(38.2%)		
Re-intubation	0		1	(1.5%)		
Pleural effusion					0.693	
Conservative treatment	0		1	(1.5%)		
Drainage	1	(2.4%)	1	(1.5%)		
Coagulation disorder	1	(2.4%)	6	(8.8%)	0.412	
Myocardial infarction	0		1	(1.5%)	1.000	

AV – atrioventricular; IABP – intra-aortic balloon pump.

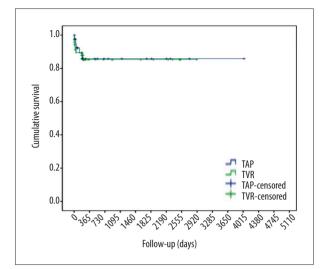


Figure 1. Kaplan-Meier Survival Curve for tricuspid valve repair (TAP) vs. tricuspid valve replacement surgery (TVR): Log Rank p=0.919, Breslow p=0.834, Tarone-Ware p=0.880.

studies include a variety of patients in their analysis, especially those in need of multiple cardiac interventions. In this study however, only isolated tricuspid valve pathologies were analyzed, thus our patient cohort was more uniform concerning preoperative characteristics and direct comparison between the test groups is feasible without the need for statistical alterations such as pair-matched analysis or propensity score matching. Age, gender, BMI, and the severity of the disease were comparable in both groups. Known preoperative risk factors like right ventricular function and pulmonary hypertension were also similar, as well as known operative confounding factors like cross clamping time or the urgency of the procedure.

The overall results of this study show that the outcome after tricuspid surgery is acceptable with 30-day survival of 97.6% in the TAP group and 91.1% in the TVR group and long-term survival of 85.8% in the TAP and 87.8% in the TVR group. These rates are in accordance with, if not better, than in previous published studies [8,14,15]. However, our promising results might be overestimated by a relatively small number of

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patients and therefore lower statistical power. Nonetheless the outcomes are somewhat disappointing when compared with survival after left sided valve replacement operations, even after exclusion of patients with complex cardiac pathologies needing multiple surgical corrections. The same observations were also reported by Moraca et al. with survival rates at 1-, 5-, and 10 years of 80%, 72%, and 66% for repair and 85%, 79%, and 49% respectively for the replacement group, without statistical significance [6]. The incidence of major adverse cardiac events was also similar in both groups, as were clinical surrogate parameters, except for the use of noradrena-line: which was significantly more needed in the tricuspid replacement group. This may be due to the fact that this patient group had a longer CPB time and thus suffered more vasople-gia in the early postoperative period.

Conclusions

Thus the main finding of this study is that there is no clear benefit when comparing tricuspid repair to replacement. Furthermore, freedom from reoperation was also non-significant. Hence the growing enthusiasm towards corrective procedures needs to be put into perspective and focus needs to be shifted towards the timing of the procedure to maintain right ventricular function. Current guidelines recommend surgical intervention when patients become symptomatic [1], which might lead to significant progression of the disease and further deterioration of right ventricular function. However, it has been shown that TR reduces exercise capacity and negatively affects long-term outcome, irrespective of pulmonary hypertension or left ventricular function [16–19]. Echocardiographic parameters of right ventricular function could be consulted when determining optimal timing of surgical intervention [4]. The decision should lean towards valve replacement in patients with reasonable suspicion of recurrent regurgitation.

Disclosures

None.

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