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Propensity score matching analysis of valvesparing versus aortic root replacement in type A aortic dissection patients

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This study compared long-term survival and reintervention rates between Valve-Sparing Root Replacement (VSRR, n = 244) and Aortic Root Replacement (ARR, n = 499) in 743 patients undergoing Type A acute a ortic dissection (AAD), given the lack of prospective comparative data. Multivariable analysis is identifying advanced age, high Body Mass Index (BMI), Marfan syndrome, severe aortic regurgitation, bicuspid aortic valve, increased aortic root diameter, and reduced aortic cross-clamp time (ACC) as significant factors associated with ARR. After Propensity Score Matching (PSM), VSRR is showing significantly higher 5-year survival rates than ARR (80.2% vs. 64.1%, P = 0.001), validated by Inverse Probability of Treatment Weighting (IPTW) analysis. Reintervention rates are being found comparable, with endocarditis more prevalent in ARR and aortic regurgitation in VSRR. Subgroup analysis indicated that patients aged less than 60 years and those with a BMI greater than 24 in the VSRR group exhibited significantly improved survival probabilities compared to the ARR group. These findings support the wider utilization of valvesparing root replacement (VSRR) in appropriately selected patients, highlighting its potential advantages for suitable candidates.

The surgical management of acute type A aortic dissection remains a formidable challenge¹. Despite widespread reporting of perioperative results, there is still no unanimous agreement in the literature on the optimal proximal repair surgical technique, often relying on surgeon discretion or institutional traditions²⁻¹¹. ARR surgery, which simultaneously replaces the valve and aortic root, potentially eliminates the need for future valve repairs but poses unique challenges to patients, such as indefinite anticoagulant therapy required for mechanical valves, increasing the

risk of blood clots and bleeding¹². Conversely, biological valves do not require lifelong anticoagulants but may deteriorate over time, necessitating repeat surgery. In contrast, VSRR surgery, which focuses on replacing the aortic root while preserving the native valve, is a favorable option for patients with functional aortic valves^{13,14}. However, its comparative complexity and demand for high technical proficiency may lead to prolonged extracorporeal circulation, intensified myocardial ischemia, and increased perioperative mortality^{2,3,7,10,11}. While several studies have shown

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promising mid-term aortic valve function following VSRR surgery, the long-term prognosis remains uncertain^{2,3,9-11,15,16}.

The perioperative results data predominantly originate from single institutions, which are inherently prone to selection bias. Furthermore, many of these single-institution studies are relatively small-scale and lack crucial long-term survival data. The necessity of prospective randomized trials in comparing the efficacy of ARR and

VSRR for AAD management is paramount. However, such trials are currently nonexistent, owing to the arduous, expensive, and time-consuming nature of their conduct. In the absence of randomized data, we opted to undertake an observational study in a multicenter database to assess the effectiveness of ARR and VSRR. In this work, we addressed the patient characteristic heterogeneity between the ARR and VSRR groups by utilizing propensity score matching (PSM) to adjust for baseline disparities and minimize confounding effects^{17,18}. Furthermore, the robustness of our findings was reinforced through sensitivity analysis utilizing inverse probability of treatment weighting (IPTW).

Results

Patient characteristics

From 2013 to 2019, a total of 743 patients with acute aortic dissection (AAD) were identified. Of these patients, 499 (67.16%) underwent aortic root replacement (ARR) surgery as their primary treatment, while 244 (32.84%) underwent valve-sparing root replacement (VSRR) surgery. The demographic characteristics of the two groups differed significantly. Specifically, the mean age of patients in the ARR group was 60.09 years, with a male preponderance of 319 individuals (63.93%). In contrast, the VSRR group had a younger mean age of 54.99 years, and a slightly higher proportion of males, accounting for 171 patients (70.08%). Baseline patient characteristics are presented in Table 1. Patients undergoing ARR surgery demonstrated advanced age, elevated body mass index, and a higher prevalence of comorbidities such as Marfan syndrome, aortic regurgitation, chronic kidney disease, peripheral vascular disease, cerebrovascular disease, and bicuspid aortic valve, in comparison to those treated with VSRR, Furthermore, they exhibited a significantly larger aortic root diameter and curtate ACC (aortic cross-clamp) time, as summarized in Table 1.

Confounding factors

We conducted a comprehensive analysis using both univariate and multivariable logistic regression models to identify confounders pertaining to aortic valve replacement (ARR). As summarized in Table 2, the univariate analysis highlighted significant predictors, such as age at initial diagnosis (P < 0.001), BMI (P < 0.001), chronic kidney disease (P = 0.008), peripheral vascular disease (P = 0.032), cerebrovascular disease (P < 0.001), Marfan syndrome (P < 0.001), aortic regurgitation (P < 0.001), bicuspid aortic valve (P < 0.001), enlarged aortic root diameter (P < 0.001), and decreased aortic cross-clamp time (ACC, min).

Upon multivariable analysis, we confirmed that advanced age at diagnosis (P<0.001), increased BMI (P<0.001), Marfan syndrome (P<0.001), severe aortic regurgitation (P<0.001), bicuspid aortic valve (P<0.001), and a marked increase in aortic root diameter (P<0.001), along with reduced ACC (min), were independently and significantly associated with a heightened relevance for ARR. Notably, chronic kidney disease, peripheral vascular disease, and cerebrovascular disease did nottabfigdemonstrate independent associations with ARR and were thus excluded from the propensity score matching (PSM) analysis.

Outcomes

The standardized mean differences analysis revealed that all covariate imbalances were under 10%, indicating an acceptable matching

Table 1 | The baseline clinical characteristics of patients

Subgroup	ARR group (n = 499)		VSSR group (<i>n</i> = 244)		P- value
	n or mean	% or σ	n or mean	% or σ	
Preoperative data					
DeBakey					0.299
I	364	72.95%	169	69.26%	
II	135	27.05%	75	30.74%	
Age(years)	60.09	7.64	54.99	2.54	<0.00
BMI	26.78	3.14	25.03	3.31	0.001
Sex					0.096
Male	319	63.93%	171	70.08%	
Female	180	36.07%	73	29.92%	
Race(Asian)	499	100.00%	244	100.00%	/
Education level			,		0.532
Master degree or above	4	0.80%	0	0.00%	
Undergraduate course	56	11.22%	24	9.84%	
Junior college	106	21.24%	58	23.77%	
Secondary and high schools	146	29.26%	66	27.05%	
Junior high school and below	187	37.47%	96	39.34%	
History of smoking					0.159
One cigarette a few days	13	2.61%	6	2.46%	
One cigarette a day	15	3.01%	8	3.28%	
2-3 cigarettes a day	20	4.01%	9	3.69%	
4-10 cigarettes a day	35	7.01%	15	6.15%	
10-20 cigarettes a day	231	46.29%	106	43.44%	-
More than 20 cigarettes a day	51	10.22%	22	9.02%	
History of alcohol intake	170	34.07%	88	36.07%	0.592
Hypertension	414	82.97%	210	86.07%	0.267
Diabetes	56	11.22%	21	8.61%	0.254
Chronic kidney diseases	81	16.23%	22	9.02%	0.007
Peripheral vascular diseases	45	9.02%	11	4.51%	0.037
Cerebrovascular diseases	59	11.82%	8	3.28%	<0.00
LVEF	52.85	5.97	53.15	6.1	0.527
Marfan syndrome	34	6.81%	7	2.87%	<0.00
Malperfusion syndrome	88	17.64%	47	19.26%	0.613
Brain	12	2.40%	5	2.05%	
Coronary artery	16	3.21%	9	3.69%	
Renal	23	4.61%	10	4.10%	
Superior mesenteric artery	2	0.40%	2	0.82%	
Lower extremity	35	7.01%	21	8.61%	
Coronary Artery Stenosis					0.477
Moderate(50-70%)	18	3.61%	6	2.46%	
Severe(>70%)	17	3.41%	9	3.69%	
Aortic regurgitation					<0.00
No or Mild	82	16.43%	137	56.15%	
Moderate	219	43.89%	82	33.61%	
Severe	198	39.68%	25	10.25%	
Aortic root diameter(mm)	44.47	4.48	42.52	4.52	<0.00
Bicuspid aortic valve	65	13.03%	37	15.16%	0.001
Moderate/severe peri- cardial effusion	101	20.24%	48	19.67%	0.856
Preoperative shock(systolic blood pressure <80 mmHg)	34	6.81%	14	5.74%	0.636

Table 1 (continued) | The baseline clinical characteristics of patients

Subgroup	group ARR group (n = 499)		VSSR group (<i>n</i> = 244)		<i>p</i> . value	
	n or mean	% or σ	n or mean	% or σ		
Intraoperative data						
Operatig Time(min)	306.89	70.38	315.76	63.58	0.097	
CPBT(min)	181.59	45.05	186.96	43.27	0.122	
ACC (min)	117.56	18.21	123.88	18.68	<0.001	
Concomitant procedures						
Partial aortic arch replacement	123	24.65%	70	28.69%	0.248	
Complete aortic arch replacement	326	65.33%	164	67.21%	0.611	
CABG	15	3.01%	9	3.69%	0.661	
TVP	14	2.81%	12	4.92%	0.235	
MVP	17	3.41%	13	5.33%	0.143	
Blood transfusion						
Erythrocytes (U)	4.17	2.19	3.88	2.01	0.084	
Cryoprecipitate (U)	0.57	0.47	0.56	0.46	0.914	
Plasma (U)	556.99	341.26	596.31	350.44	0.144	
Postoperative characteristic	s and Outo	omes				
Neurological deficits	73	14.63%	30	12.30%	0.43	
Renal insufficiency	133	26.65%	59	24.18%	0.532	
Low cardiac output syndrome	12	2.40%	10	4.10%	0.249	
Tracheostomy	77	15.43%	39	15.98%	0.831	
Sepsis	67	13.43%	28	11.48%	0.485	
Hepatic dysfunction	115	23.05%	58	23.77%	0.854	
Respiratory dysfunction	39	7.82%	17	6.97%	0.768	
Reintubation	32	6.41%	13	5.33%	0.626	
Stroke	19	3.81%	8	3.28%	0.717	
Endocarditis	11	2.20%	6	2.46%	0.827	
Reintervention for	70	14.03%	41	16.80%	0.319	
Spontaneous hemorrhage	10	2.00%	10	4.10%		
Distal anastomotic fistula	9	1.80%	2	0.82%		
Endocarditis	36	7.21%	3	1.23%		
Aortic regur- gitation(Moderate or Severe)	12	2.40%	18	7.38%		
Aortic stenosis(Moderate or Severe)	0	0.00%	2	0.82%		
Recurrent aortic dissection	3	0.60%	6	2.46%		
ICU stay (days)	9.29	10.50	8.84	9.84	0.595	
Hospitalization time (day)	20.68	9.84	19.52	9.73	0.129	
Death rate						
In-hospital mortality	38	7.62%	26	10.66%	0.165	
Overall mortality	152	30.46%	62	25.41%	0.093	

 σ standard deviation, BMI Body Mass Index, LVEF Left ventricular ejection fraction, CPBT Cardiopulmonary bypass time, ACC aortic cross-clamp, CABG Coronary artery bypass grafting, MVP Mitral valvuloplasty, TVP Tricuspid valvuloplasty.

process. Further examination of the propensity score distributions before and after matching showed a more uniform distribution post-matching, suggesting enhanced comparability between groups (Fig. 1 and Supplementary Table 1). After mitigating potential biases through PSM analysis, patients undergoing VSRR exhibited a higher incidence of low cardiac output syndrome, hepatic dysfunction, root aneurysm requiring reintervention, spontaneous hemorrhage, and moderate to

severe aortic stenosis compared to ARR patients. Additionally, VSRR patients had higher in-hospital and overall mortality rates than ARR patients (Table 3).

Survival analysis

Initially, survival analyses conducted on the unmatched cohorts of patients undergoing valve-sparing root replacement (VSRR) and aortic root replacement (ARR) did not yield statistically significant differences in survival outcomes (Fig. 2). However, upon application of propensity score matching, a distinct advantage in survival emerged for the VSRR group compared to the ARR group, with a *P*-value of 0.001 indicating statistical significance (Fig. 3). More precisely, the 5-year overall survival (OS) rate for patients in the VSRR group stood at 80.2%, which was notably and significantly higher than the 64.1% observed in the ARR group, as evident from the data presented in Table 4. This disparity in survival rates was further corroborated by the inverse probability of treatment weighting (IPTW) analysis, which robustly demonstrated an association between VSRR and increased survival, as depicted in Fig. 4.

Subgroup analysis of the impact of potential confounding variables on treatment outcomes

Subgroup analysis revealed that, within the subgroup of patients aged <60 years and those with a BMI exceeding 24, the VSRR group demonstrated a significant improvement in survival probability compared to the ARR group. Specifically, for patients under 60 years old, the hazard ratio was 0.638 (95% CI: 0.413 to 0.987) with a p-value of 0.042. For individuals with a BMI between 23.9 and 28, the hazard ratio was 0.494 (95% CI: 0.267 to 0.913), yielding a p-value of 0.024. Furthermore, in patients with a BMI g>28, the hazard ratio was 0.454 (95% CI: 0.213 to 0.969), with a p-value of 0.041. Additionally, an examination of heterogeneity across subgroups was conducted. The results indicated that the survival probability improvement in the VSRR group compared to the ARR group was statistically significant across subgroups with varying degrees of aortic regurgitation (P for interaction = 0.031), as depicted in Supplementary Table 2.

Discussion

The current study uniquely aims to bridge the knowledge gap regarding the clinical merits of VSRR compared to ARR. By integrating cohorts from three prominent aortic centers, our findings offer three significant insights. Firstly, after adjusting for baseline differences through propensity score matching (PSM), VSRR exhibited a superior survival rate compared to ARR. Further analysis revealed that, within specific subgroups, VSRR demonstrated significantly improved survival probabilities compared to ARR. Specifically, patients aged <60 years and those with a BMI >24 in the VSRR group showed higher survival rates. Secondly, it is noteworthy that the rate of reintervention following VSRR remains relatively high. Finally, as a large retrospective study conducted in an Asian region, we believe that our findings provide crucial new data to support the field.

Despite ARR's position as the primary treatment for AAD, the utilization of VSRR has been steadily increasing^{1,2,19}. Nonetheless, to date, no randomized controlled trials comparing survival rates between these two procedures have been reported. Although observational studies are not a perfect substitute for randomized clinical trials, they serve as a valuable complement to address knowledge gaps and inquiries that may not be feasible or addressed in randomized trials. However, direct comparisons between VSRR and ARR are hampered by significant selection bias, as VSRR tends to be performed in younger, healthier patients, whereas ARR is more widely applicable, encompassing older and sicker patients. These limitations are evident in the significant differences in baseline characteristics observed in unmatched comparisons. To address these limitations, propensitymatched analyses were conducted on a multicenter database

Table 2 | Logistic regression analysis of confounding factors for the Selection of Aortic Root Replacement (ARR)

Subgroup	Univariate		Multivariate		
	OR (95% CI)	P-value*	OR (95% CI)	P-value	
DeBakey					
1	REF				
II	0.836(0.597-1.170)	0.295			
Age(years)	1.247(1.195-1.301)	<0.001	1.238(1.179-1.299)	<0.001	
BMI	1.178(1.121-1.238)	<0.001	1.234(1.150-1.324)	<0.001	
Sex					
Male	REF				
Female	0.757(0.544-1.052)	0.097			
Race(Asian)	1	/			
Education level	1.043(0.899-1.210)	0.577			
Master degree or above				,	
Undergraduate course					
Junior college					
Secondary and high schools					
Junior high school and below					
History of smoking					
One cigarette a few days	0.793(0.290-2.170)	0.651			
One cigarette a day	0.916(0.372-2.259)	0.849			
2-3 cigarettes a day	0.773(0.335-1.781)	0.546			
4-10 cigarettes a day	0.736(0.378-1.433)	0.368			
10-20 cigarettes a day	0.788(0.549-1.132)	0.197			
More than 20 cigarettes a day	0.741(0.418-1.314)	0.305			
History of alcohol intake	1.092(0.793-1.504)	0.591			
Hypertension	1.268(0.824-1.951)	0.28		,	
Diabetes	0.745(0.440-1.261)	0.745			
Chronic kidney diseases	1.955(1.188-3.219)	0.008	1.025(0.338-3.102)	0.966	
Peripheral vascular diseases	2.100(1.066-4.135)	0.032	1.219(0.137-4.515)	0.998	
Cerebrovascular diseases	3.956(1.859-8.418)	<0.001	1.335(0.247-3.936)	0.974	
LVEF	0.992(0.967-1.017)	0.527	,		
Marfan syndrome	11.380(4.965-26.081)	<0.001	19.639(6.261-61.602)	<0.001	
Malperfusion syndrome	1.114(0.752-1.650)	0.589	,		
Brain					
Coronary artery					
Renal					
Superior mesenteric artery					
Lower extremity					
Coronary Artery Stenosis					
No or Mild	REF				
Moderate(50–70%)	0.668(0.262-1.706)	0.399			
Severe(>70%)	1.061(0.466-2.418)	0.888			
Aortic regurgitation	1.001(0.400 2.410)	0.000			
Mild	1.037(0.624-1.723)	0.888	REF		
Moderate	0.274(1.166-1.43)	<0.001	3.616(1.877-6.966)	<0.001	
Severe	3.098(2.077-4.621)	<0.001	18.712(9.554-36.648)	0.001	
Aortic root diameter(mm)	1.100(1.062-1.140)	<0.001	1.115(1.062-1.172)	<0.001	
Bicuspid aortic valve	2.115(1.385-3.229)	0.001	6.075(3.121-11.824)	<0.001	
Moderate/severe pericardial effusion	0.965(0.657-1.417)	0.856	0.070(0.121-11.024)	₹0.001	
Preoperative shock(systolic blood pressure <80 mmHg)		0.856			
· · · · · · · · · · · · · · · · · · ·	0.832(0.438-1.582)		0.070/0.005.0.007\	0.001	
ACC (min)	0.982(0.974-0.990)	<0.001	0.976(0.965-0.987)	<0.001	

CI confidence interval, σ standard deviation, LVEF Left ventricular ejection fraction, ACC aortic cross-clamp.

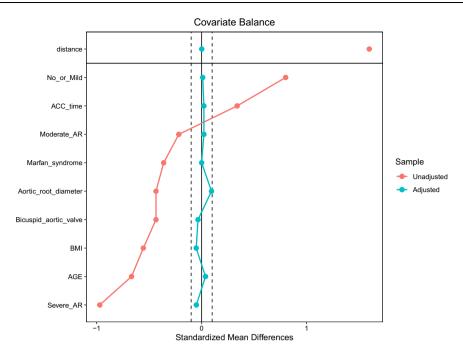


Fig. 1 | **Standardized mean difference in PSM models.** This figure presents the standardized mean differences (SMD) for various covariates related to the study, both before and after propensity score matching (PSM). Each panel represents a different variable: distance, no or mild aortic regurgitation, ACC time, moderate, Marfan syndrome, aortic root diameter, bicuspid aortic valve, BMI, AGE, and severe

AR (both unadjusted and adjusted). The SMD values indicate the degree of imbalance between the groups. Covariate balance was evaluated by analyzing the standardized difference of means, with an absolute value of the mean difference <0.1 considered indicative of successful matching. No_or_Mild no or mild aortic regurgitation, AR aortic regurgitation.

comprising 743 patients who underwent either ARR or VSRR surgery. Prior to matching, no statistical difference in 5-year survival was observed between the two groups. However, following propensity score matching, the VSRR group exhibited better survival outcomes. The results indicated that patients treated with VSRR exhibited an 80.2% greater OS benefit compared to those receiving ARR. This survival benefit remained consistentfollowing the application of inverse probability of treatment weighting (IPTW) analysis.

The present study indicates a potential survival benefit of VSRR over ARR in certain cases, a finding that has not been robustly supported in the current literature. A meta-analysis conducted by Elbatarny et al.¹⁵ revealed that, in comparison to composite valve transplantation, retaining the aortic valve is not only safe but also reduces the risk of late mortality, thromboembolism/stroke, and bleeding. This suggests that aortic valve preservation should be considered for patients with well-preserved valve morphology. Similarly, Levine et al.³ reported that VSRR is associated with improved survival and a reduced need for subsequent reinterventions compared to bioprosthetic composite valve grafts (CVG).

Conversely, Mario Gaudino¹⁰, Sergey Leontyev⁹, and Eilon Ram² failed to detect significant variations in long-term survival between VSRR and aortic root replacement (ARR). Gaudino et al. 10 reported that surgical type does not affect early or late survival, with mechanical CVG still being the durability benchmark. However, biocomposite valve grafts and valve-preserving reconstructions offer viable options for patients unsuitable for long-term anticoagulation. Leontyev et al.9 found the David operation resulted in reduced blood loss compared to the Bentall procedure, without increasing reoperation risk. This operation is preferred in patients with suitable anatomical features due to its avoidance of bleeding and other long-term artificial valve complications. Eilon Ram et al.² subsequently reported favorable early and medium-term outcomes for both VSRR and CVG in carefully selected patients, with no significant survival differences between the two approaches. Despite valve retention's higher risk of aortic insufficiency, no differences in reoperation rates were observed.

In our comprehensive analysis, we found no evidence to support the previously reported concerns of increased reoperation rates and decreased durability associated with David's surgery²⁰, as our study did not observe any notable difference in the overall reoperation rates between Aortic Root Replacement (ARR) and Valve-Sparing Root Replacement (VSRR) during the follow-up period, following propensity score matching. However, a subtle yet significant distinction emerged when we delved into the subgroup analysis of the causes of re-surgery. Specifically, endocarditis emerged as the primary reason for reintervention in the ARR group, whereas moderate to severe aortic regurgitation was the predominant factor in the VSRR cohort. Regarding other long-term outcomes, our study failed to detect any statistically significant differences in the incidence of complications such as stroke between the two surgical procedures. Nevertheless, in the immediate postoperative phase, patients undergoing VSRR exhibited a heightened susceptibility to low cardiac output syndrome and liver insufficiency. Notably, these VSRR patients were typically younger, had pre-existing moderate to severe aortic regurgitation, and frequently experienced cardiac insufficiency or even heart failure during postoperative surveillance. Furthermore, a subset of these VSRR patients engaged in high-risk behaviors, including alcoholism and inadequate sleep, during the postoperative follow-up period, which could potentially exacerbate liver damage. Our further investigation uncovered a robust correlation between these early postoperative complications and elevated early mortality rates. Additionally, our findings are consistent with previous studies conducted by other researchers, which have reported a higher prevalence of endocarditis among Bentall patients^{21,22}. This underscores the importance of maintaining a heightened level of vigilance in this patient population to mitigate potential adverse outcomes.

This study has some limitations. As a retrospective, three-center study conducted in high-volume aortic centers, the associations described here cannot be interpreted as causal. Additionally, the generalizability of the presented results may be limited. However, given the challenges of conducting randomized clinical trials to

Table 3 | Intraoperative, postoperative, and follow-up characteristics in Matched Population

Subgroup	ARR Group (n = 249)		VSRR Group (n = 147)		<i>P</i> -value
	n or mean	% or σ	n or mean	% or σ	
Intraoperative data					
Operation Time(min)	302.85	72.49	310.09	68.65	0.328
CPBT(min)	177.64	40.78	181.67	47.54	0.372
ACC (min)	117.14	17.03	120.88	19.93	0.158
Concomitant procedures					
Partial aortic arch replacement	67	26.91%	41	27.89%	0.832
Complete aortic arch replacement	182	73.09%	105	71.43%	0.72
CABG	11	4.42%	6	4.08%	0.873
TVP	5	2.01%	4	2.72%	0.646
MVP	4	1.61%	3	2.04%	0.751
Blood transfusion					,
Erythrocytes (U)	4.14	2.11	3.90	2.00	0.276
Cryoprecipitate (U)	0.59	0.47	0.60	0.48	0.77
Plasma (U)	576.71	351.54	622.45	365.33	0.223
Postoperative characteristics and outcomes					
Neurological deficits	38	15.26%	7	4.76%	0.102
Renal insufficiency	63	25.30%	36	24.49%	0.857
Low cardiac output syndrome	2	0.80%	8	5.44%	0.004
Tracheostomy	35	14.06%	24	16.33%	0.54
Sepsis	38	15.26%	15	10.20%	0.153
Hepatic dysfunction	49	19.68%	49	33.33%	0.002
Respiratory dysfunction	20	8.03%	13	8.84%	0.778
Reintubation	19	7.63%	10	6.80%	0.76
Stroke	10	4.02%	4	2.72%	0.584
Endocarditis	9	3.61%	3	2.04%	0.377
Reintervention for	29	11.65%	18	12.24%	0.873
Spontaneous hemorrhage	6	2.41%	4	2.72%	
Distal anastomotic fistula	1	0.40%	2	1.36%	
Endocarditis	16	6.43%	0	0.00%	,
Aortic regurgitation(Moderate or Severe)	6	2.41%	10	6.80%	
Aortic stenosis(Moderate or Severe)	0	0.00%	0	0.00%	
Recurrent aortic dissection	0	0.00%	2	1.36%	
ICU stay (days)	9.83	11.58	8.73	9.82	0.334
Hospitalization time (day)	20.11	9.97	19.76	9.94	0.735
Death rate					
In-hospital mortality	23	9.24%	7	4.76%	<0.001
Overall mortality	89	35.74%	29	19.73%	0.001

 $\sigma \ standard \ deviation, \ \textit{CABG} \ Coronary \ artery \ by pass \ grafting, \ \textit{MVP} \ Mitral \ valvuloplasty, \ \textit{TVP} \ Tricuspid \ valvuloplasty.$

compare these procedures, we believe our study provides an important addition to the literature. We also acknowledge that preoperative and intraoperative decision-making as well as postoperative care may vary among surgeons and institutions, potentially influencing outcomes. Thus, the influence of unmeasured confounders remains a possibility. Furthermore, the approach to ARR was based on surgeon preferences, although the techniques used for each type of surgery were consistent across the entire cohort. Nonetheless, we consider the comparison between ARR and VSRR to be highly relevant in contemporary practice.

Conclusion

Upon comprehensive analysis utilizing propensity score matching, Valve-Sparing Root Replacement (VSRR) has emerged as the superior surgical option, exhibiting improved 5-year survival outcomes in comparison to Aortic Root Replacement (ARR). This

conclusion is fortified by the additional validation achieved through the application of inverse probability of treatment weighting (IPTW) analysis. Additional subgroup analysis revealed significantly higher survival probabilities for VSRR compared to ARR, specifically in patients aged <60 years and those with a BMI >24. Collectively, our findings present persuasive arguments in favor of a more widespread adoption of VSRR for appropriate patient cohorts. This strategic shift in surgical approach has the potential to elevate patient outcomes and augment OS rates in the context of aortic root disease management.

Materials and methods

Ethics approval and Informed consent to participate

The authors bear full responsibility for the accuracy and integrity of this work, ensuring that any inquiries related to it are thoroughly investigated and resolved. The patient data used in the study was not

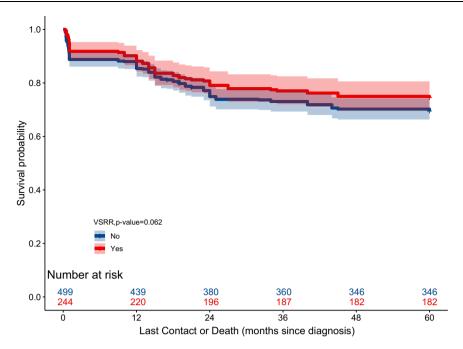


Fig. 2 | **Overall survival in the unmatched analysis of AAD patients suffering from ARR (blue) or VSRR (red).** This figure illustrates the survival probability over time for patients undergoing valve-sparing root replacement (VSRR) and aortic root replacement (ARR). The horizontal axis represents the time in months since diagnosis until death or last contact, while the vertical axis indicates the survival probability. The color coding distinguishes between the two patient groups: blue

represents the "No" group (ARR), and red represents the "Yes" group (VSRR). Additionally, the chart displays the number of patients at risk at various time points. However, survival analyses conducted on the unmatched cohorts of these patients did not yield statistically significant differences in survival outcomes, with a p value of 0.062, which is greater than the commonly accepted significance level of 0.05.

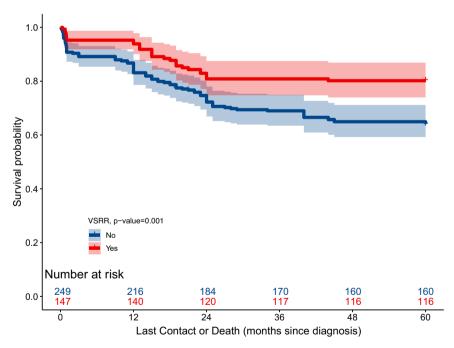


Fig. 3 | Overall survival in the propensity score matched analysis of AAD patients suffering from ARR (blue) or VSRR (red). This figure illustrates the survival probability over time in months since diagnosis until death or last contact. The horizontal axis depicts the time period, while the vertical axis indicates the survival probability. The chart employs color coding to differentiate between two

variables: blue represents the "No" group (ARR), and red represents the "Yes" group (VSRR). Additionally, the chart displays the number of patients at risk at various time points. After applying propensity score matching, a significant survival advantage emerged for the VSRR group compared to the ARR group, with a p value of 0.001, indicating statistical significance (*p* value < 0.05).

anonymized. Prior to the publication of this study, informed consent was obtained from all participants for the inclusion of their data and any accompanying images. The research adhered strictly to the principles outlined in the Declaration of Helsinki (2013 revision). The study

and its database were reviewed and approved by ethics boards, with the following reference numbers: [2024] Scientific Research No. 112, 2024LWB265, and [2024] Ethics Review No. (172). Regarding patient consent for data use, it was obtained during the patients' treatment

process. Furthermore, to maintain adherence to ethical standards, we took the additional step of re-contacting the patients or their authorized representatives to obtain specific consent for the utilization of their data in this particular study.

Data source

The data pertaining to patients with acute aortic dissection (AAD) who underwent surgical treatment at three tertiary Grade A hospitals (Fujian Medical University Union Hospital, Zhangzhou Affiliated Hospital of Fujian Medical University and the First Affiliated Hospital of Xiamen University) between February 1, 2013, and February 1, 2019, was retrieved from various databases, encompassing case management systems, imaging systems, and surgical records. The clinical information, demographic characteristics, and baseline data were extracted from electronic medical records, and the operative notes were rigorously reviewed. All imaging modalities, specifically transesophageal echocardiography (TEE) and computed tomography, were analyzed to validate the anatomical diagnosis. The study encompasses all eligible individuals within the designated study period, adhering to specific inclusion criteria: age over 18, clinical diagnosis of

Table 4 | Overall Survival with ARR Versus VSRR in AAD Patients

Year	Unmatch	red (%)	PSM	PSM		IPTW		
	ARR	VSRR	ARR	VSRR	ARR	VSRR		
1	82.8%	88.0%	83.1%	93.9%	82.4%	85.9%		
2	74.6%	77.1%	72.2%	82.9%	73.4%	83.6%		
3	72.5%	75.3%	69.0%	81.2%	69.5%	82.1%		
4	68.4%	73.4%	64.9%	80.9%	66.3%	81.2%		
5	67%	73.0%	64.1%	80.2%	65.8%	80.0%		

PSM Propensity Score Matching, σ standard deviation, IPTW Inverse Probability of Treatment Weighting.

AAD confirmed by CT angiography (CTA) or ultrasound, emergency surgery within 48 h of admission, and informed consent from patients. The exclusion criteria for the study encompassed infective endocarditis, prior AV surgery, and the utilization of non-David V or Bentall root replacements, as depicted in Fig. 5. Notably, the presence of moderate or severe aortic regurgitation did not constitute an absolute contraindication for valve-sparing root replacement (VSRR), thus allowing for its consideration in select cases.

In this study, we have comprehensively documented the key variables pertaining to patient characteristics, intraoperative specifics, and in-hospital outcomes. The following aspects are noteworthy: During the data collection process, "race" information is often self-reported by participants/respondents, which helps ensure the accuracy and authenticity of the data. The "education level" assessment may combine multiple data sources, including self-reports of participants, census data, and administrative records. In categorizing people into different categories, we follow standardized classification criteria and use objective, quantifiable indicators whenever possible. However, there are some exceptions where data is missing, specifically for EF (18 missing, 2.3%), CPB time (11 missing, 1.4%), and cross-clamp time (4 missing, 0.5%). The median follow-up duration for the entire cohort was 5.6 years, with a slight variation of 5.2 years for patients undergoing VSRR and 5.8 years for those undergoing ARR. When updated follow-up information was unavailable, patients were censored at the date of their last contact. The primary endpoint of this study was to assess all-cause mortality. Additionally, we considered any reintervention necessitated due to aortic valve-related issues as a secondary endpoint. The study design, results, and implications are summarized in Fig. 6.

Surgical technique

Although the surgical indication was individually determined by the attending surgeon, it generally adhered to the contemporary

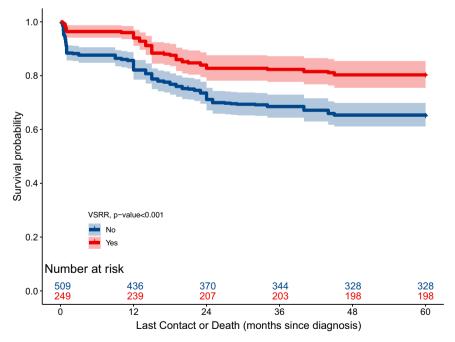


Fig. 4 | **Overall survival in the inverse probability of treatment weight-adjusted analysis in AAD patients suffering from ARR (blue) or VSRR (red).** This figure depicts the survival probability over time for two groups, distinguished by color coding. The horizontal axis represents the time in months since diagnosis until death or last contact, while the vertical axis indicates the survival probability. Blue represents the "No" group (ARR), and red represents the "Yes" group (VSRR). The

chart also displays the number of patients at risk at various time points. Upon application of the inverse probability of treatment weighting (IPTW) analysis, a statistically significant advantage in survival emerged for the VSRR group compared to the ARR group, with a p value of < 0.001 (p< 0.001), indicating a clear distinction between the two groups.

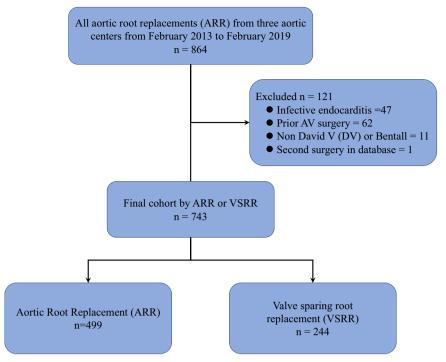


Fig. 5 | **Consort diagram of patients with acute type A aortic dissection undergoing VSRR or ARR root replacement.** A total of 864 surgeries were performed during February 2013 to February 2019. Of these, 121 surgeries were excluded for various reasons: 47 due to infective endocarditis, 62 due to prior AV surgery, 11 due to non-David V (DV) or Bentall surgery, and 1 due to being a second

surgery in the database. The final cohort for analysis, by either ARR or valve-sparing root replacement (VSRR), included 743 surgeries. Among these, 499 surgeries involved aortic root replacement (ARR) with valve replacement, while 244 surgeries involved valve-sparing root replacement (VSRR). The symbols used in the flowchart are as follows: • represents excluded cases. AV aortic valve.

guidelines published at the time of surgery^{23,24}. All surgical interventions were performed using a median sternotomy approach, central cannulation, moderately hypothermic cardiopulmonary bypass (32°C), aortic cross-clamping, and myocardial protection achieved through cold antegrade blood cardioplegia. Specifically, the David I technique¹⁴ was utilized for VSARR, while the modified Bentall technique²⁵ was adopted for ARR. In cases requiring concomitant hemiarch/transverse aortic procedures, the selection of the arterial cannulation site and the cerebral protection method was left to the discretion of the operating surgeon.

Statistical analysis

The relationships between treatment modality and patient demographic and clinical characteristics were examined using the Pearson $\chi 2$ or Fisher's exact test for categorical data and the Wilcoxon rank sum test for ordinal and continuous data. Regarding the primary outcome, two analytical models were formulated: a propensity score-matched model and an unmatched univariable analysis utilizing the Kaplan-Meier estimator of inverse probability of treatment weight (IPTW). Additionally, a propensity score model for the receipt of ARR was established via stepwise variable selection within a multivariable logistic regression framework. This model incorporated candidate variables that demonstrated significant associations with treatment modality through univariable analysis, with a threshold of P < 0.05required for initial inclusion and subsequent retention in the final model. Based on these propensity scores, a stabilized inverse probability of treatment weight was calculated. It is noteworthy that in the context of weighted analysis utilizing IPTW (Inverse Probability of Treatment Weighting), the efficiency and accuracy of statistical analysis are enhanced through the adjustment of weights. This process leads to an apparent increase in the effective sample size or information content within each group under the weighted analysis. It is crucial to emphasize that this augmentation represents a statistical increase, rather than a physical one, underscoring the refined precision and sensitivity gained through the application of IPTW^{26,27}.

To establish the propensity-score-matched model for OS, a greedy, nearest neighbor matching algorithm was employed to match patients treated with valve-sparing root replacement (VSRR) to patients treated with aortic root replacement (ARR), with a 1:2 ratio, based on their propensity scores. The maximum permissible difference in propensity scores was set at 2%. Covariate balance was evaluated by analyzing the standardized difference of means, with an absolute value of the mean difference < 0.1 considered indicative of successful matching. Subsequently, Kaplan-Meier estimators were calculated for both groups and were compared using the log-rank test. In the final OS model, IPTWadjusted Kaplan-Meier estimators were calculated for the entire patient cohort and were contrasted between treatment groups via the log-rank test. Participants were further stratified into subgroups according to specific criteria: age, BMI, degree of aortic regurgitation, presence or absence of Marfan syndrome, and bicuspid aortic valve. Cox regression analysis was subsequently conducted on the data pertaining to each subgroup. To ascertain whether there was an interaction effect between the grouping variables and the treatment factors, heterogeneity among the subgroups was assessed using a test for interaction (P for interaction).

The primary objective of this study was to compare the OS of patients with acute type A aortic dissection (AAD) treated conservatively with ARR versus those surgically treated with VSRR.OS

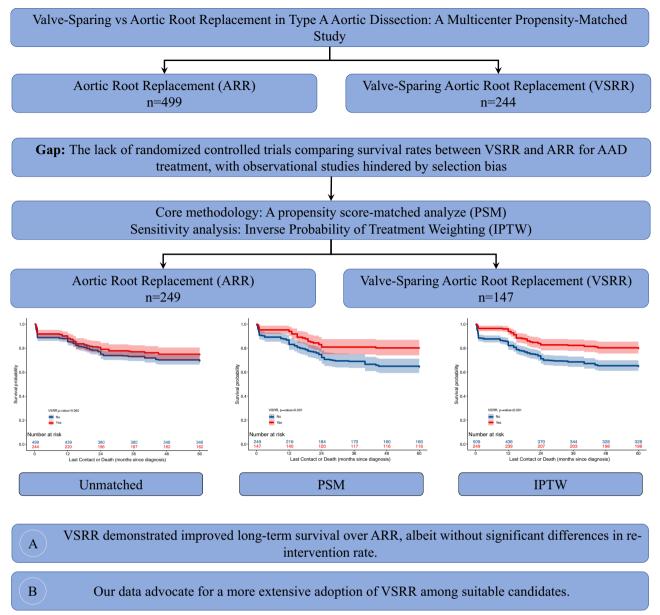


Fig. 6 | **Schematic depiction of the study cohort.** This figure illustrates a multicenter, propensity-matched study comparing survival rates between valve-sparing aortic root replacement (VSRR) and aortic root replacement (ARR) for the treatment of type A aortic dissection (AAD). The study enrolled a total of 743 patients, with 499 in the VSRR group and 244 in the ARR group. Due to the lack of randomized controlled trials comparing these two surgical approaches, observational studies have been hindered by selection bias. To address this, the core methodology employed a propensity score-matched analysis (PSM), resulting in 249

patients in the VSRR group and 147 patients in the ARR group after matching. Sensitivity analysis was conducted using inverse probability of treatment weighting (IPTW) to further validate the results. A, B highlight the key clinical implications of the study, which demonstrate that VSRR demonstrated improved long-term survival over ARR, albeit without significant differences in re-intervention rates. Our data advocate for a more extensive adoption of VSRR among suitable candidates, highlighting its potential benefits in the treatment of type A aortic dissection.

was measured from the date of diagnosis to the date of death as a result of any cause. A significance level of P < 0.05 indicated statistically significant differences, with all P-values being two-sided. All necessary calculations were executed utilizing R software, version 4.4.0, as well as SPSS version 26.

Reporting summary

Further information on research design is available in the Nature Portfolio Reporting Summary linked to this article.

Data availability

Data supporting the findings of this study are available in the article and its Supplementary information. The data underlying this article is

under restricted access due to the inclusion of confidential patient information. For any questions, please contact the corresponding author.

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Author contributions

L. Chen (First Author): Conceptualization, Methodology, Software, Investigation, Resources, Supervision, Formal Analysis, Validation Writing—Original Draft. Y.C Pan, H.J Zhang and Chunsheng Wang: Data Curation, Visualization, Investigation, Writing—Original Draft. Y Chen, Z.H Qiu and H Lu: Visualization, Writing—Review and Editing. L.W Chen (Corresponding Author): Conceptualization, Funding Acquisition, Resources, Supervision, Writing—Review and Editing.

Competing interests

The authors declare no competing interests.

Additional information

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