## 160. Machine Learning-Derived Echocardiographic Phenotypes Predict Clinical Outcome and Success Repair of Tricuspid Annuloplasty

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## **Body**

**Background:** Persistent undertreatment of tricuspid regurgitation (TR) reflects uncertainties with its quantification and management. Morphological and functional components of TR, which may account for the success of tricuspid valve intervention, have not been evaluated. This study sought to identify echocardiographic phenotypes in TR and assess their association with outcomes after tricuspid annuloplasty (TA).

**Methods:** Morphomic and functional network profiling were performed in patients undergoing TA between 2012 and 2019. Principal component (PC) analysis with varimax rotation was used to condense echocardiographic data into PCs, from which clusters were derived using K-means clustering. Clusters and PCs were correlated with clinical outcomes (composite of heart failure hospitalization and all-cause mortality) and residual TR, adjusted for comorbidities, medications, and EuroSCORE II.

**Results:** Morphomic and functional data from 290 patients (age 63±9 years, 63% female) were profiled and subsequently condensed into PCs. PC 1 described high loadings of tricuspid valve and right atrial morphology; PC 2 described high loadings of right ventricular morphology; PC 3 described high loadings of left ventricular topology. Based on these components, two clusters representing "small" (Cluster 1; n=253) and "large right atria and ventricle" (Cluster 2; n=37) phenotypes were derived (Table, Figure A).

Compared with Cluster 1, Cluster 2 was associated with a higher risk of adverse outcomes (adjusted hazard ratio 2.04; 95% Confidence Interval [CI] 1.17-3.56; P=0.012) (Figure B). Nonetheless, TR remained associated with adverse outcomes after adjusting for the clusters (adjusted hazard ratio 1.005; 95% CI 1.001-1.009; P=0.030). Cluster 2 was also associated with an increased risk of residual TR (Odds Ratio 2.31; 95% CI 1.06-5.00; P=0.034) after TA.

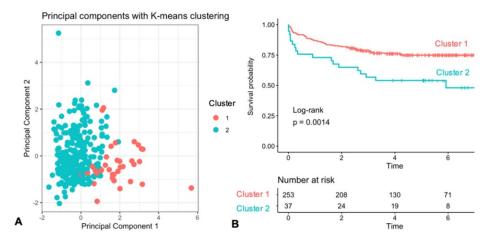
**Conclusion:** Amongst the two distinct phenotypes of TR, a "large right atria and ventricle" is associated with worse clinical outcomes and a lower chance of repair success. Such association cannot be purely attributed to cardiac morphology alone, supporting other complementary tricuspid valve features in driving poor postoperative outcomes.

**Table** 

	Cluster 1 (n=253)	Cluster 2 (n=37)	P values	T	Cluster 1 (n=253)	Cluster 2 (n=37)	P values
Demographics, Symptomatic Status, and Anthropometric Characteristics				AVR	91 (36.0)	15 (40.5)	0.721
Age, years	64.0 ± 9.5	61.9 ± 6.5	0.203	CABG	20 (7.9)	2 (5.4)	0.838
Male	94 (37.2)	13 (35.1)	0.956	LV, LA, and left-sided valvular disease			
Body mass index, kg/m <sup>2</sup>	22.74 (4.47)	22.26 (5.17)	0.557	≥ Moderate MR	152 (60.1)	7 (18.9)	<0.001
NYHA functional class III/IV	106 (41.9)	13 (35.1)	0.547	≥ Moderate MS	134 (53.0)	19 (51.4)	0.994
Cardiovascular risk factors and comorbidities				≥ Moderate AR	48 (19.0)	4 (10.8)	0.327
Hypertension	60 (23.7)	4 (10.8)	0.12	≥ Moderate AS	69 (27.3)	16 (43.2)	0.072
Diabetes Mellitus	52 (20.6)	6 (16.2)	0.692	LVEDV, mL	99.88 (40.69)	78.27 (30.63)	0.002
Dyslipidemia	54 (21.3)	2 (5.4)	0.038	LVESV, mL	42.34 (22.30)	32.32 (15.77)	0.009
Atrial fibrillation	208 (82.2)	32 (86.5)	0.682	LVEF, %	58.48 (8.83)	58.68 (8.61)	0.899
Heart failure	147 (58.1)	17 (45.9)	0.224	LA area, mm <sup>2</sup>	36.27 (17.50)	53.54 (24.90)	<0.001
Smoking	36 (14.2)	4 (10.8)	0.758	RV and RA			
Stroke	36 (14.2)	15 (40.5)	<0.001	RV basal diameter, cm <sup>2</sup>	4.62 (1.36)	5.09 (1.17)	0.052
Chronic rheumatic heart disease	174 (68.8)	30 (81.1)	0.181	RV mid-cavity diameter, cm <sup>2</sup>	3.74 (1.19)	4.25 (1.00)	0.014
Medications				RV longitudinal diameter, cm <sup>2</sup> RV end-diastolic area, cm <sup>2</sup>	5.13 (1.22)	6.18 (1.16)	<0.001
			0.079	1	16.11 (5.59)	24.70 (7.37)	
Beta blockers	` '	` '		RV end-systolic area, cm <sup>2</sup> RV sphericity index	8.71 (3.65) 4.19 (1.07)	13.39 (4.32) 5.37 (0.94)	<0.001
2	105 (41.5)	11 (29.7)	0.236	. ,	` '	` '	0.001
Calcium channel blockers	51 (20.2)	4 (10.8)	0.258	TAPSE, mm	16.48 (3.06) -16.37 (5.49)	14.68 (2.30) -12.96 (5.06)	<0.001
Diuretics	49 (19.4)	19 (51.4)	<0.001	RV global longitudinal strain, %  RA area, mm <sup>2</sup>	21.78 (8.09)	-12.96 (5.06) 53.80 (23.20)	<0.001
<u>Laboratory measurements</u>				PASP, mmHg	, ,	` ,	0.001
Hemoglobin, g/dL	12.36 (2.03)	11.08 (1.66)	< 0.001	Tricuspid Valve	48.80 (12.55)	43.84 (10.49)	0.023
Creatinine, umol/L	7.42 (2.81)	7.98 (3.58)	0.276	TR severity			<0.001
Urea, mmol/L	88.05 (27.36)	95.70 (43.10)	0.146	Moderate TR	140 (55.3)	1 (2.7)	<0.001
eGFR, mL/min/1.73m <sup>2</sup>	69.71 (20.45)	74.65 (26.80)	0.189	Severe TR	113 (44.7)	36 (97.3)	-
Valvular surgery risk-scoring systems				EROA, mm <sup>2</sup>	43.74 (24.72)	151.79 (115.42)	<0.001
EuroSCORE II	4.91 (5.82)	6.80 (4.99)	0.062	RVol, mL/beat	43.97 (23.66)	117.23 (70.99)	<0.001
STS Score	3.02 (2.71)	4.71 (3.79)	0.001	Leaflet tenting height, cm	0.60 (0.15)	0.87 (0.18)	<0.001
Surgical Details				Leaflet tenting area, cm <sup>2</sup>	0.97 (0.44)	1.77 (0.70)	<0.001
MVR	164 (64.8)	19 (51.4)	0.16	IVC diameter	2.13 (0.50)	3.05 (0.71)	<0.001
MV Repair	67 (26.5)	0 (0.0)	0.001	Right atrial pressure	8.55 (4.52)	15.95 (3.95)	<0.001
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Abbreviations: ACEI, angiotensin-converting enzyme inhibitors; AR, aortic regurgitation; ARB, angiotensin II receptor blockers; AS, aortic stenosis; AVR, aortic valve replacement; CABG, coronary artery bypass graft; CRHD, chronic rheumatic heart disease; eGFR, estimated glomerular filtration rate; EROA, effective regurgitant orifice area; EgroSCORE II, European System for Cardiac Operative Risk Evaluation II; LV, left ventricle; LA, left artium; LVEDV, left ventricular end-diastolic volume; LVEF, left ventricular ejection fraction; LVESV, left ventricular end-systolic volume; LVEF, left ventricular ejection fraction; PASP, pulmonary artery systolic pressure; RA, right artium; RV, right ventricle; RVol, regurgitant volume; STS score, Society of Thoracic Surgeons Predicted Risk of Mortality Score; TAPSE, tricuspid annular plane systolic excursion; TR, tricuspid regurgitation.

## **Figure**



**Clinical Implications:** better understand the morphological and functional components of tricuspid regurgitation and their implications on the successful repair and clinical outcomes of tricuspid annuloplasty.