Choosing Right Patient for Transcatheter Tricuspid Valve Intervention

Vinayak Bapat Chief of Cardiac Surgery Abbott Northwestern Hospital Minneapolis Heart Foundation





Disclosure Statement of Financial Interest

Within the past 12 months, I or my spouse/partner have had a financial interest, arrangement or affiliation with the organization(s) listed below.

Financial Relationship

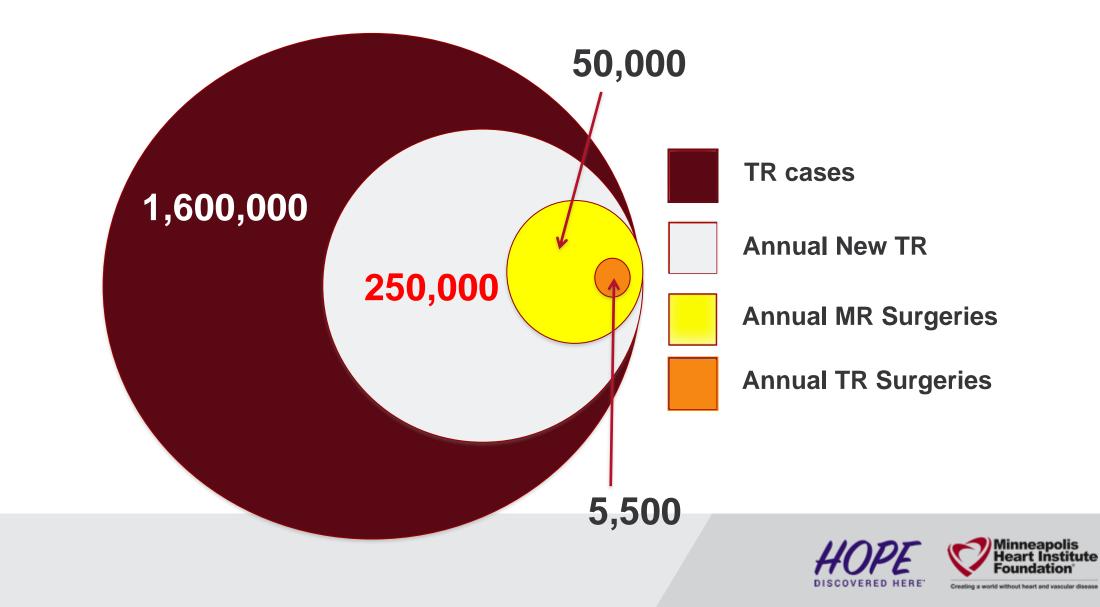
Consulting Activities

Company

Abbott, Boston Scientific, Edwards Lifesciences, Medtronic, 4C, Anteris

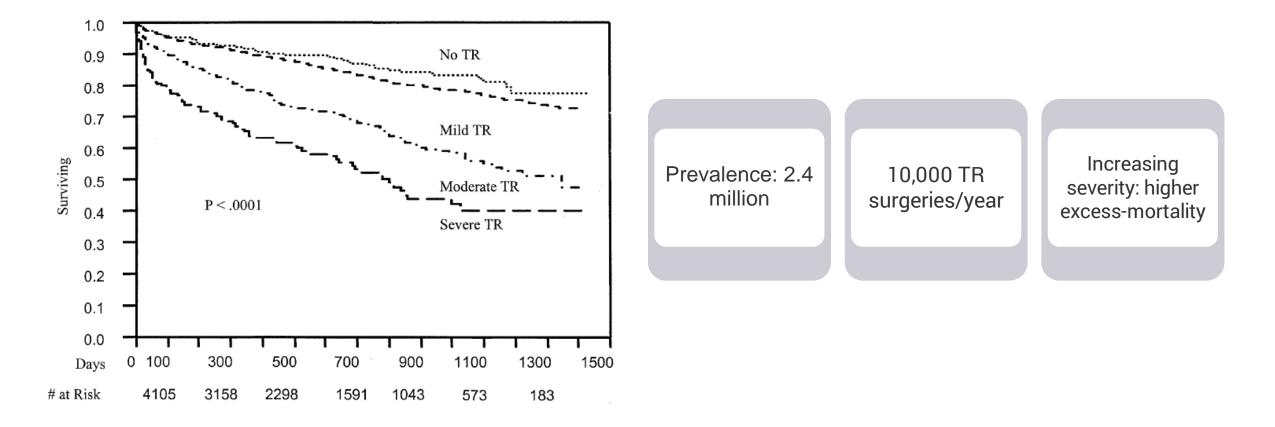


TR is currently undertreated



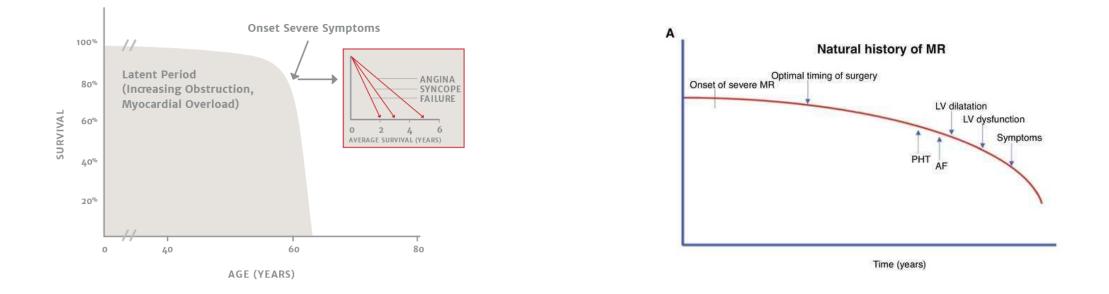
Untreated Tricuspid Regurgitation

• 40% Mortality Rate for Severe TR





Natural history of TR is not well documented



In TR the symptoms do not guide us in timing an early intervention



Challenges with Tricuspid Regurgitation

- Symptoms
 - 1. Decreased CO fatigue, decreased exercise tolerance

Considered too early for intervention

- 2. Right Heart Failure Ascites, LE edema
 - May be too late for intervention



Diagnostic Challenges with TR

- Regurgitation can be dynamic and very volume dependent
- Volume overload is well-tolerated for years
- Poor understanding about grading the severity of TR on Echo
 - Is Mild, moderate and severe a good grading?



Early Diagnosis and Follow Up are Critical to Reduce Mortality

- TR can progress:
 - From trivial/mild to moderate/severe: 5.3 <u>+</u> 2.9 yrs.
 - Follow up is critical

• Risk of death is higher with TR severity as well as in the early period after TR diagnosis

$0.1 \mod 1052$ 50.5 \bullet $3.60 (3.37 to 0.3.7 to 0.4.7 to 0.4.$		Deaths	Death rate per 100 person-years							Hazard ratio vs. no TR (95% CI)
$1-3 \mod 8$ 978 25.6 + $2.05 (1.92 to)$ $3-6 \mod 8$ 67 17.1 + $1.50 (1.40 to)$ $6-12 \mod 8$ 1166 13.9 + $1.32 (1.25 to)$ $1-2 years$ 1359 11.7 + $1.13 (1.06 to)$ >2 years 1798 12.1 + $1.10 (1.05 to)$ Moderate - - $6.10 (2.43 to)$ $0-1 \mod 8$ 866 42.2 + $2.61 (2.43 to)$ $3-6 \mod 8$ 842 31.0 + $2.61 (2.43 to)$ $6-12 \mod 8$ 886 42.2 + $2.61 (2.43 to)$ $3-6 \mod 8$ 911 21.2 + $1.54 (1.44 to)$ $1-2 years$ 1035 18.1 + $1.36 (1.27 to)$ >2 years 1239 18.4 + $1.35 (1.29 to)$ $1-3 \mod 8$ 114.3 - $46.10 (5.48 to)$ - $1-3 \mod 8$ 319 32.8 - $2.26 (2.02 to)$ - $1-2 years$ 401 26.1 - $1.84 (1.65 to)$	Mild										
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6-12 months 391 32.8 → 2.26 (2.02 to 1.10) 1-2 years 401 26.1 → 1.84 (1.65 to 1.10)	1-3 months	339	56.8				-	-		3.31 (2.94 to 3.72)	
1-2 years 401 26.1	3-6 months	319	42.0				-			2.62 (2.32 to 2.96)	
	6-12 months	391	32.8				-			2.26 (2.02 to 2.52)	
>2 years 414 22.5	1-2 years	401	26.1			-+	-			1.84 (1.65 to 2.05)	
	>2 years	414	22.5			+				1.45 (1.29 to 1.62)	
Hazard ratio vs. no TR				Hazard	1 1		2	4	8	3	

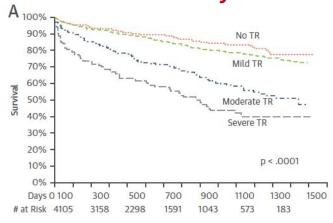


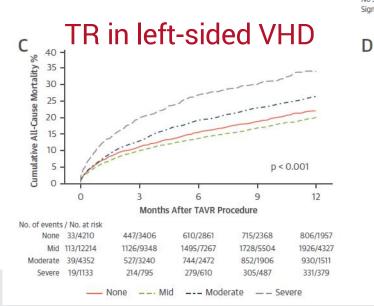
TR – Morphologic Types vary but prognosis does not

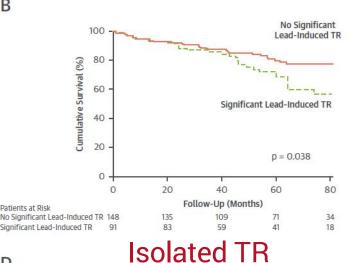
Secondary TR

B

Lead-induced TR







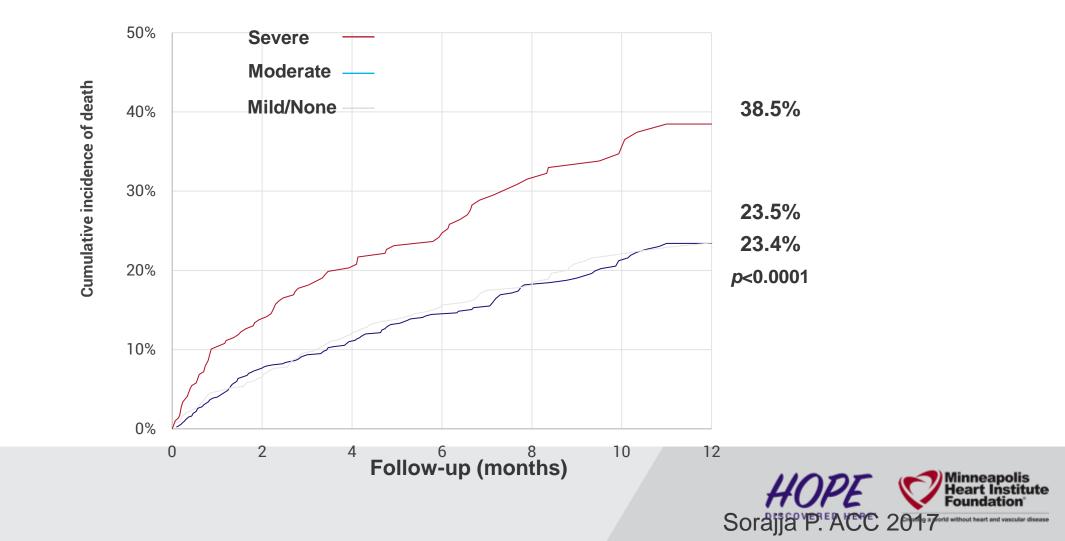
100 87 ± 2% Cumulative Survival (%) 80 70 ± 6% 60 $66 \pm 6\%$ 40 38 ± 7% 20 p < 0.001 0 -10 0 2 6 8 4 Years Number at Risk 353 308 252 194 70 31 Total 23 285 253 210 163 46 ERO <40 68 55 42 31 24 8 ERO ≥40 --- ERO ≥40 mm² ---- ERO <40 mm

As TR progresses, survival is significantly worse with 5-year survival of 50%



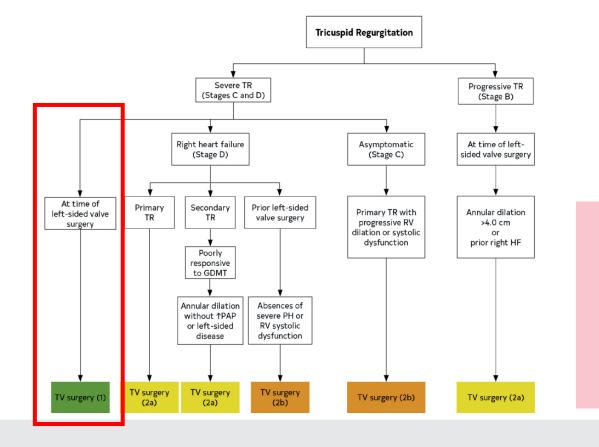
Prihadi et al JACC CV Imaging 2019

US TVT Registry: Impact of TR on MitraClip Death



Guidelines Indicate TR Surgery for Selected Patients with Isolated TR and Left-Sided Valve Lesions

• Recommendations for Intervention



• Recommendations for Medical Therapy

COR	LOE	RECOMMENDATIONS
2a	C-EO	1. In patients with signs and symptoms of right-sided HF attributable to severe TR (Stages C and D), di- uretics can be useful.
2a	C-EO	2. In patients with signs and symptoms of right-sided HF attributable to severe secondary TR (Stages C and D), therapies to treat the primary cause of HF (eg, pulmonary vasodilators to reduce elevated pulmonary
		artery pressures, GDMT for HF with reduced LVEF, or rhythm control of AF) can be useful (1,2)

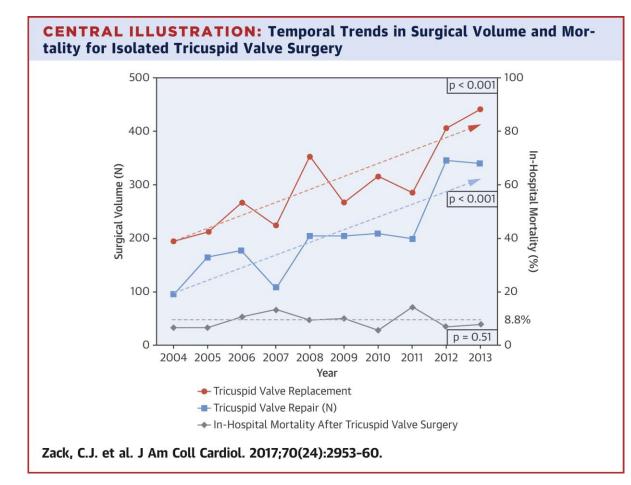
Surgery is indicated for patients with severe **TR**, but many have significant RV failure

Should we consider earlier treatment?





Isolated TV surgery



2004-2013 National Trend 5005 surgeries Steady increase in number 2004: 290 2013: 780 In hospital mortality : 8.8%

Repair: 40.8% Replacement: 59.2%

Mortality:

Repair: 5.9% Replacement (M): 13.6% Replacement (T): 9.1%



Hence



Prolonged Medical management

Late Consideration for intervention





New tricuspid therapies

Mechanism	New Technologies
Annuloplasty (Direct and Indirect)	TriAlign Millepede
Leaflet Devices	Forma MitraClip
Stented Valves in IVC/SVC	Trinity /Sapien NVT
Valve Replacement	





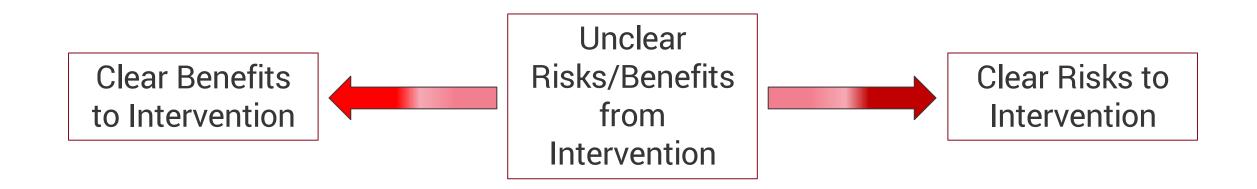
Results can be Near Perfect



But can they predict recovery and Survival benefit?

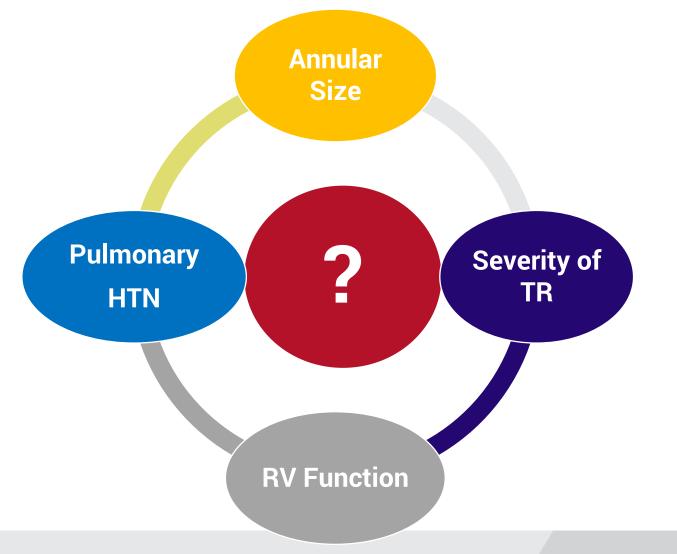


But we still need to Understand



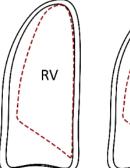


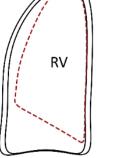
When to Intervene is bit unclear





Understanding : RV Systolic Function





Pattern 1 P Normal RV size RV Normal RV function Norma

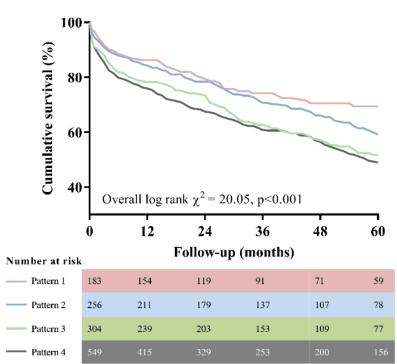
Pattern 2 RV dilation Normal RV function

Pattern 3Pattern 4Normal RV sizeRV dilationRV dysfunctionRV dysfunction

RV

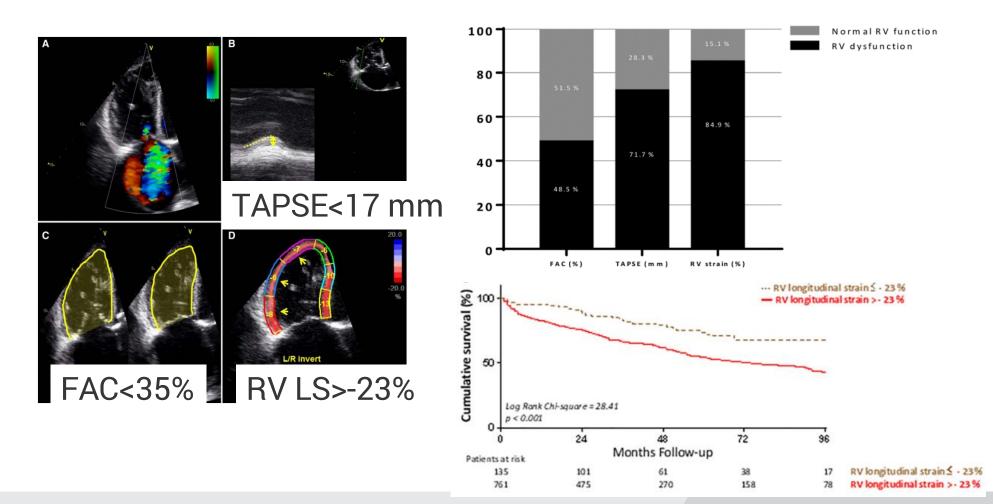
Pattern 4 RV dilation

RV





RV Systolic Function

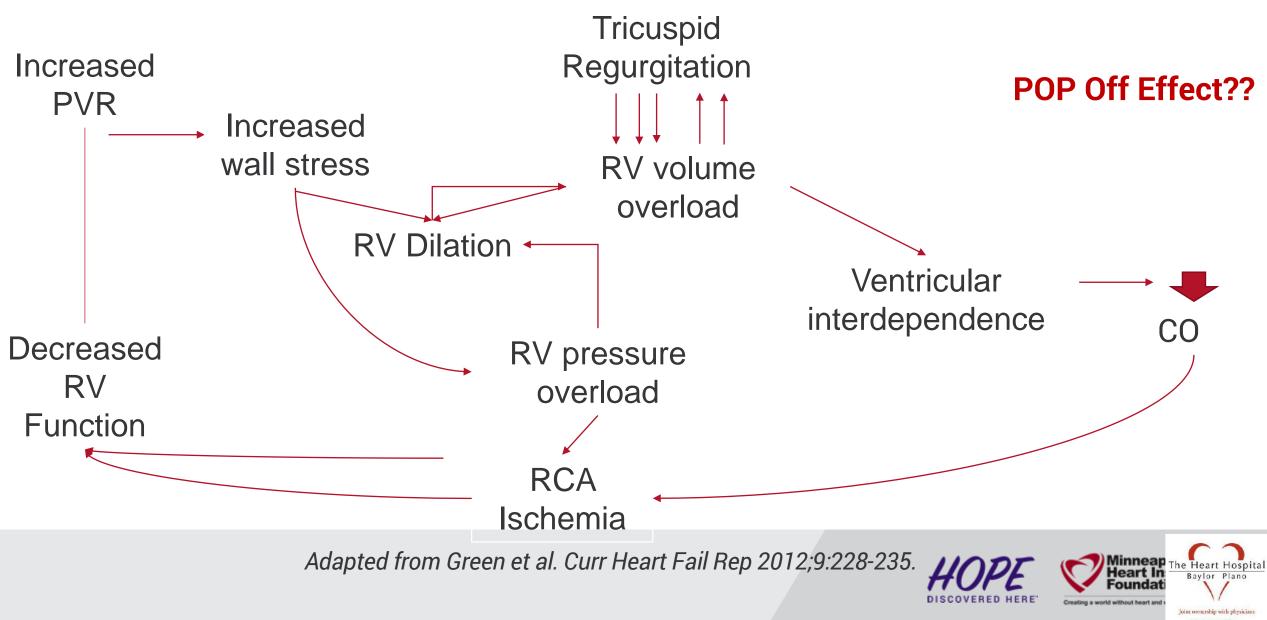




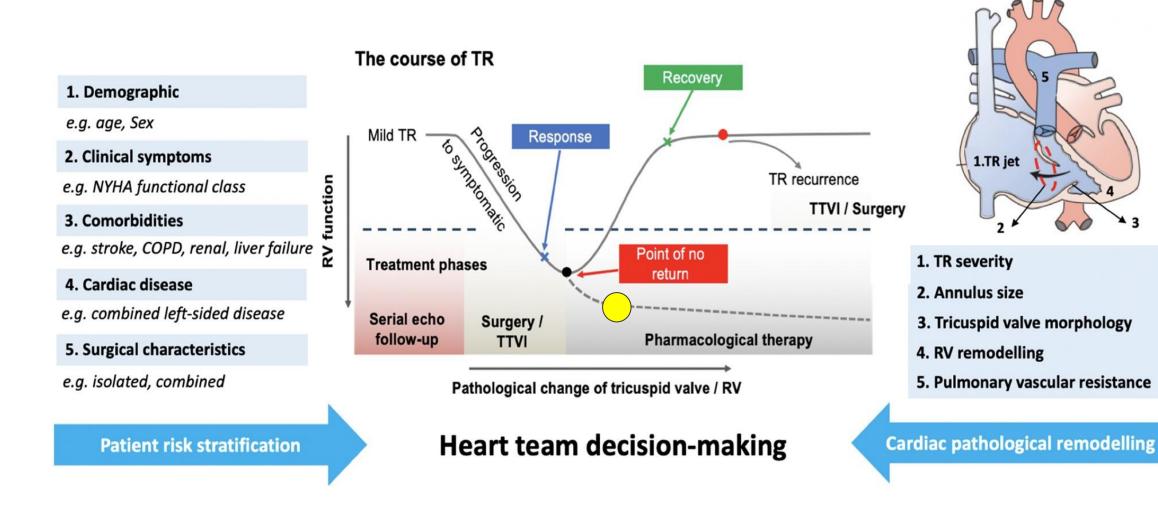
Minneapolis Heart Institute

Foundation

Pathophysiology of RV Failure



Heart Team Approach to Determine Best Timing for Treatment

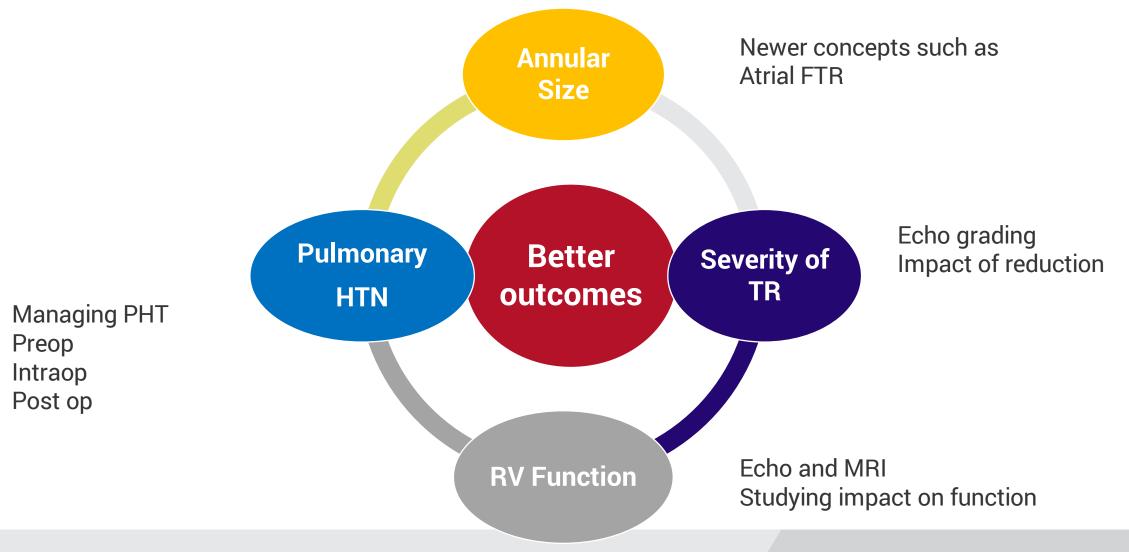


Chang CC, Veen KM, Hahn RT, et al. Uncertainties and challenges in surgical and transcatheter tricuspid valve therapy: a state-of-the-art expert review. *EUROPEAN HEART JOURNAL*. 2020;41(20):1932



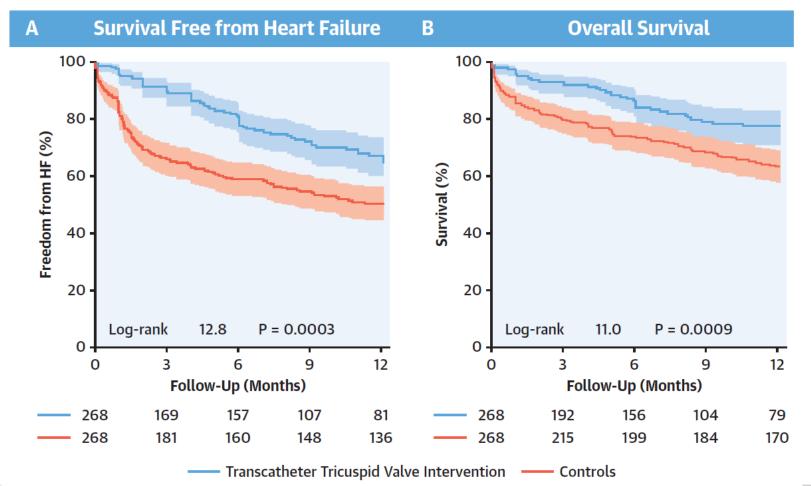


Heart Team 3.0





Significantly Improved Clinical Outcomes with Transcatheter TV Therapy versus Medical Therapy



Tamasso M. et al., 2019 Transcatheter Versus Medical Treatment of Patients With Symptomatic Severe Tricuspid Regurgitation



Improvements and Impact

- Critical piece
 - Early diagnosis and monitoring progression
 - Trigger for intervention
 - Procedure planning
 - Quantifying improvements



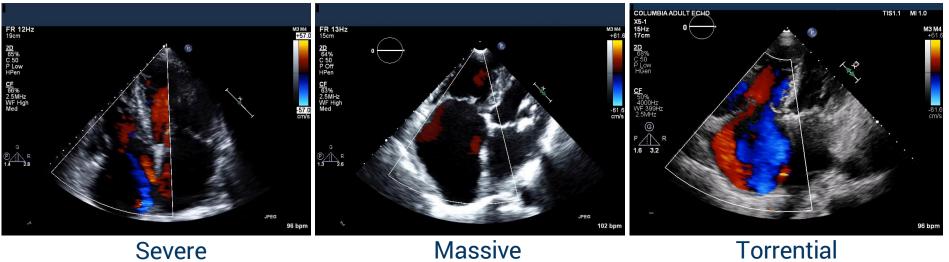
Extended Grading Scheme

Table I	Proposed	expansion of the	'Severe' grade	е
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Variable	Mild	Moderate	Severe	Massive	Torrential
VC (biplane) EROA (PISA) 3D VCA or quantitative EROAª	<3 mm <20 mm ²	3-6.9 mm 20–39 mm ²	7–13 mm 40–59 mm ² 75–94 mm ²	14–20 mm 60–79 mm ² 95–114 mm ²	≥21 mm ≥80 mm ² ≥115 mm ²

VC, vena contracta; EROA, effective regurgitant orifice area; 3D VCA, three-dimensional vena contracta area. ^a3D VCA and quantitative Doppler EROA cut-offs may be larger than PISA EROA.

> Rebecca T. Hahn, and Jose L. Zamorano. "The Need for a New Tricuspid Regurgitation Grading Scheme." European Heart Journal - Cardiovascular Imaging, 2017

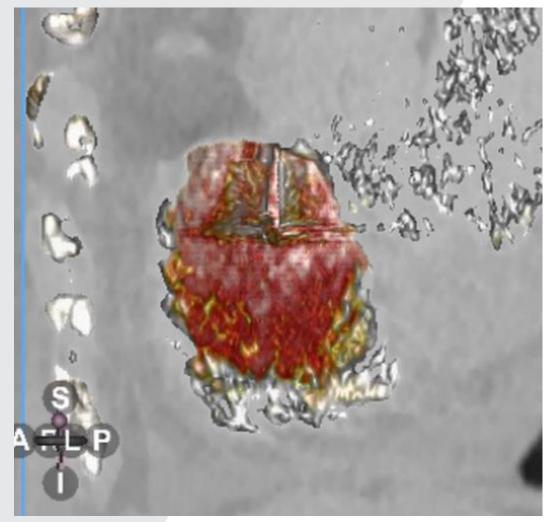


Torrential



Procedure planning: Case example









Examples







Should we intervene earlier?



Elephant in the room



Minneapolis Heart Institute Foundation

Triluminate Trial Results

6. TRILUMINATE Pivotal: TEER With the TriClip Benefits Patients With Severe TR

The trial establishes that TEER for tricuspid regurgitation is safe and effective, but to the surprise of some observers, quality-of-life improvements were what drove the primary endpoint, rather than the "harder" gains seen in the mitral space.

Only Conclusion : We need better Devices and Earlier Interventions?



"Prophylactic" Tricuspid Repair for Functional Tricuspid Regurgitation

Nicholas R. Teman, MD, Lynn C. Huffman, MD, Marguerite Krajacic, RN, Francis D. Pagani, MD, PhD, Jonathan W. Haft, MD, and Steven F. Bolling, MD Department of Cardiac Surgery, University of Michigan Health System, Ann Arbor, Michigan

- Moderate TR repaired at time of L-sided valve surgery vs. late FTR patients requiring reoperative TR
- 66 pts, propensity matched

Mortality	Prophylactic TVr (n = 42) No. (%)	Redo TVr (n = 21) No. (%)	p Value
30-day	0 (0)	3 (14.3)	0.033
1-year	3 (7.1)	4 (19.0)	
2-year	5 (11.9)	5 (23.8)	
Overall	6 (14.3)	6 (28.6)	

Table 2. Actuarial Survival After Tricuspid Valve Repair

TVr = tricuspid valve repair.

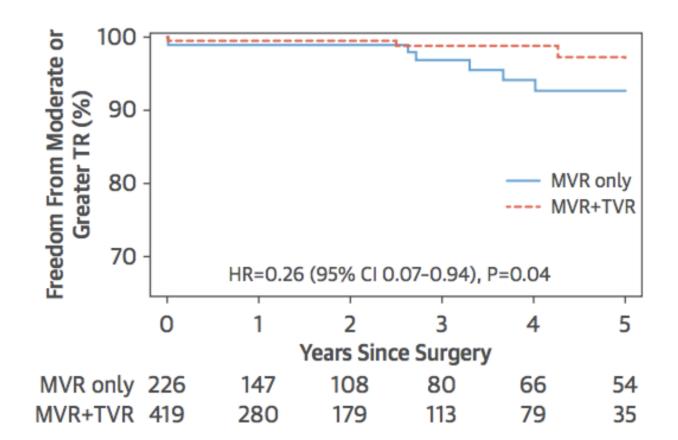
(Ann Thorac Surg 2014;97:1520-5)



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Impact of Concomitant Tricuspid Annuloplasty on Tricuspid Regurgitation, Right Ventricular Function, and Pulmonary Artery Hypertension After Repair of Mitral Valve Prolapse

Joanna Chikwe, MD, Shinobu Itagaki, MD, Anelechi Anyanwu, MD, David H. Adams, MD





Future Outlook

As therapy and understanding of the disease process and it's impact grows





