# The Role of TAVI in Current Era.

Cheol Woong Yu, MD, PhD

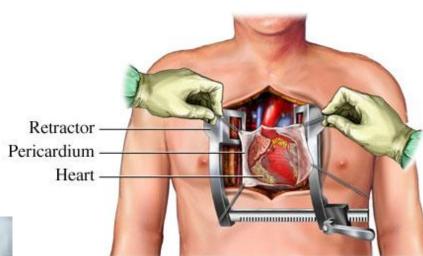
Korea University Anam Hospital Division of Cardiology





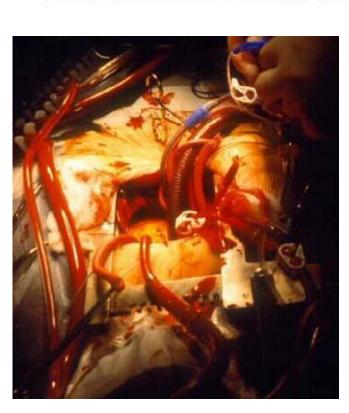


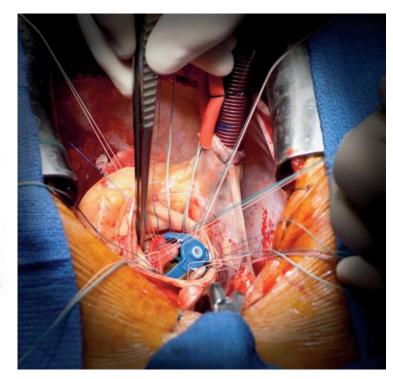


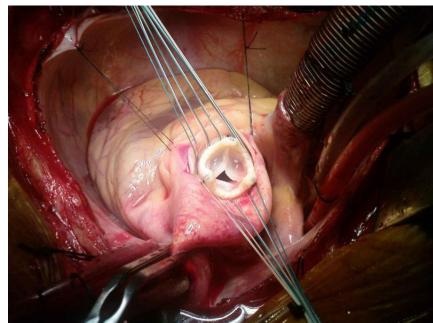










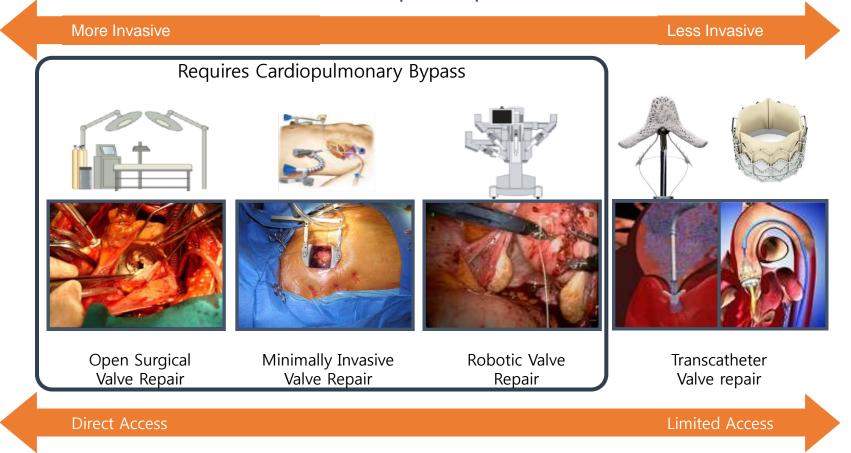






## From invasive to non invasive is a general trend

#### Valve Repair Options



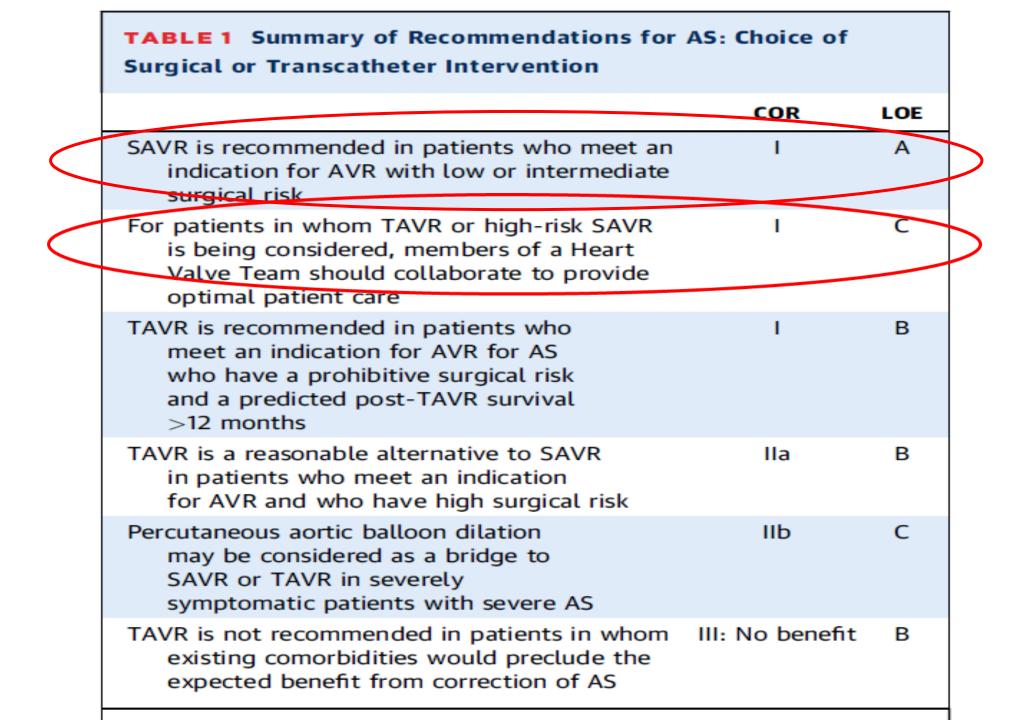
# **Current Status:** What about guidelines for TAVI?

• The Joint Task Force on the Management of Valvular Heart Disease of the European Society of Cardiology and the European Association for Cardio-Thoracic Surgery defined indications for TAVI in the 2012 Guidelines on the Management of Valvular Heart Disease.

• The corresponding 2014 U.S. Guidelines define similar indications

# Both recommend

- TAVI in patients with severe symptomatic AS who are not suitable to undergo conventional AVR as assessed by a heart team, if they are likely to gain improvement in their quality of life (QoL) and if they have a life expectancy >1 year given their comorbidities [Class of Recommendation (COR) I, Level of Evidence (LOE) B].
- TAVI should also be considered in high-risk patients with severe symptomatic AS who are suitable for surgery but in whom TAVI is favoured by a Heart Team as a COR IIa LOE B recommendation.

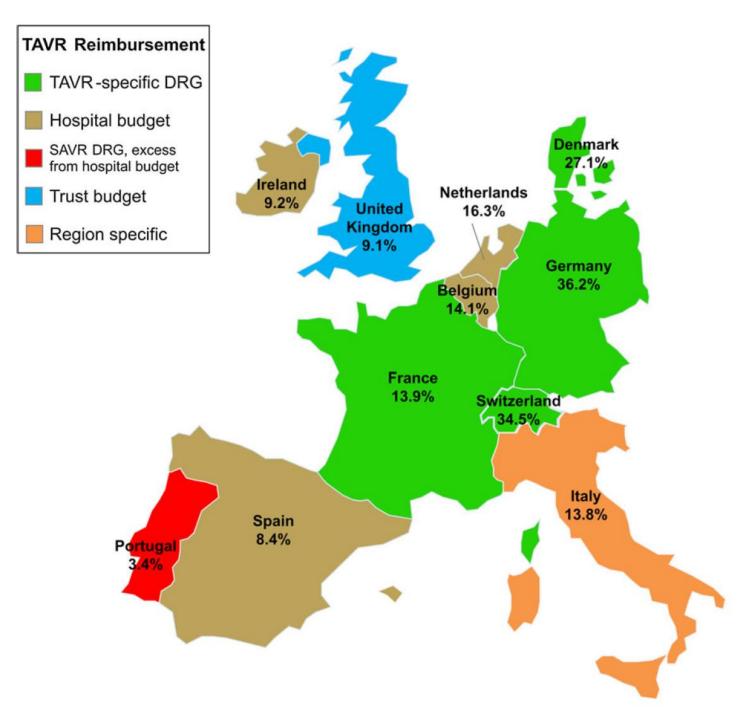


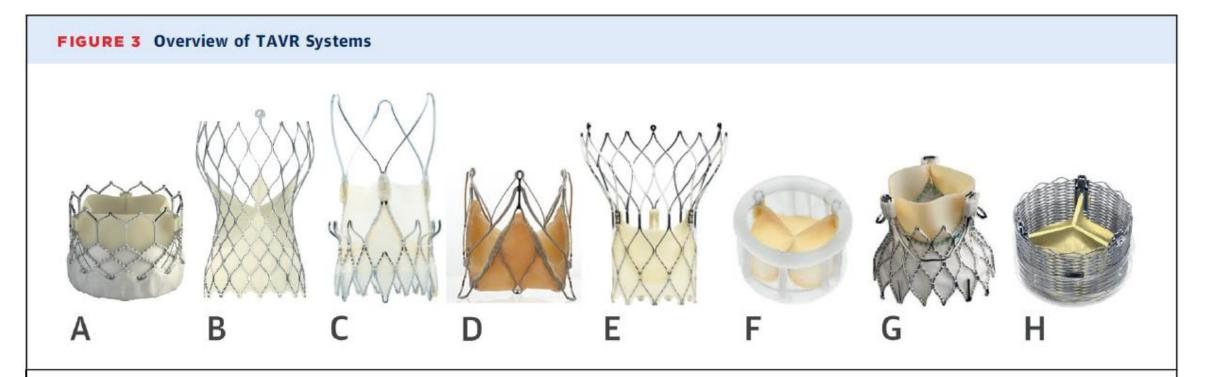
Under these current treatment indications, a significant clinical unmet need still exists worldwide



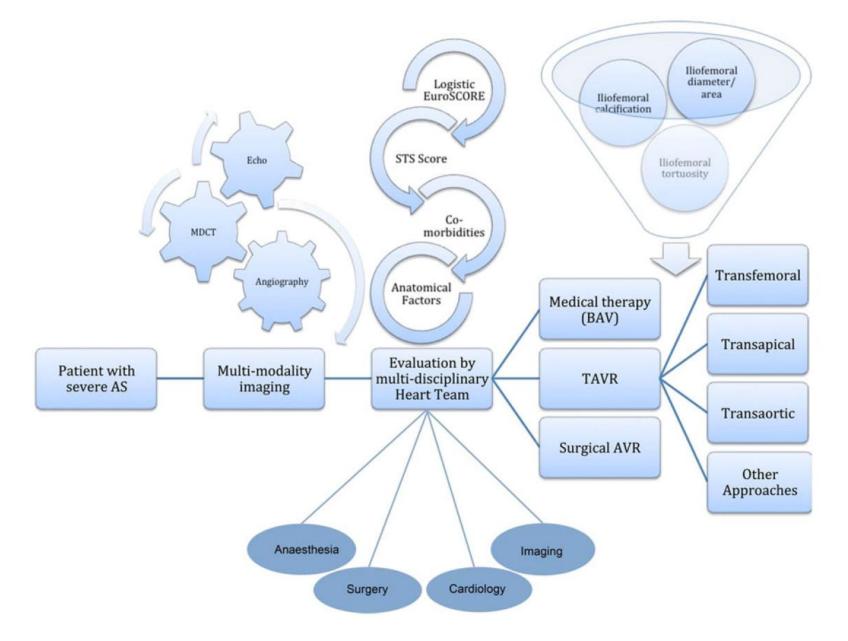
Country	Candidates (95%CI) 263 (115-152)				
Austria					
Belgium	402 (172-232)				
Czech Republic	316 (136-581)				
Denmark	179 (78-325)				
Finland	192 (82-349)				
France	2,265 (990-4,160)				
Germany	3,952 (1,684-7,227)				
Greece	529 (226-954)				
Italy	2,679 (1,145-4,958)				
Ireland	110 (46-203)				
Luxembourg	15 (6-27)				
Norway	131 (55-24)				
Poland	1,220 (512-2,226)				
Portugal	463 (197-844)				
Spain	1,737 (728-3,155)				
Sweden	318 (133-582)				
Switzerland	270 (115-495)				
The Netherlands	526 (224-965)				
The United Kingdom	2,217 (896-3,904)				
Total 19 European countries	17,712 (7,590-32,691)*				
The United States	8,205 (3,470-15,139)				
Canada	970 (408-1,777)				
otal North America	9,189 (3,898-16,682)*				

Annual number of elderly patients with severe AS who are potential TAVR candidates in different countries under current treatment indications; from Osnabrugge et al.





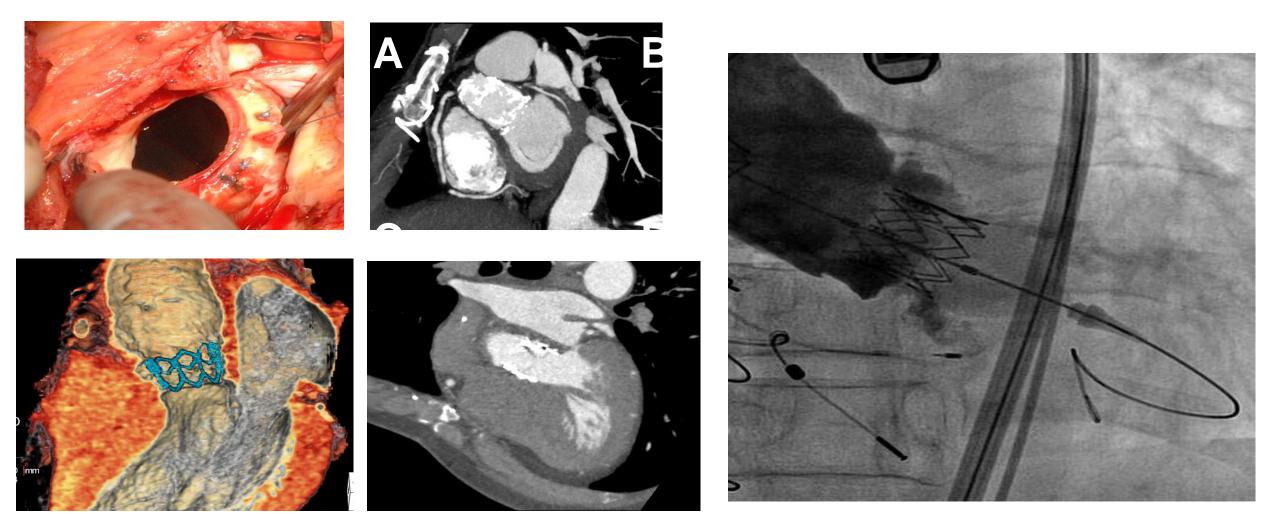
Currently, 8 transcatheter aortic valve replacement (TAVR) systems are commercially available in Europe (A-H), whereas 2 TAVR systems are approved by the U.S. Food and Drug Administration in the United States (A, B). (A) Edwards Lifesciences Sapien 3 Valve (Edwards Lifesciences, Irvine, California); (B) Medtronic CoreValve Evolut R (Medtronic, Minneapolis, Minnesota); (C) Symetis Acurate *neo* Valve (Symetis, Ecublens VD, Switzerland); (D) JenaValve (JVT Research & Development Corporation, Irvine, California); (E) St. Jude Medical Portico Valve (St. Jude Medical, St. Paul, Minnesota); (F) Direct Flow Medical Valve (Direct Flow Medical, Inc., Santa Rosa, California); (G) Medtronic Engager Valve (Medtronic, Minneapolis, Minnesota); and (H) Boston Scientific Lotus Valve (Boston Scientific, Marlborough, Massachusetts).



Optimal patient selection is best accomplished by a Heart Team, who must consider all of the patient's comorbidities (COR 1, LOE C)

Transcatheter Aortic Valve-In-Valve Implantation for Severe Bioprosthetic Stenosis after Bentall Operation Using a Homograft in a Patient with Behçet's Disease Hyung Joon Joo, Soon Jun Hong, Cheol Woong Yu

Department of Cardiology, Cardiovascular Center, Korea University Anam Hospital, Seoul, Korea



# How should we treat this patient?



# And a with a wit

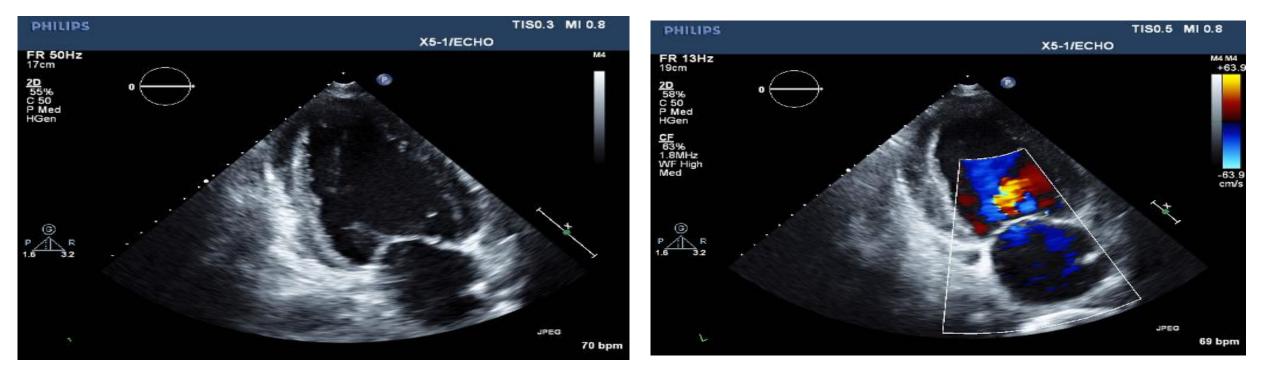
**M/80** 

CC. :NYHA Fc IV dyspnea and mental change Past Hx; HTN, HBsAg(+) Transfer to our hospital due to severe AS with pul edema. V/S : unstable..

Total bilirubin; 3.1 AST/ALT 3592/2938 BUN/Creatinine 72/2.23 Uric acid 13.0

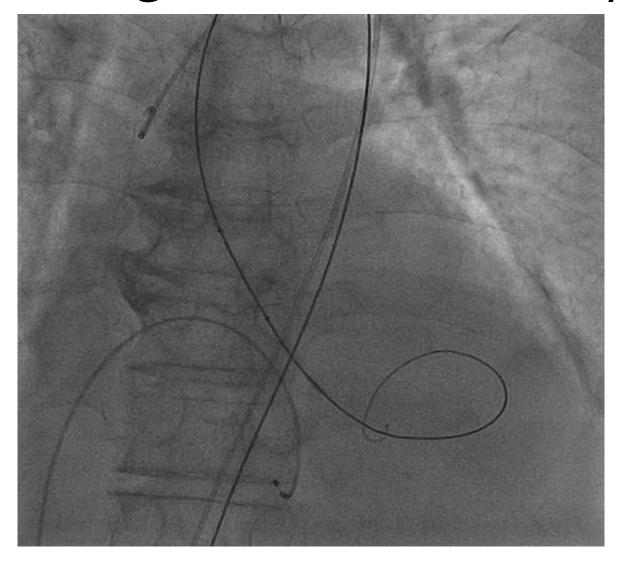
Severe AS with multiorgan failure!

# Severe LV systolic dysfunction and moderate to severe MR



# Logistic Euroscore:46.7% STS score:17.1%

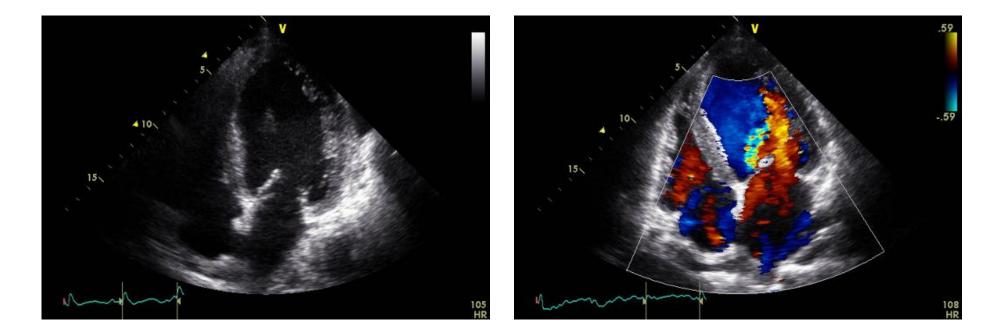
# Diagnostic and theraputic BAV!





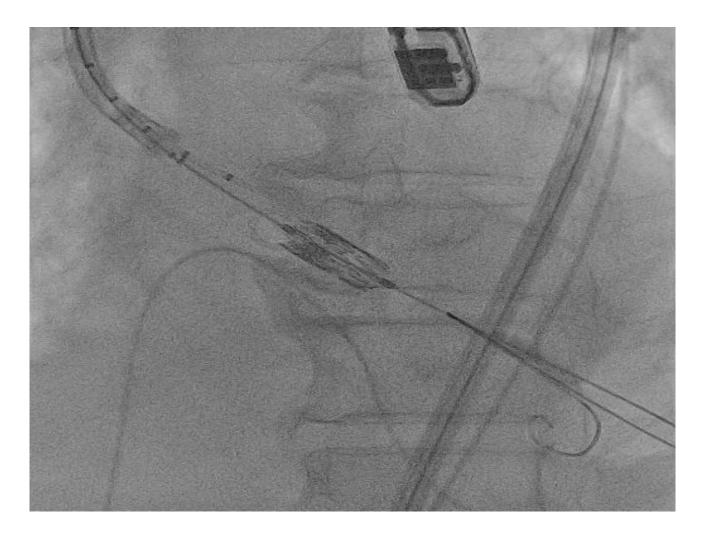
Total bilirubin; 1.2 AST/ALT 28/17 BUN/Creatinine 25/1.17 Uric acid 7.3

# Post BAV Echo

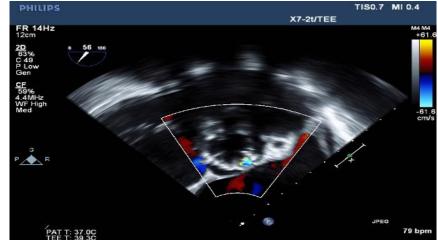


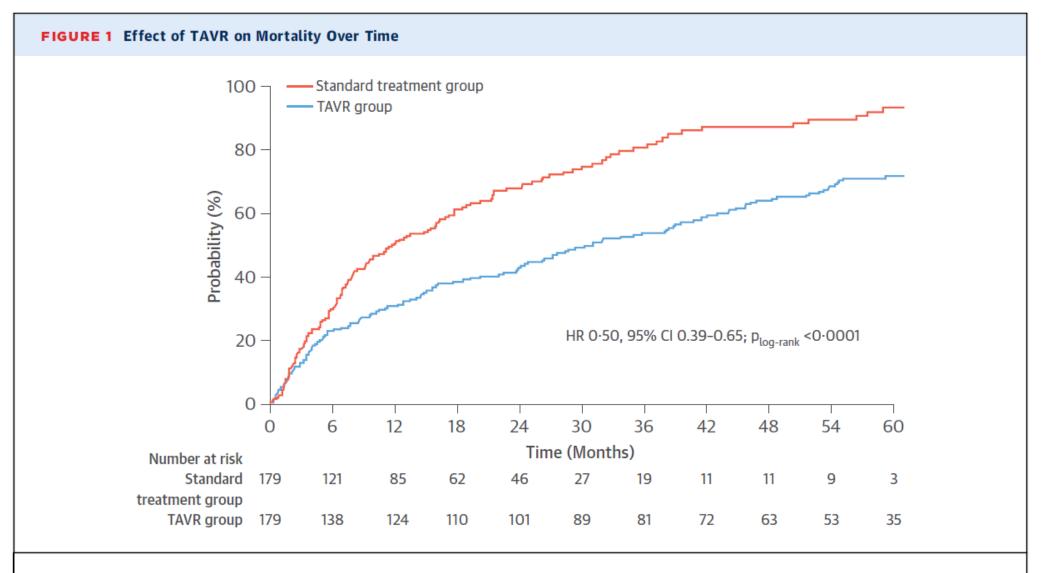
#### The problems are from AS, not comorbidities with AS

# Elective TAVR



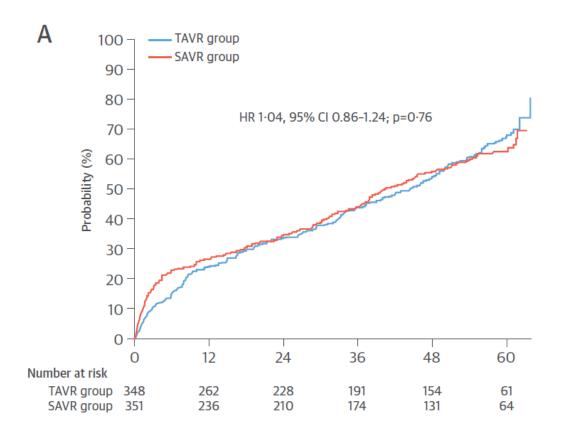




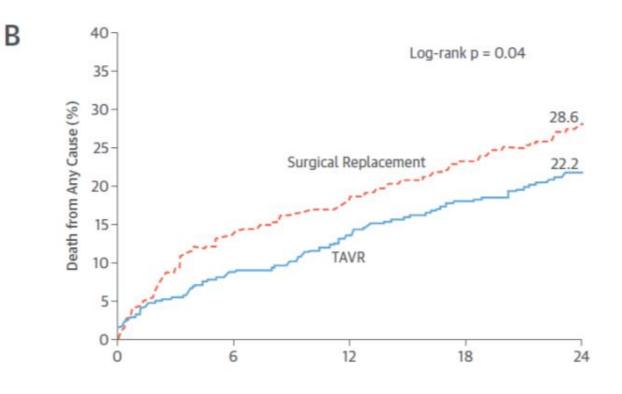


Kaplan-Meier analysis of all-cause mortality for the intention-to-treat population from the PARTNER 1B (Placement of Aortic Transcatheter Valve Trial 1B) cohort, comparing transcatheter aortic valve replacement (TAVR) versus standard therapy over 5 years. CI = confidence interval; HR = hazard ratio.

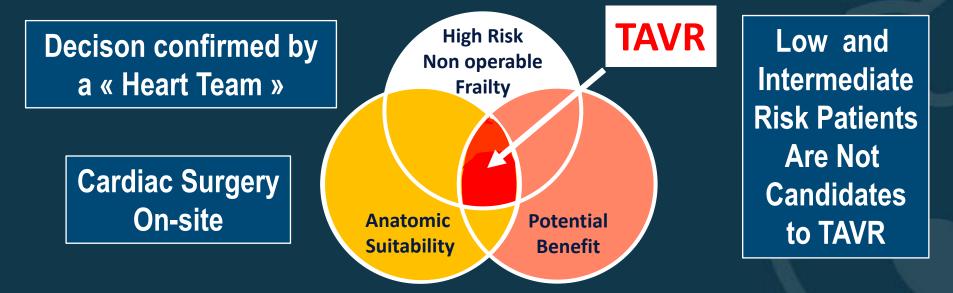
#### PARTNER 1A cohort, comparing TAVR vs SAVR over 5 years



# TAVR vs SAVR in the CoreValve U.S. Pivotal Trial



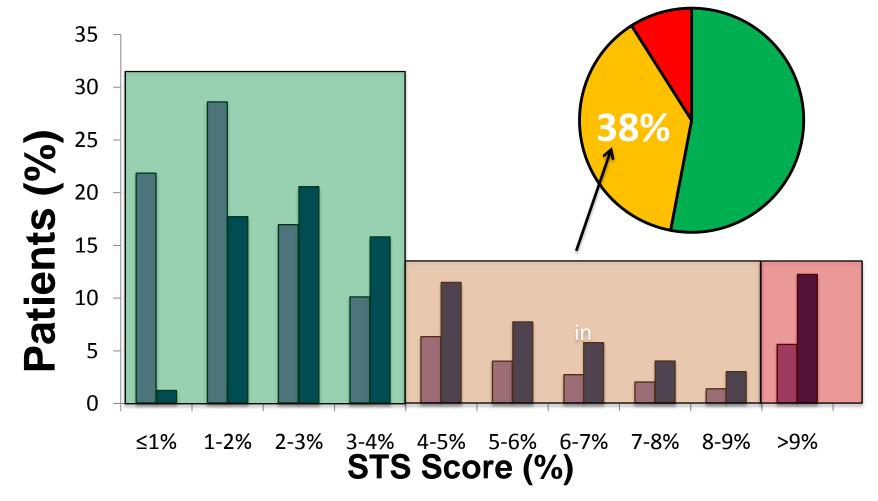
# Indications of TAVR are frozen in the past ESC Guidelines 2012 / ACC/AHA Guidline 2014



#### **PARTNER US:**

 TAVR: New technology, Ist generation devices (Edwards SAPIEN) Early experience of teams
 SAVR: Most experienced cardiac surgeons Well established treatment for 50 years

# **Estimated operative risk for AVR** 180,000 surgical patients in the US



Personal communication with M. Mack

# Elements required to expand the indications to lower risk patients

 Improved devices and strategies making TAVR
 SAFER, simpler

• Clinical evidences

ongoing

• Evidence-based trials PARTNER II: SAPIEN XT SURTAVI: CoreValve

Cost-effectiveness
 done

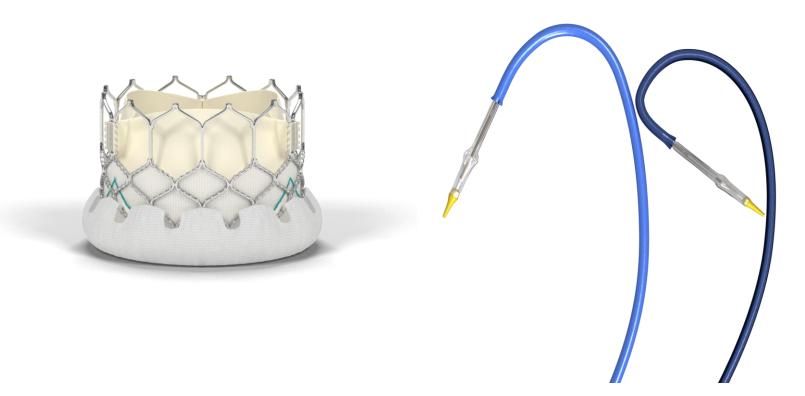
Assessment of Valve + Platform durability on longterm

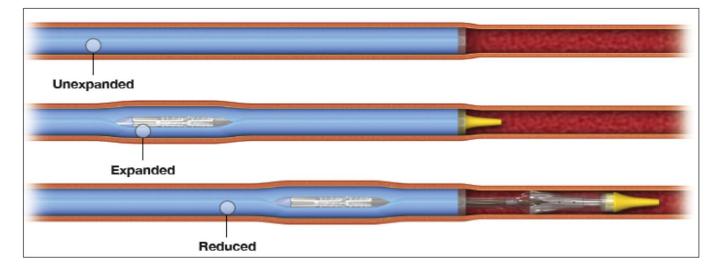
## 5 years~ upto 9 years

### PARTNER SAPIEN Platforms Device Evolution

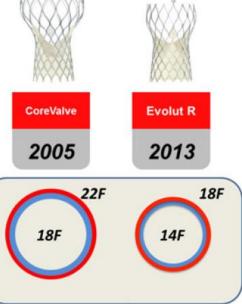




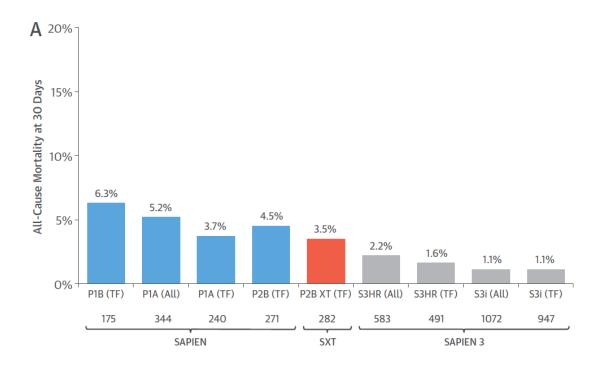


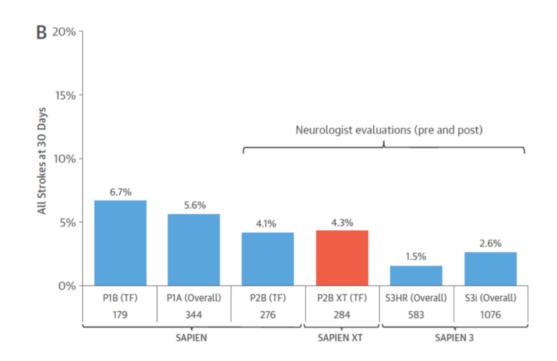




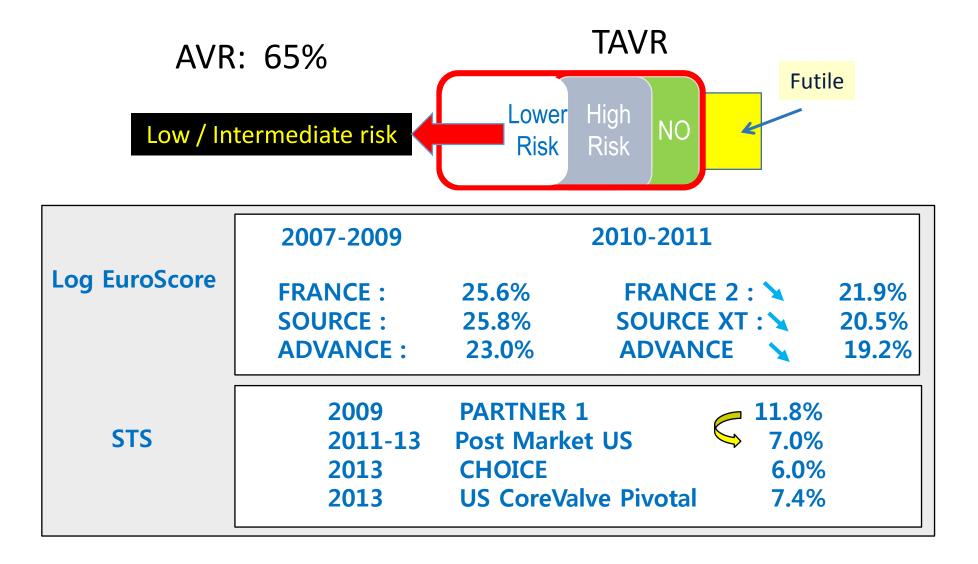


## Temporal Trends in Mortality and Stroke After TAVR





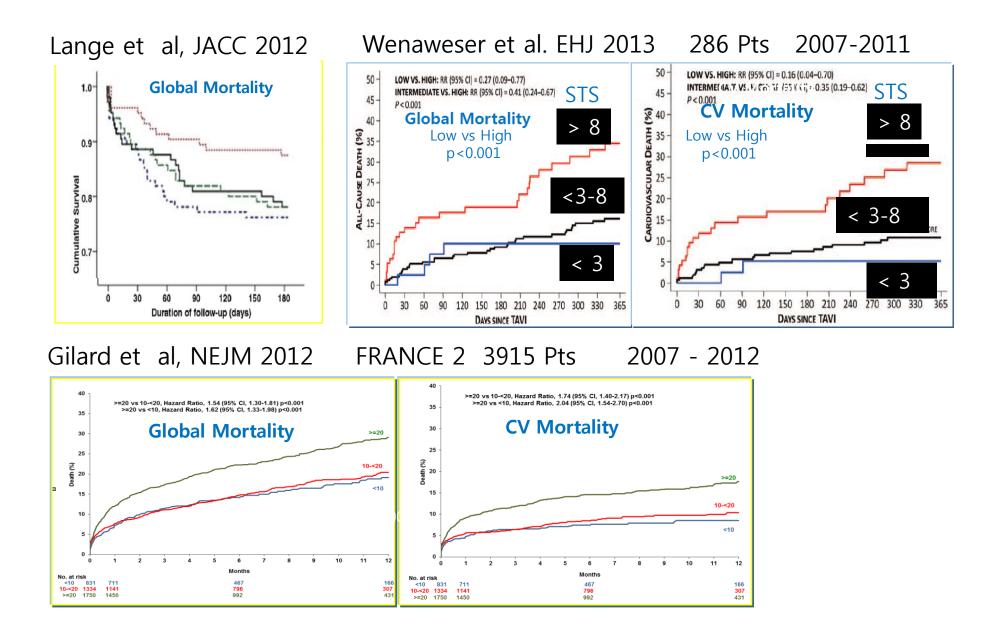
# Trend to treat lower risk patients in most recent series



# **Treating lower risk patients**

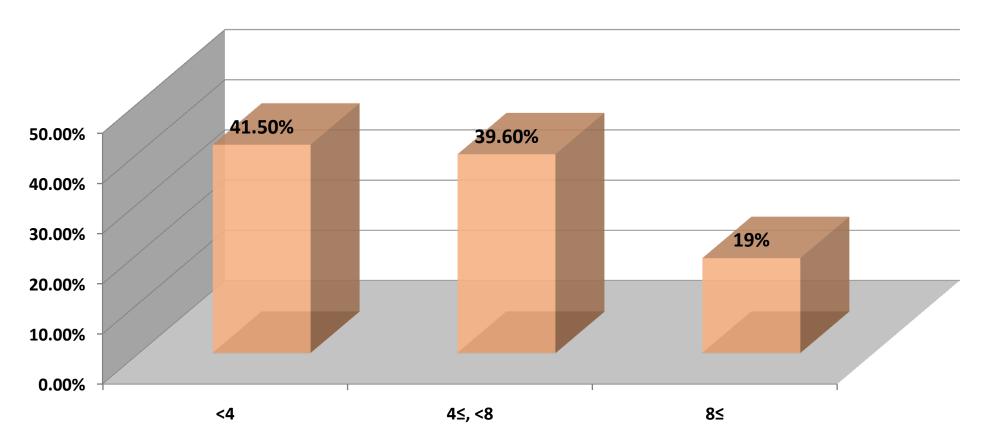
- It happens!
- What is the evidence?
- What is needed?
  - RCT
  - Better risk scores
  - Decrease complications
  - Longer follow-up
  - More data on specific subgroups
  - Better technology

# **Higher Survival in Lower Risk Patients**



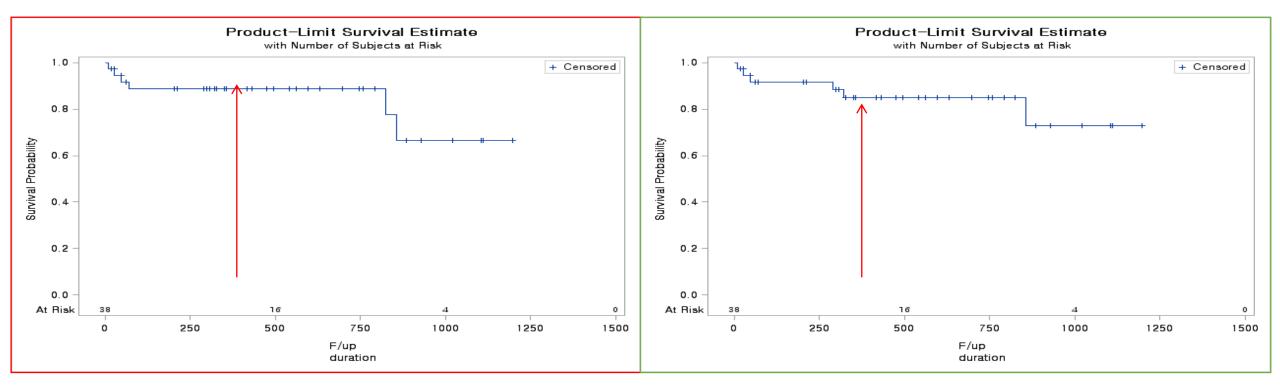
# Op risk assessment except for unoperable conditions Mean: 5.6 ± 4.1

**STS** score



Korean Sapient Valve registry data

### Clinical Outcomes at 12 months (n=50)



MACCE (n=9/50, 18%) 3 CV death 3 ischemic stroke 1 hemorrhagic stroke

> 2 infective endocarditis : 6~7 months after TAVR 1 underwent SAVR 1 underwent medical therapy

Mortality (n=7/50, 14%) 3 CV death 4 non-CV death

Korean Sapient Valve registry data

# **TAVR in Lower Surgical Risk Patients**

**MINI-FOCUS ON TAVI** 

**CLINICAL RESEARCH** 

#### A 3-Center Comparison of 1-Year Mortality Outcomes Between Transcatheter Aortic Valve Implantation and Surgical Aortic Valve Replacement on the Basis of Propensity Score Matching Among Intermediate-Risk Surgical Patients

Nicolo Piazza, MD, PHD,\*+ Bindu Kalesan, PHD,+ Nicolas van Mieghem, MD, Stuart Head, MSc, Peter Wenaweser, MD, Thierry P. Carrel, MD,# Sabine Bleiziffer, MD,\*+ Peter P. de Jaegere, MD, PHD,§ Brigitta Gahl,# Robert H. Anderson, MD, PHD,\*\* Arie-Pieter Kappetein, MD, PHD,|| Ruediger Lange, MD, PHD,\*+ Patrick W. Serruys, MD, PHD,§ Stephan Windecker, MD,¶ Peter Jüni, MD‡ Munich, Germany; Bern, Switzerland; Rotterdam, the Netherlands; Montreal, Canada; and Newcastle-Upon-Tyne, United Kingdom

#### STRUCTURAL HEART DISEASE

Acute and Late Outcomes of Transcatheter Aortic Valve Implantation (TAVI) for the Treatment of Severe Symptomatic Aortic Stenosis in Patients at High- and Low-Surgical Risk

GERHARD SCHYMIK, M.D.,<sup>1</sup> HOLGER SCHRÖFEL, M.D.,<sup>2</sup> JAN S. SCHYMIK,<sup>3</sup> RAINER WONDRASCHEK,<sup>1</sup> TIM SÜSELBECK, M.D.,<sup>4</sup> RÜDIGER KIEFER,<sup>2</sup> VERONIKA BALTHASAR, M.D.,<sup>2</sup> ARMIN LUIK, M.D.,<sup>1</sup> HERBERT POSIVAL, M.D.,<sup>2</sup> and CLAUS SCHMITT, M.D.<sup>1</sup>

From the <sup>1</sup>Medical Clinic IV, Municipal Hospital Karlsruhe, Germany; <sup>2</sup>Clinic for Cardiac Surgery Karlsruhe, Germany; <sup>3</sup>University of Munich, Germany; and <sup>4</sup>Department of Medicine, University Medical Centre Mannheim, Germany

#### Improvements in Transcatheter Aortic Valve Implantation Outcomes in Lower Surgical Risk Patients

#### A Glimpse Into the Future

Ruediger Lange, MD, PHD, Sabine Bleiziffer, MD, Domenico Mazzitelli, MD, Yacine Elhmidi, MD, Anke Opitz, MD, Marcus Krane, MD, Marcus-Andre Deutsch, MD, Hendrik Ruge, MD, Gernot Brockmann, MD, Bernhard Voss, MD, Christian Schreiber, MD, Peter Tassani, MD, PHD, Nicolo Piazza, MD, PHD

Munich, Germany

#### Clinical outcomes of patients with estimated low or intermediate surgical risk undergoing transcatheter aortic valve implantation

Peter Wenaweser<sup>1†\*</sup>, Stefan Stortecky<sup>1†</sup>, Sarah Schwander<sup>1</sup>, Dik Heg<sup>2</sup>, Christoph Huber<sup>3</sup>, Thomas Pilgrim<sup>1</sup>, Steffen Gloekler<sup>1</sup>, Crochan J. O'Sullivan<sup>1</sup>, Bernhard Meier<sup>1</sup>, Peter Jüni<sup>2</sup>, Thierry Carrel<sup>3</sup>, and Stephan Windecker<sup>1,2</sup>

#### Transcatheter vs surgical aortic valve replacement in intermediate-surgical-risk patients with aortic stenosis: A propensity score-matched case-control study

Azeem Latib, MB ChB, <sup>a,b,f</sup> Francesco Maisano, MD, <sup>c,f</sup> Letizia Bertoldi, MD, <sup>b</sup> Andrea Giacomini, MD, <sup>c</sup> Joanne Shannon, MD, <sup>a</sup> Micaela Cioni, MD, <sup>c</sup> Alfonso Ielasi, MD, <sup>b</sup> Filippo Figini, MD, <sup>a,b</sup> Kensuke Tagaki, MD, <sup>a</sup> Annalisa Franco, MD, <sup>d</sup> Remo Daniel Covello, MD, <sup>d</sup> Antonio Grimaldi, MD, <sup>d</sup> Pietro Spagnolo, MD, <sup>c</sup> Gill Louise Buchannan, MD, <sup>b</sup> Mauro Carlino, MD, <sup>b</sup> Alaide Chieffo, MD, <sup>b</sup> Matteo Montorfano, MD, <sup>b</sup> Ottavio Alfieri, MD, <sup>c</sup> and Antonio Colombo, MD <sup>a,b</sup> *Milan, Italy* 

Transcatheter aortic valve implantation versus surgical aortic valve replacement for severe aortic stenosis: Results from an intermediate risk propensity-matched population of the Italian OBSERVANT study

Paola D'Errigo <sup>a</sup>, Marco Barbanti <sup>b,c,\*</sup>, Marco Ranucci <sup>d</sup>, Francesco Onorati <sup>e</sup>, Remo Daniel Covello <sup>f</sup>, Stefano Rosato <sup>a</sup>, Corrado Tamburino <sup>b,c</sup>, Francesco Santini <sup>e</sup>, Gennaro Santoro <sup>g</sup>, Fulvia Seccareccia <sup>a</sup> and on behalf of the OBSERVANT Research Group

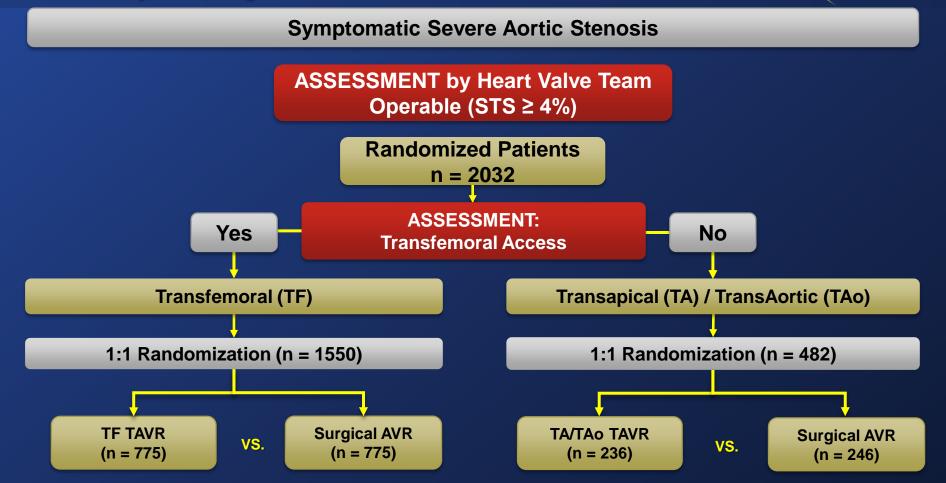
#### Intermediate or low risk patients are already being treated.....

# What is needed for treating lower risk patients?

RCT

### The PARTNER 2A Trial Study Design

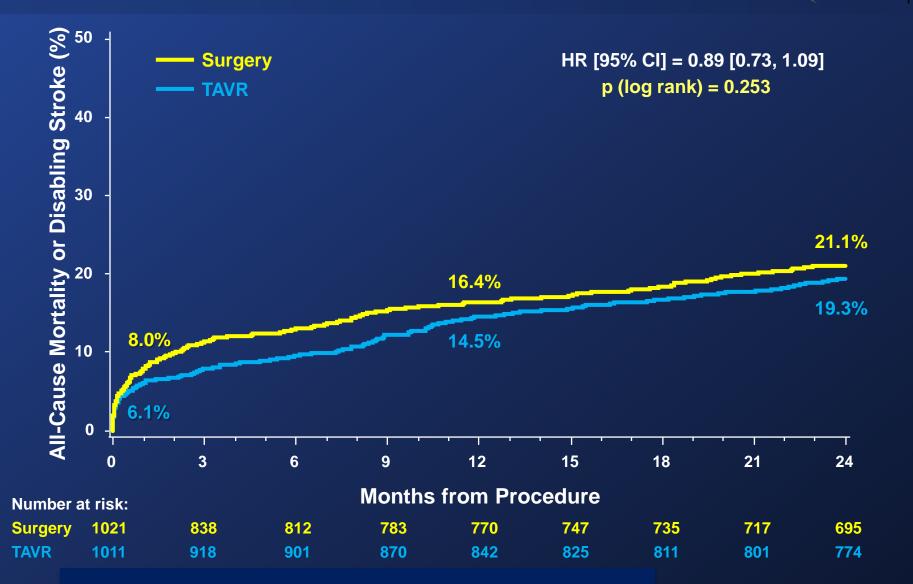




Primary Endpoint: All-Cause Mortality or Disabling Stroke at Two Years

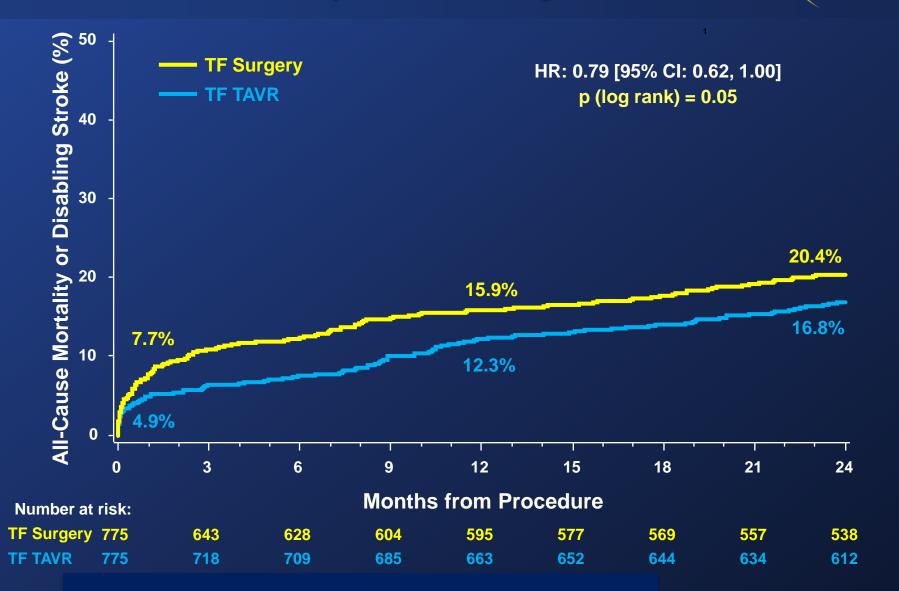
#### **Primary Endpoint (ITT)** All-Cause Mortality or Disabling Stroke





#### **TF Primary Endpoint (ITT)** All-cause Mortality or Disabling Stroke





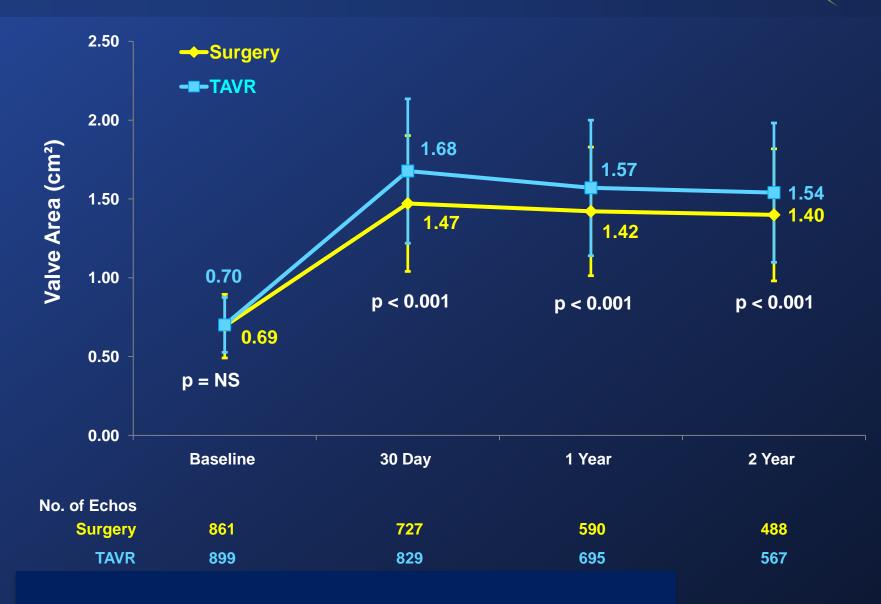
### Other Clinical Endpoints (ITT) At 30 Days and 2 Years



Events (%)	30 Days			2 Years		
	TAVR (n = 1011)	Surgery (n = 1021)	p-value*	TAVR (n = 1011)	Surgery (n = 1021)	p-value*
Rehospitalization	6.5	6.5	0.99	19.6	17.3	0.22
МІ	1.2	1.9	0.22	3.6	4.1	0.56
Major Vascular Complications	7.9	5.0	0.008	8.6	5.5	0.006
Life-Threatening / Disabling Bleeding	10.4	43.4	<0.001	17.3	47.0	<0.001
AKI (Stage III)	1.3	3.1	0.006	3.8	6.2	0.02
New Atrial Fibrillation	9.1	26.4	<0.001	11.3	27.3	<0.001
New Permanent Pacemaker	8.5	6.9	0.17	11.8	10.3	0.29
Re-intervention	0.4	0.0	0.05	1.4	0.6	0.09
Endocarditis	0.0	0.0	NA	1.2	0.7	0.22

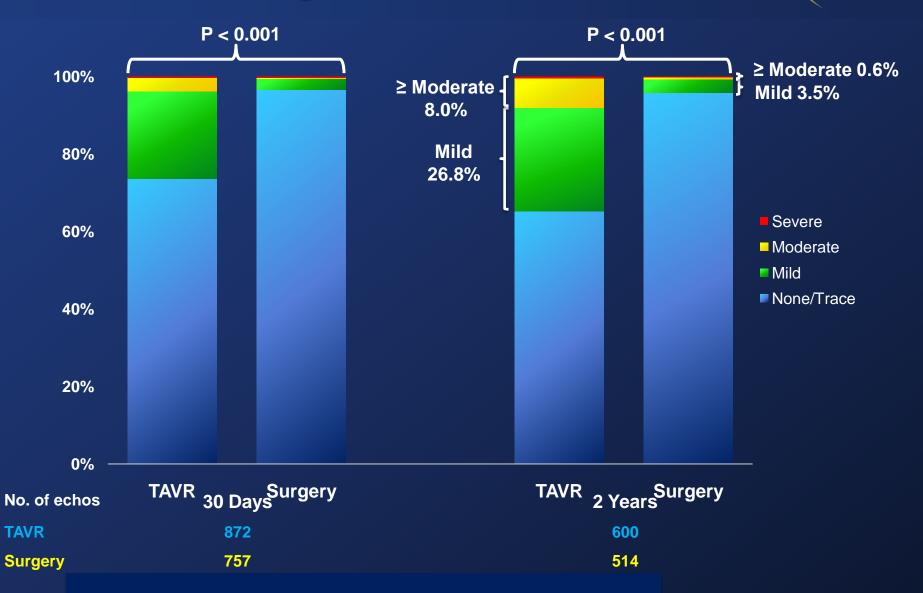
#### Echocardiography Findings (VI) Aortic Valve Area





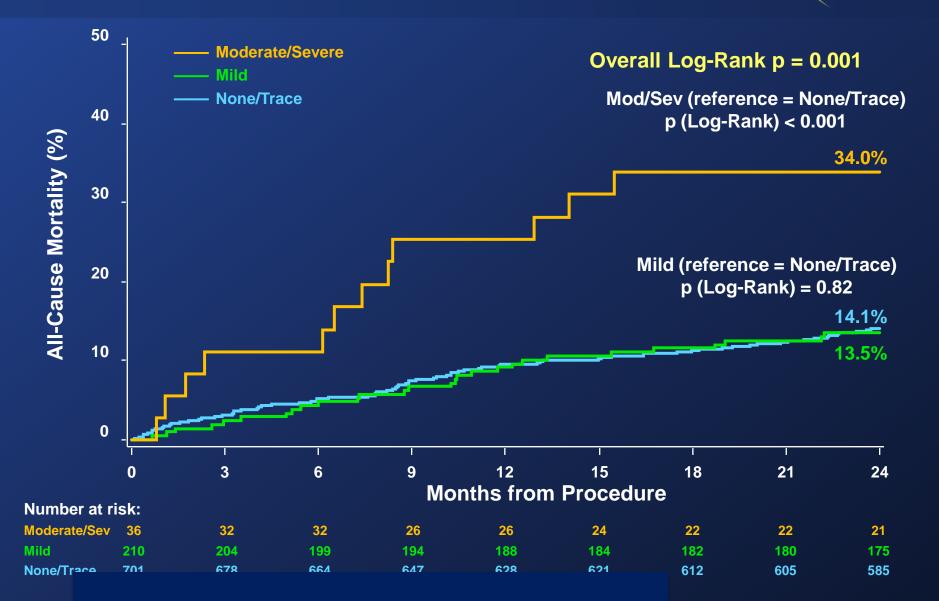
### Paravalvular Regurgitation (VI) 3-Class Grading Scheme





### Severity of PVR at 30 Days and All-cause Mortality at 2 Years (VI)



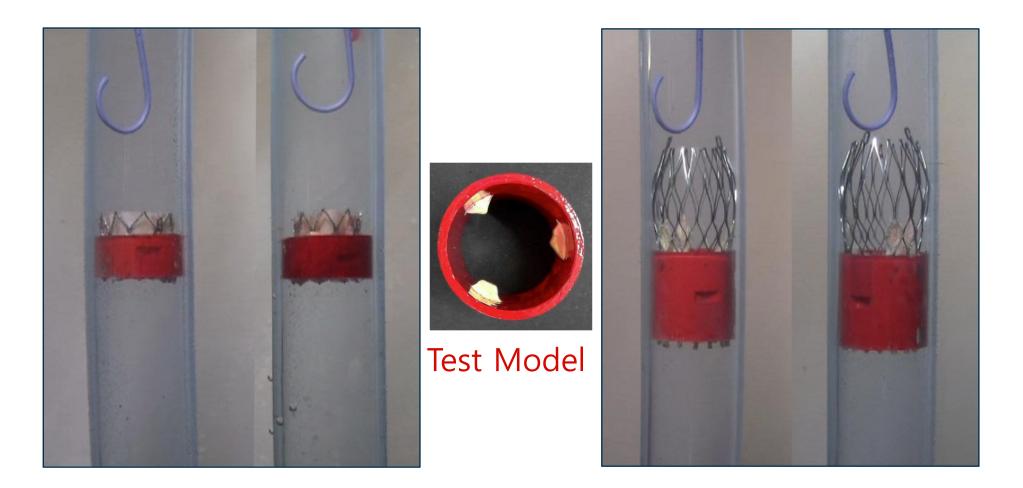


## Expandable Skirt Technology Highly-conformable, on-demand seal technology





## Elimination of paravalvular regurgitation In vitro testing



## Expandable Skirt Technology No change in device profile

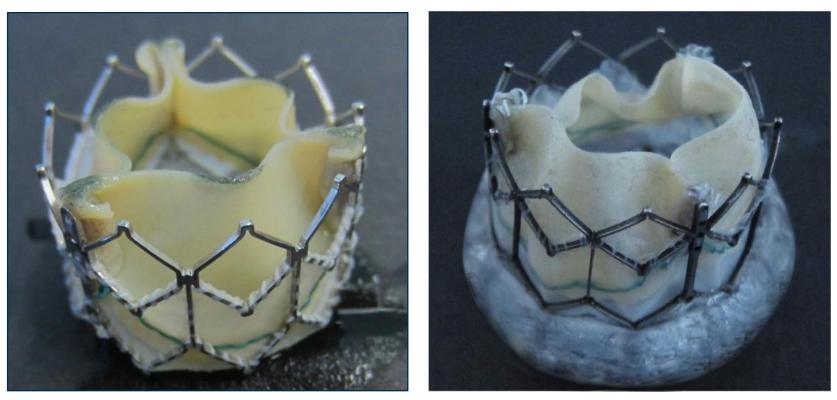




# Unexpanded state Expanded state

#### **EXPANDABLE Skirt**

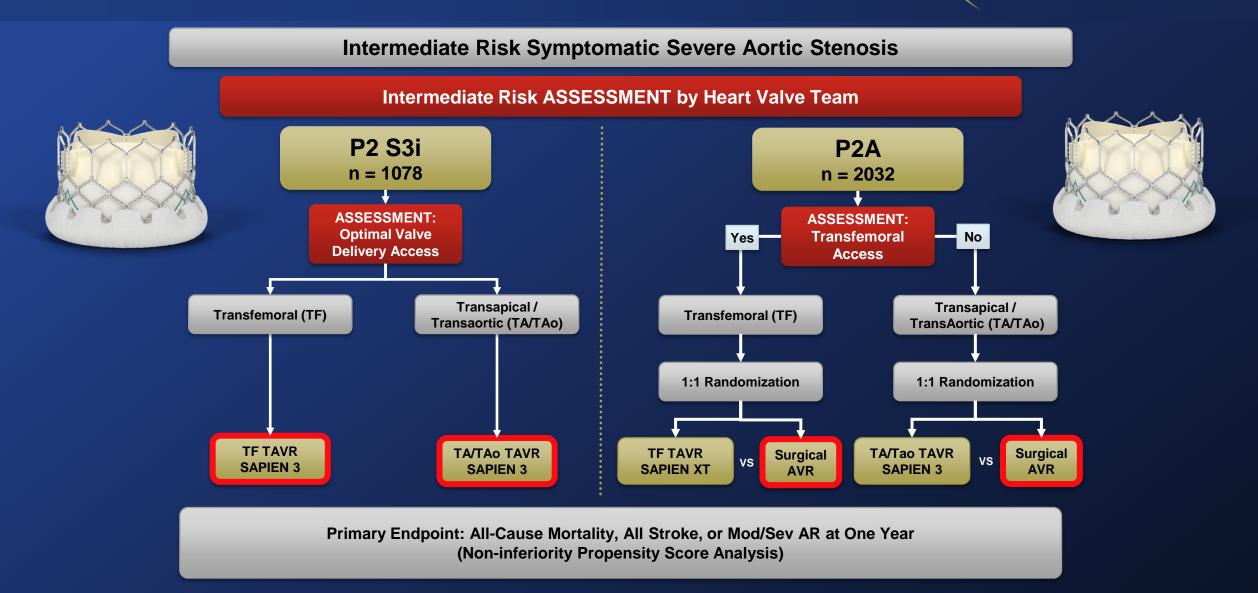
## Expandable Skirt Technology No change in device profile



**EXPANDABLE Skirt** 

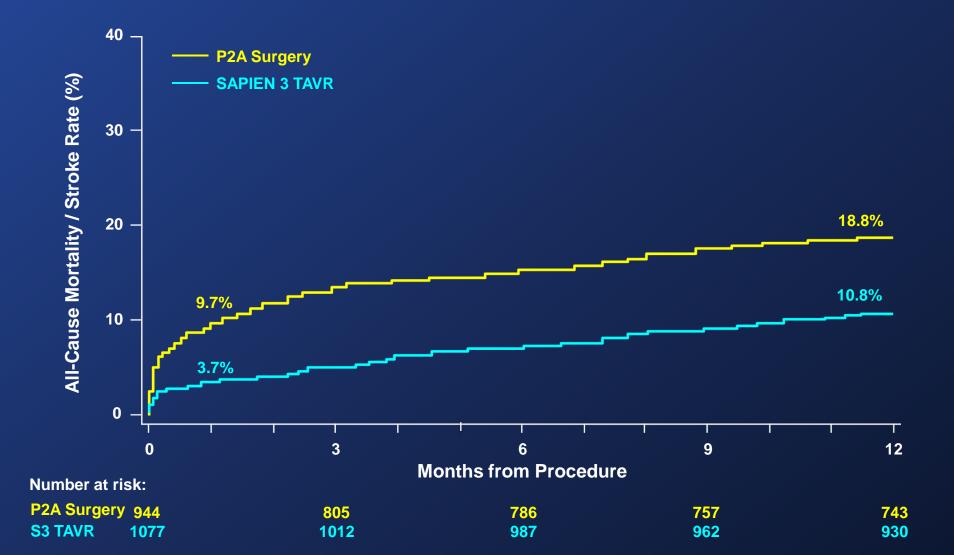
### The PARTNER 2A and S3i Trials Study Design





#### Unadjusted Time-to-Event Analysis All-Cause Mortality and All Stroke (AT)





The PARTNER 2A Trial Clinical Implications



- The results from PARTNER 2A support the use of TAVR as an alternative to surgery in intermediate risk patients, similar to those included in this trial.
- In patients who are candidates for transfemoral access, TAVR may result in additional clinical advantages.
- Long-term durability assessments of transcatheter bioprosthetic valves are still lacking and extrapolation of these findings to low-risk patients requires further clinical trial validation.

## **SAPIEN Platform Has Now Demonstrated Durability to 9 Years**

#### 1<sup>st</sup> generation



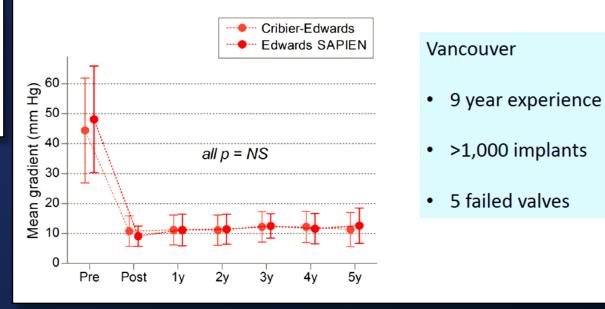
2<sup>nd</sup> generation SAPIEN

Horse pericardium Untreated

Bovine pericardium Anticalcification treatment

#### Valves explanted after 7 years

#### Vancouver 9 year durability experience



Webb at ACC 2014: New Balloon Expandable Aortic Valves

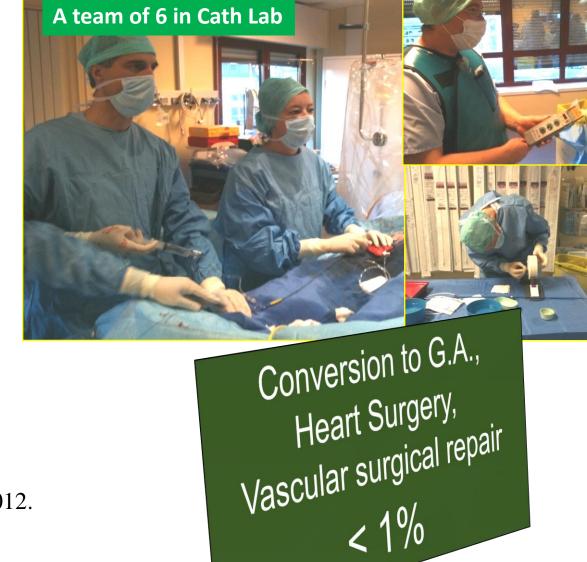
## Minimalism : Procedural Considerations for TAVR

### Strong trend to maximally simplify TAVR procedures in real world practice

- Preferential percutaneous transfemoral access,
- Reduced use of general anesthesia,
- Less intra-procedural TEE,
- Eliminating pre-dilatation,
- Decreased use of complex and costly hybrid cath lab/OR
- Early discharge programs.

## Change in Strategy: Minimalist TF-TAVR approach SAPIEN XT – Edwards SAPIEN

- Conscious sedation
- No TEE
- Percutaneous access
- Preclosing
- Procedure: 60 min
- Discharge: Day 1 to 3
- Back home



TF > 80%

Durand et al, JACC Cardiovasc Interv 2012.

# The 3MM strategy in Vancouver

- Multidisciplinary
- Multimodality
- Minimalist
  ✓ TF access
  ✓ next day discharge
- Carefully selected TF pts
- 60% (31/52) discharged in 1 d (mean LOS 1.8 d)
- Final 17 cases wo GA
- 30-day mortality 1.9%
- 2 pts (3.8%) readmitted within 30 days

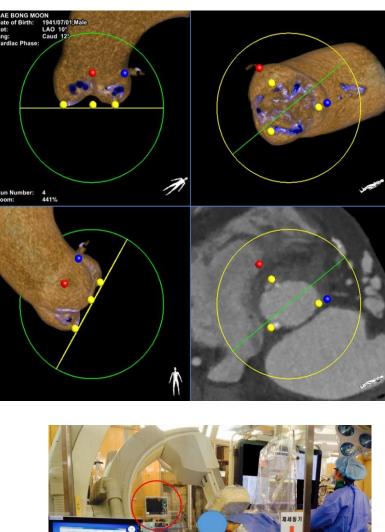
# THE VANCOUVER SUN



A 97 years old women discharged the sam e day after a TAVR procedure stretching while waiting for the bus to get home… rea d more on page 3.

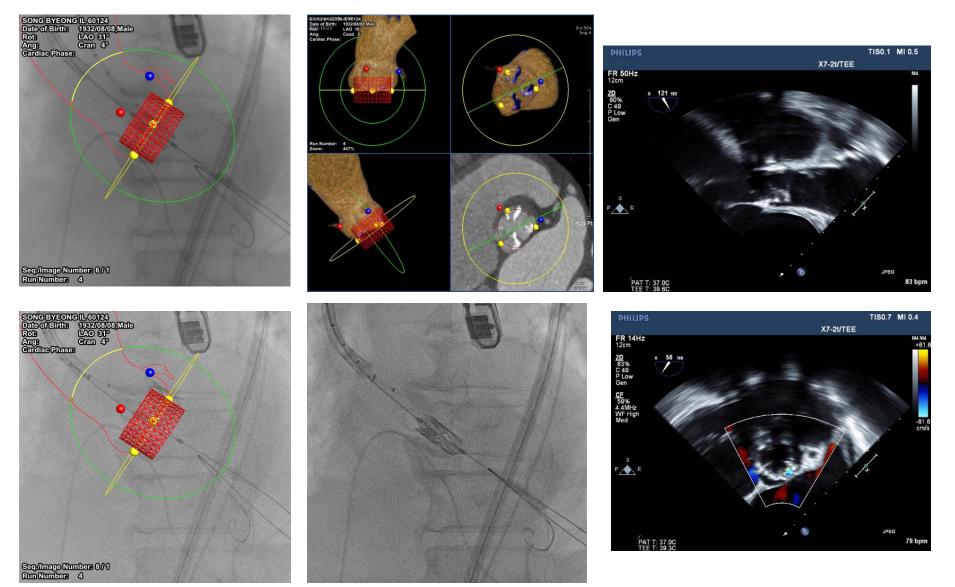
## Under the only sedation



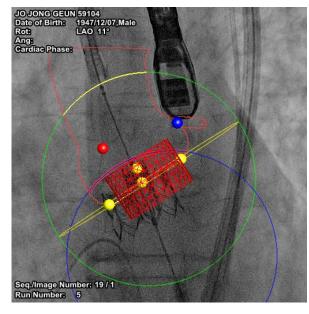




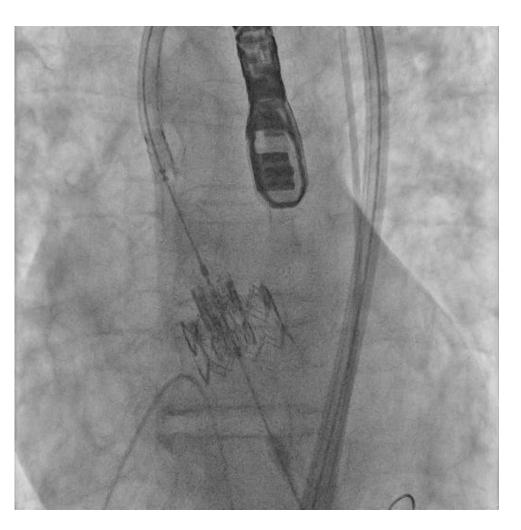
#### Valve deployment using Heart Navigating system for Quadircuspid aortic valve stenosis



## Valve in-valve guided by heart navigation system

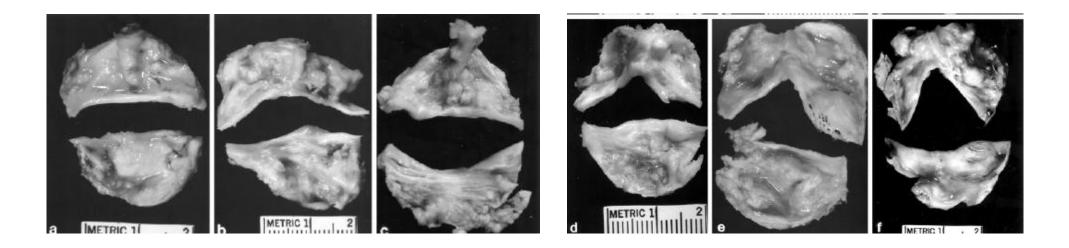






## **Bicuspid Aortic Valve disease and TAVR**

- ✓ Bicuspidy is regarded as a relative CI to TAVR due to the risk of uneven expansion of the b ioprosthesis.
- ✓ Exclusion criteria in clinical trials
- ✓ Procedural outcomes were comparable btw bicuspic & tricuspid, in terms of successful im plantation, significant AR after TAVI, 30-day combined safty endpoint



#### Transcatheter Aortic Valve Implantation for Patients With Severe Bicuspid Aortic Valve Stenosis

 Kentaro Hayashida, MD, PhD, FESC; Erik Bouvier, MD; Thierry Lefèvre, MD, FSCAI, FESC; Bernard Chevalier, MD, FSCAI, FESC; Thomas Hovasse, MD; Mauro Romano, MD;
 Philippe Garot, MD, FESC; Yusuke Watanabe, MD; Arnaud Farge, MD; Patrick Donzeau-Gouge, MD; Bertrand Cormier, MD; Marie-Claude Morice, MD, FESC

- Background—Bicuspid aortic valve (BAV) is regarded as a relative contraindication to transcatheter aortic valve implantation attributable to the risk of uneven expansion of the bioprosthesis. The purpose of this study was to evaluate the efficacy and safety of transcatheter aortic valve implantation in patients with BAV.
- *Methods and Results*—Of 470 patients included in our prospective transcatheter aortic valve implantation database (October 2006–January 2012), 229 consecutive patients undergoing both echocardiography and multidetector computed tomography were analyzed. We compared clinical outcomes in patients with vs patients without BAV. In this series of 229 patients, BAV was detected by multidetector computed tomography in 21 patients (9.2%). BAV was identified by transthoracic and transoesophagal echocardiography in only 9 of these 21 patients. Patients were 83.1±6.6 years old, and European system for cardiac operative risk evaluation score was  $20.0\% \pm 11.4\%$ . The BAV group was similar to the non-BAV group except for diabetes mellitus (4.8% vs 24.0%; P=0.05). The aortic annulus diameter in BAV patients was not significantly larger by multidetector computed tomography (24.7±3.0 vs 23.7±1.9 mm; P=0.07). The CoreValve was used more frequently in the BAV group (47.6% vs 16.3%; P=0.002). There was no significant difference in device success (100% vs 92.8%; P=0.37), risk of annulus rupture (0% vs 1.4%; P=1.00), or valve migration (0% vs 1.4%; P=1.00) in BAV patients compared with non-BAV patients. Postprocedural mean gradient (10.0±3.4 vs 9.7±4.1 mmHg; P=0.58), aortic regurgitation  $\geq 2$  of 4 (19.0% vs 14.9%; P=0.54), 30-day mortality (4.8% vs 8.2%; P=1.00), and 30-day combined safety end point (14.3% vs 13.5%; P=1.00) were also similar in both groups.
- Conclusions—In selected BAV patients, transcatheter aortic valve implantation may be associated with low complication rate, efficacy, and acceptable outcomes similar to those in non-BAV patients. (Circ Cardiovasc Interv. 2013;6:284-291.)

# Performance of transcatheter aortic valve implantation in patients with bicuspid aortic valve: Systematic review



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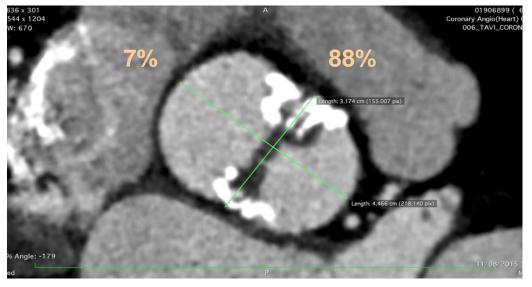
<sup>a</sup> Division of Cardiology, University of Ottawa Heart Institute, Ottawa, Ontario, Canada

<sup>b</sup> Division of Cardiology, St. Paul's Hospital, University of British Columbia, Vancouver, British Columbia, Canada

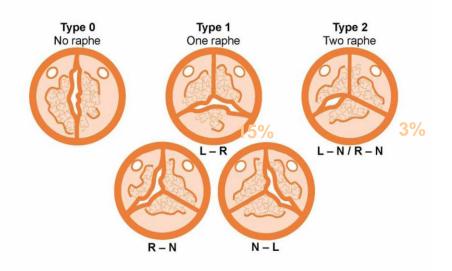
- Literature Review 92 patients
- 56% self expanding, 77% TF
- 8.6% 30 day mortality
  - 2 from aortic dissections
- PVL moderate to severe in 31%
- Long term survival good

562

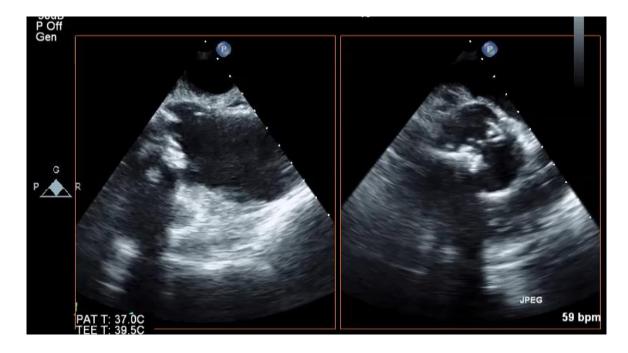
## Bicuspid Aortic Valve , Type O no Raphe

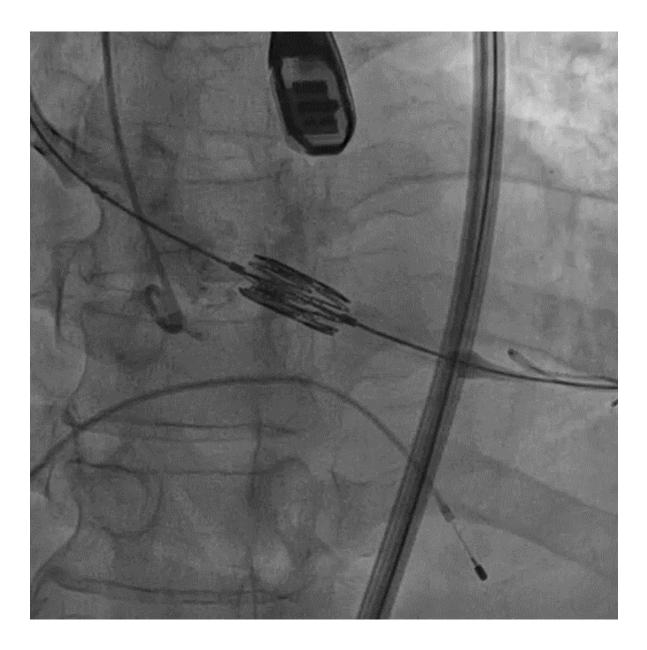


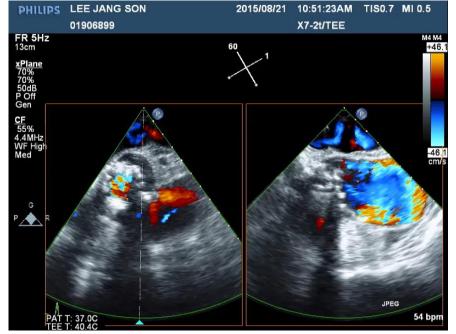
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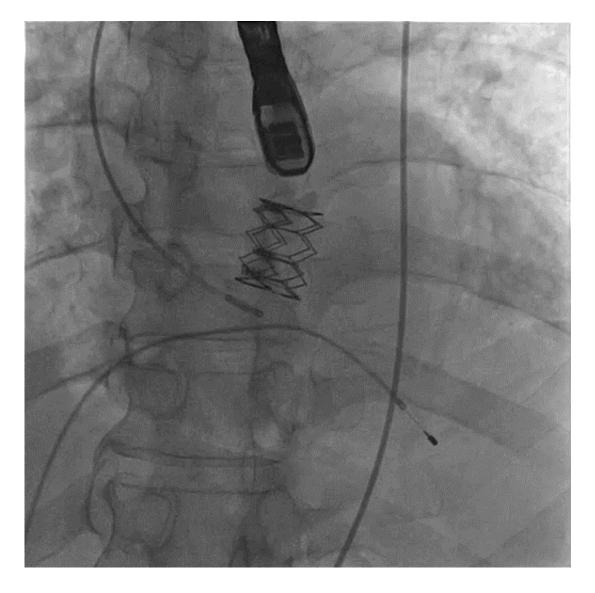
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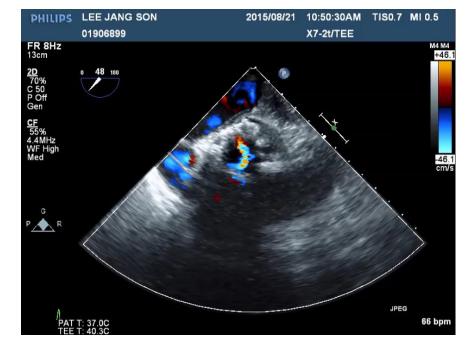


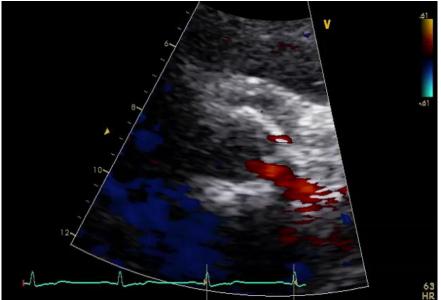




## Post TAVI







# Heart Valve Team

- The role of the Heart Team cannot be limited to preoperative assessment and choices regarding valve type and access route
- The Heart Team is essential to the management of intraoperative complications as well as postoperative care
- This includes cross-training—that is a cardiologist performing TA-TAVI (after exposure of the apex by the surgeon) or a cardiac surgeon performing TF-TAVI (assisted by an interventional cardiologist)—further promotes the ideal cooperation and collaboration of the Heart Team.

# In Summary

- Less invasive strategy is a general trend not to go against.
- TAVI is already and actively performing for the lower risk patients by virtue of increased experiences, improved device and strategy making TAVI safer and simpler.
- There are increasing data about clinical outcomes of TAVI for the lower risk patients.
- Efforts for Innovation and creation to resolve TAVI-associated problems are still ongoing
- The results from RCT support the use of TAVR as an alternative to surgery in intermediate risk patients.
- Long-term durability assessments of transcatheter bioprosthetic valves are still lacking and and more data is needed to expand TAVI indication to younger patients
- It cannot be emphasized enough that the role of heart valve team is very important for managing the AS patients.
- In the future, TAVI may be a gold standard for AS patients.

## Change the topic: How to collaborate on TAVI very well



# Thank you for your attention !!

