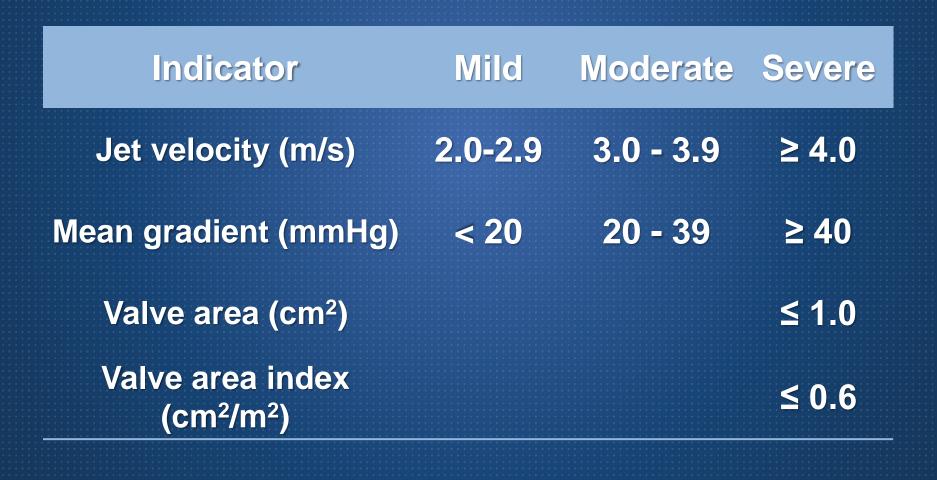
# Management of Asymptomatic Severe AS

Duk-Hyun Kang, M.D. Asan Medical Center Seoul, Korea

## 2014 AHA/ACC Guidelines Severity of AS



#### **Definition of Severe AS**

#### **Traditional Definition**

#### 2006 ACC/AHA Guidelines

Asymptomatic

Symptomatic Severe AS Asymptomatic

Symptomatic Severe AS

#### AVA < 0.75 cm<sup>2</sup>

 $AVA < 1.0 \text{ cm}^2$ 

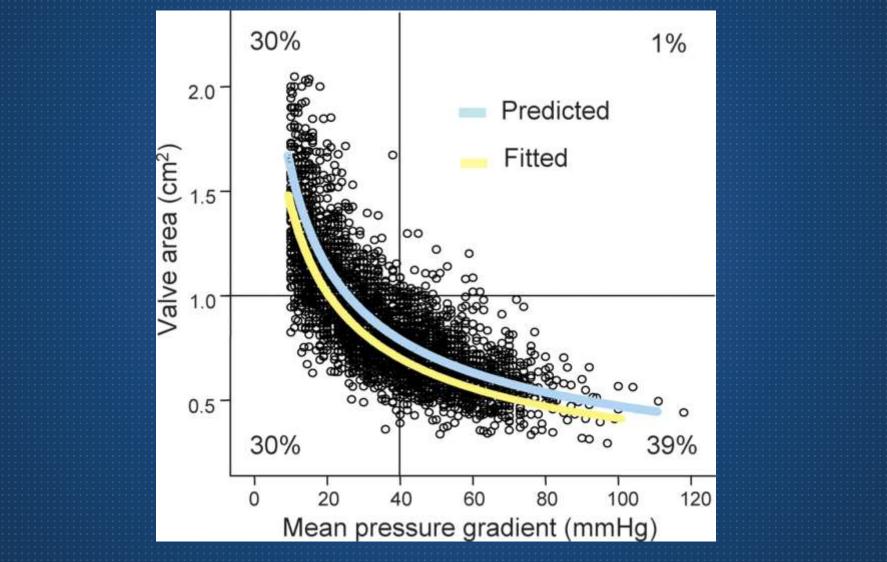
## **Relation of AV Area** to the Mean Pressure Gradient

Aortic valve area (cm<sup>2</sup>) Mean gradient (mmHg)

3.0	2.9	
2.0	6.6	
1.0	26	
0.9	32	
0.8	41	
0.7	53	
0.6	73	

Carabello BA. N Engl J Med 2002;346:677

#### **AV Area vs. Mean Pressure Gradient**



Minners J, et al. Eur Heart J 2008

# Aortic Stenosis Low Gradient

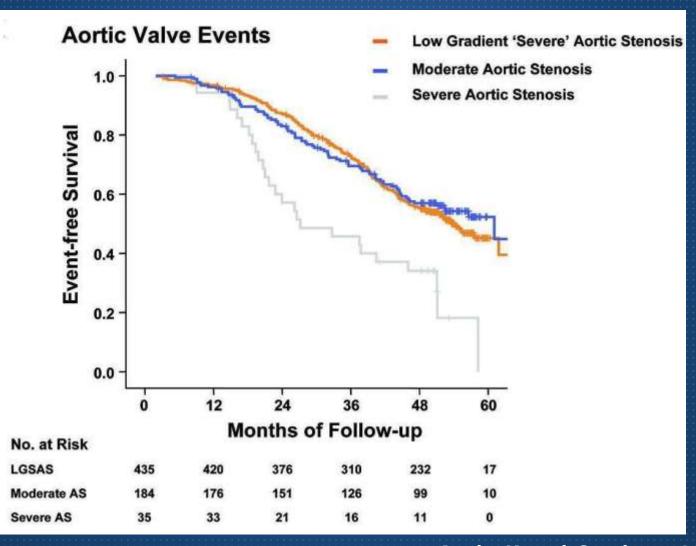
Outcome of Asymptomatic Patients with Low-Gradient "Severe" Aortic Stenosis

In 619 asymptomatic patients (SEAS study), AV events\* occurred in 48.5% pts with low-gradient "severe" AS (AVA < 1.0 cm<sup>2</sup> and mean gradient ≤ 40 mmHg) versus 44.6% with moderate AS (AVA: 1.0-1.5 cm<sup>2</sup>) during 46 months of follow-up (P= 0.37)
Outcome of low-gradient "severe" AS and normal ejection fraction similar to that of moderate AS

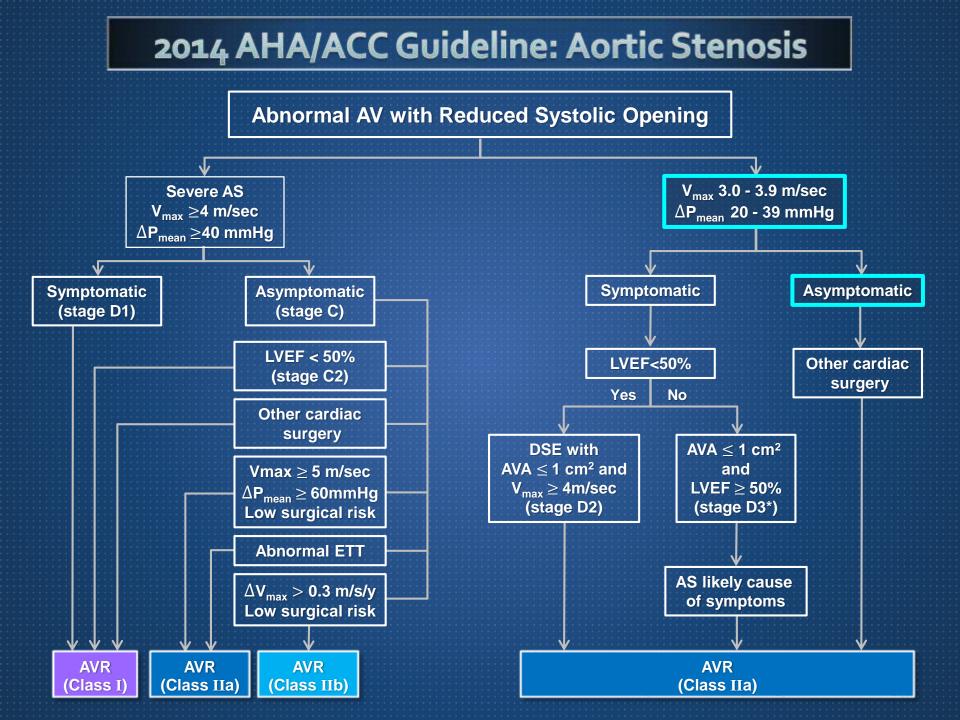
AV events\*: CV death, AVR and CHF

Jander N, et al. Circulation 2011;123:887

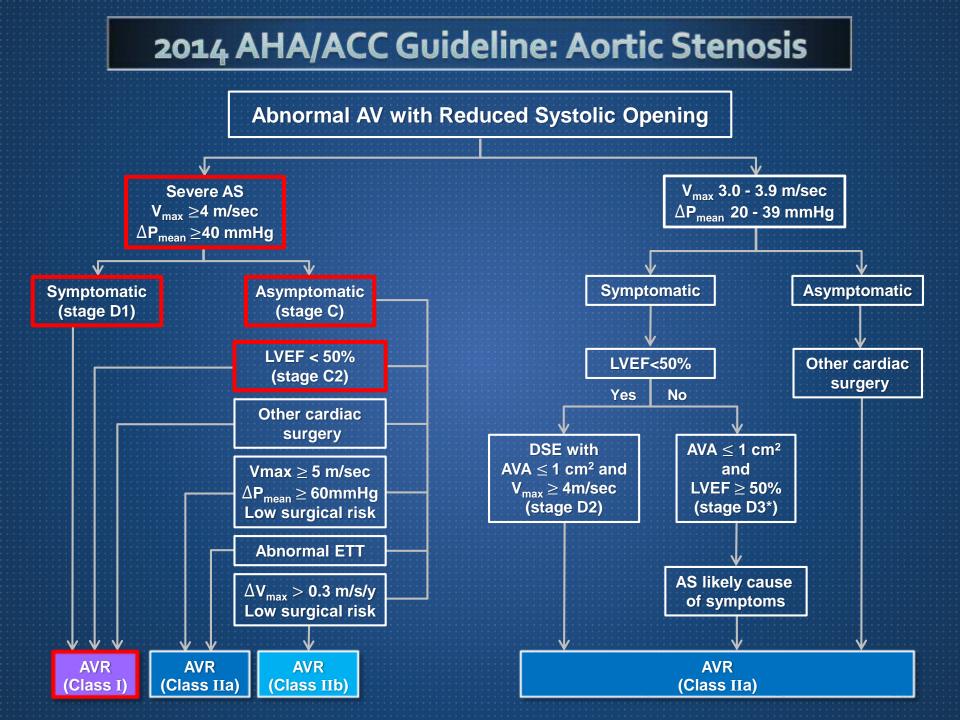
## **Outcome in Low-Gradient "Severe" AS**



Jander N, et al. Circulation 2011;123:887



# Aortic Stenosis High Gradient



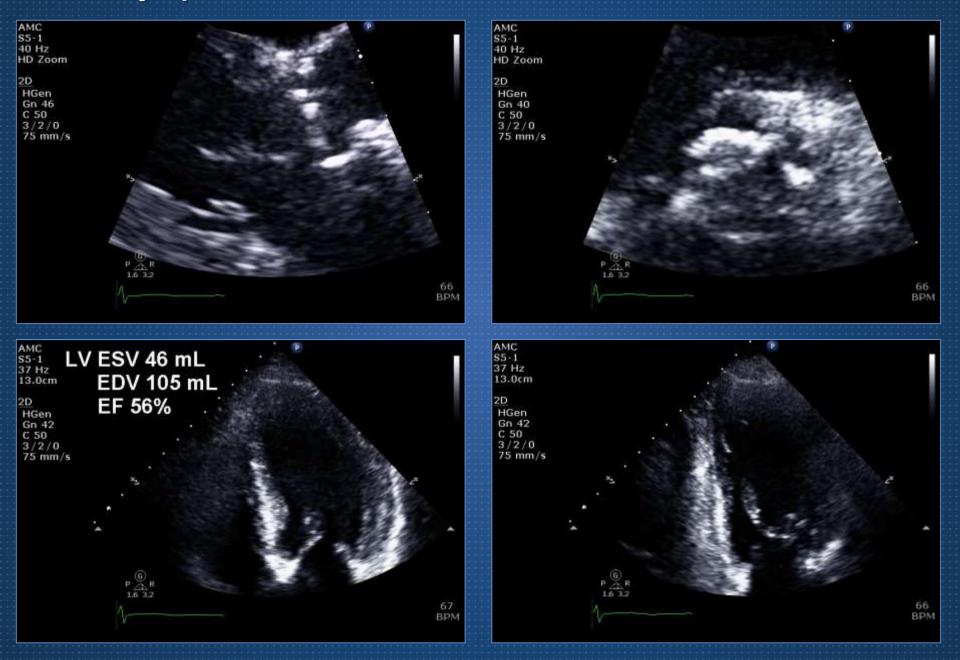
•

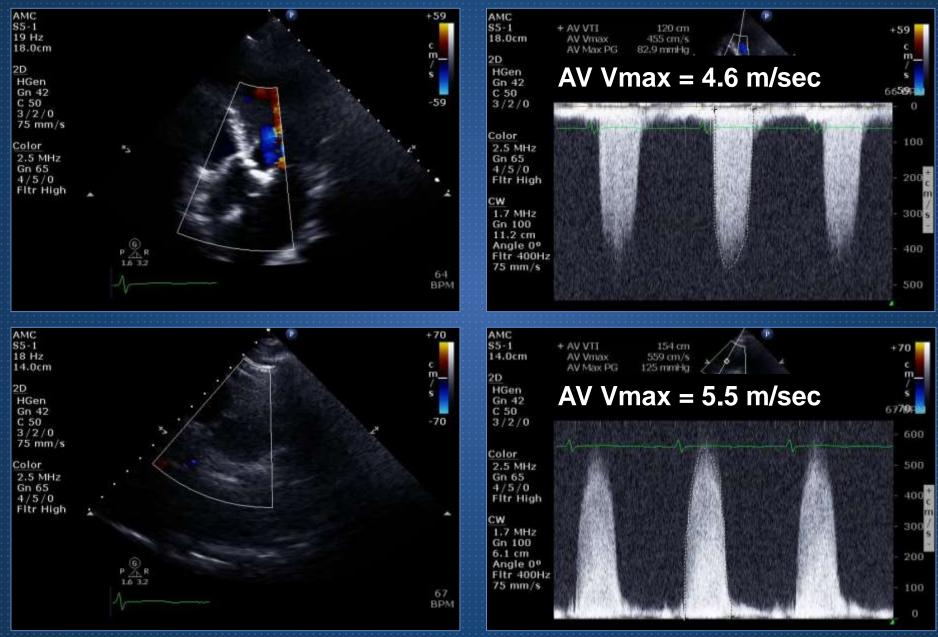
•

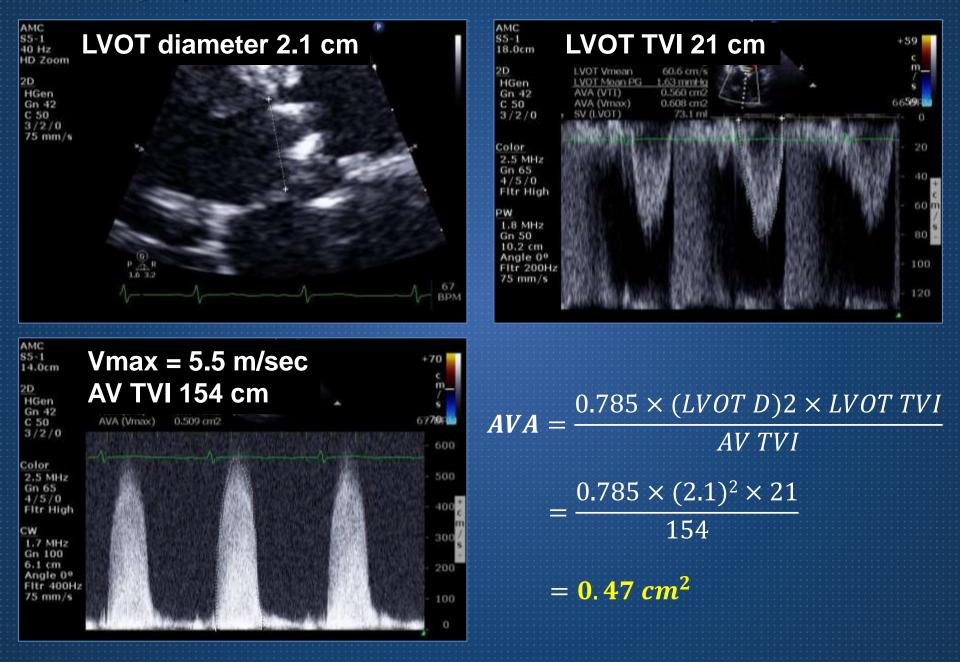
•

•

60 year-old female Referred from local clinic due to cardiac murmur No chest pain or dyspnea She regularly went to dance class without symptoms







# What will you recommend to this patient?

## 1) Watchful Waiting

#### 2) Surgical Aortic Valve Replacement

## 3) Transcatheter AV Replacement (TAVR)

## **Dilemma of Early AVR**

Balancing the real risks versus potential benefits of AV replacement in asymptomatic severe AS

Risks	Benefits		
Operative mortality	Preventing sudden death		
Prosthesis-related	Lowering cardiac mortality		
mortality and morbidity	related with refusal and		
	delay of surgery		

Comparison of Early Surgery versus Conventional Treatment in Asymptomatic Very Severe Aortic Stenosis

> Duk-Hyun Kang, Sung-Ji Park\*, Ji Hye Rim, Dae-Hee Kim, Jong-Min Song, Kee-Joon Choi, Seung Woo Park\*, Jae-Kwan Song, Jae-Won Lee, Pyo-Won Park\*

Division of Cardiology, Cardiac Surgery Asan Medical Center, Samsung Medical Center\* Seoul, South Korea

Kang DH, et al. Circulation 2010;121:1502

## **Echocardiographic Evaluation**

Etiology of AS and grading of AV calcification on 2D echo

 Maximal aortic jet velocity, mean pressure gradients and aortic valve area on Doppler

Very severe aortic stenosis
 AV area ≤ 0.75 cm<sup>2</sup> fulfilling one of criteria; peak
 aortic velocity ≥ 4.5 m/sec or mean pressure
 gradient ≥ 50 mmHg

#### **Definition of Severe AS**

#### **Traditional Definition**

#### 2006 ACC/AHA Guidelines

Asymptomatic

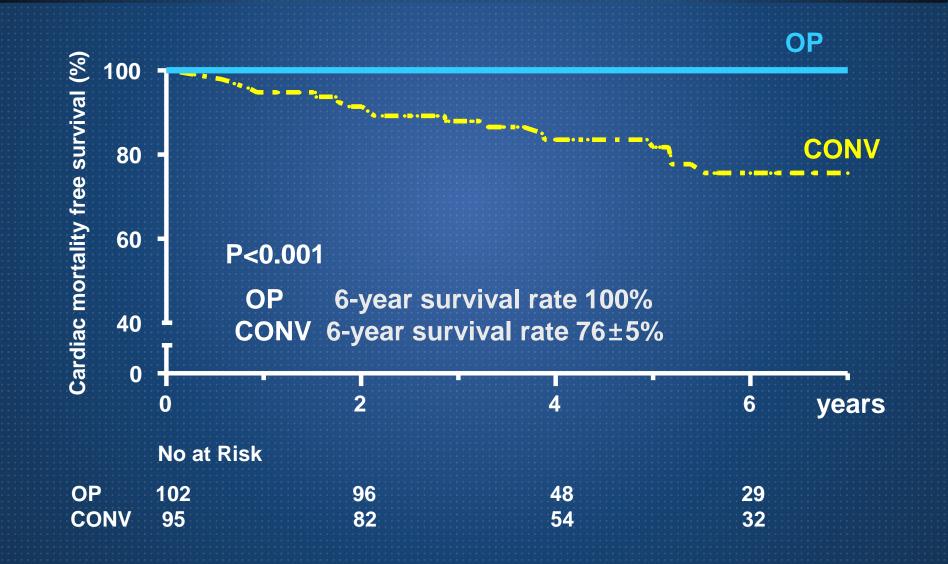
Symptomatic Severe AS Asymptomatic

Symptomatic Severe AS

#### AVA < 0.75 cm<sup>2</sup>

 $AVA < 1.0 \text{ cm}^2$ 

## Survival Free of Cardiac Death OP versus CONV group



## Conclusions

Early surgery is associated with improved longterm survival by decreasing cardiac mortality and sudden cardiac death in very severe AS A prospective randomized trial is required to confirm the efficacy of early surgery

Kang DH, et al. Circulation 2010;121:1502

# Summary of Debate in 2012

 Watchful waiting for the majority of asymptomatic patients with severe AS

 High risk patients with AVA < 0.8 cm<sup>2</sup> may benefit from preemptive AVR in selected centers with excellent operative outcomes

Circulation 2012;126:112-125

Initial Surgical versus Conservative Strategies in Patients with Asymptomatic Severe Aortic Stenosis

## **CURRENT AS Registry**

Retrospective, multicenter registry enrolled 1808 asymptomatic severe AS patients in 27 Japanese centers between 2003 and 2011

Taniguchi T, et al. JACC 2015;66:2827-38

## Incidence of Events at 5 Years Propensity-matched Cohort

	Initial AVR Group	Conservative Group	P value	
Number	291	291		
Age, years	71.6±8.7	73.1±9.3	0.047	
High gradient AS	243 (84%)	179 (62%)	< 0.001	
All-cause Death	<mark>40 (15%)</mark>	<mark>69 (26%)</mark>	0.009	
Cardiovascular Death	25 (10%)	<b>46 (19%)</b>	0.01	
AV related Death	<mark>13 (5%)</mark>	33 (14%)	0.003	
HF Hospitalization	<b>10 (4%)</b>	<b>50 (20%)</b>	< 0.001	

## Unadjusted All-cause Mortality Early Surgical AVR versus Observation

	Early	AVR	Observation			Risk Ratio	Risk Ratio	
Study	Events	Total	Events	Total	Weight	M-H Random, 95% Cl	M-H Rando	m, 95% Cl
Pellikka et al.	2	30	14	113	11.5%	0.54[0.13, 2.24]		
Pai et al.	10	99	147	239	31.4%	0.16[0.09, 0.30]		
Kang et al.	3	102	10	95	13.9%	0.28[0.08, 0.98]		
Taniguchi et al.	40	291	542	1517	43.2%	0.38[0.29, 0.52]	-	
Total (95% CI)		522		1964	100%	0.29[0.17, 0.51]	•	
Total events	55		713					
							10 100 Observation	

Généreux P, et al. JACC 2016;67:2263-88

## Adjusted All-cause Mortality Early Surgical AVR versus Observation



Généreux P, et al. JACC 2016;67:2263-88

# **Limitations of meta-analysis**

Residual confounding and selection bias

 Patients in conservative group were older and sicker; refusal of AVR in 31% during follow-up
 Prospective randomized clinical trials comparing AVR to conservative treatment is warranted

Généreux P, et al. JACC 2016;67:2263-88

Aortic Valve replAcement versus conservative Treatment in Asymptomatic seveRe Aortic Stenosis (AVATAR)

#### **12 Clinical Sites**

Belgium, Croatia, Czech Republic, France, Ireland, Lithuania, Poland, Serbia

Banovic M, Iung B, Bartunek J, et al. Am Heart J 2016;174:147-53

## **Methods**

• Primary End Point: Composite of all-cause death, AMI, stroke and **CHF** hospitalization Secondary End Point: Operative mortality, repeat AV surgery, major bleeding and thromboembolic complications

## **Methods**

#### From 2015 to 2020

312 asymptomatic patients with severe AS

#### **Conservative Tx**

(N=156)

Early surgery (N=156)

#### Follow-up of 3 Years After Enrollment

Evaluation of Transcatheter Aortic Valve Replacement Compared to SurveiLance for Patients with AsYmptomatic Severe Aortic Stenosis (EARLY TAVR)

From 2017 to 2021

1109 Asymptomatic Patients with Severe AS

#### Surveilance

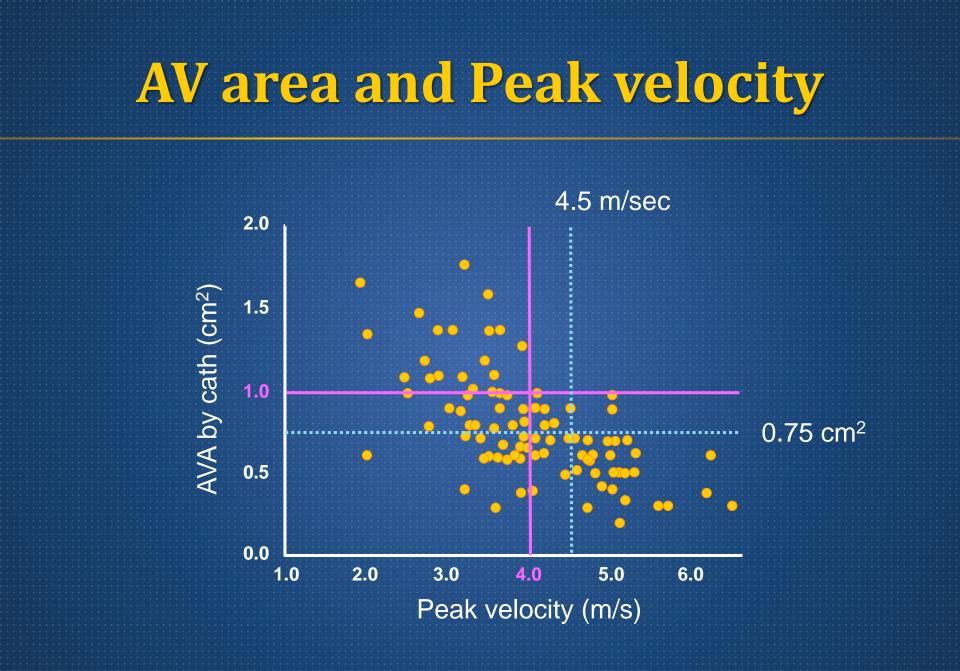
Early TAVR

Follow-up of 2 Years After Enrollment

Randomized Comparison of Early Surgery versus COnventional Treatment in Asymptomatic VERY Severe Aortic Stenosis (RECOVERY)

Duk-Hyun Kang, Sung-Ji Park\*, Geu-Ru Hong+, Sahmin Lee, Dae-Hee Kim, Hyung-Kwan Kim, Jong-Min Song, Seung Woo Park\*, Jae-Kwan Song, Jae-Won Lee, Pyo-Won Park\*

> Division of Cardiology, Cardiac Surgery AMC, SMC\*, SNUH,YUH+ Seoul, South Korea



#### Oh JK, et al. J Am Coll Cardiol 1988

## **Methods**

Primary End Point:
Composite of operative mortality, cardiac death and prosthesis related mortality during follow-up for 4 years
Secondary End Point:

All-cause mortality during follow-up

## **Methods**

#### From 2010 to April, 2019

145 Asymptomatic Patients with Very Severe AS

#### Conventional Tx CONV (N=72)

Early surgery OP (N=73)

Follow-up of 4 Years After Enrollment

## Randomized Trial of Early Surgery Versus Conventional Treatment for Asymptomatic VERY Severe Aortic Stenosis (RECOVERY)

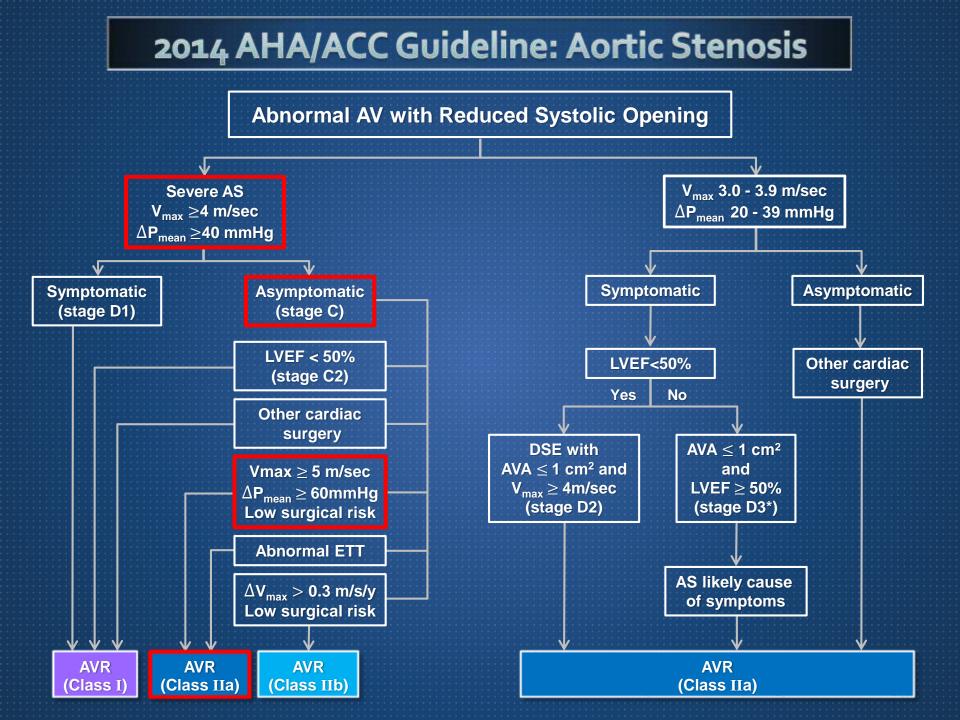


#### Duk-Hyun Kang, MD, PhD on behalf of The RECOVERY Trial Investigators

Asan Medical Center, Seoul, Korea







# **Take Home Messages**

 Watchful waiting for the majority of asymptomatic patients with severe AS Patients with very severe AS may benefit from preemptive AVR in selected centers with excellent operative outcomes Ultimately, a randomized trial will solve the debate between early AVR and watchful observation