



HARVARD MEDICAL SCHOOL  
TEACHING HOSPITAL

# Trends in Atrial Fibrillation Management

Leon Ptaszek, MD, PhD, FACC, FHRS

Cardiac Arrhythmia Service, Massachusetts General Hospital  
Assistant Professor of Medicine, Harvard Medical School

*19 August 2024*

*Disclosures: Research grant from Anumana; Consultant for Abbott, Bristol-Myers Squibb, NeuTrace, Voiant, Medtronic, Moderna, Pfizer*



MASSACHUSETTS  
GENERAL HOSPITAL

**CORRIGAN MINEHAN  
HEART CENTER**

# Objectives

- 1) **Describe advances in medical and interventional treatments for prevention of AF- related stroke.**
- 2) **Describe advances in medical and interventional treatments for minimizing the burden of AF.**
- 3) **Summarize how these advances informed the current guidelines for AF management.**

# Outline

- 1) **Clinical Case**
- 2) **Epidemiology of AF**
- 3) **Physiology of AF**
- 4) **Advances in medical and interventional treatment:**
  - *Reduce the risk of AF-related stroke*
  - *Minimize consequences of AF on heart function (symptoms, CHF)*
- 5) **Current recommendations for management of AF**

# Outline

- 1) **Clinical Case**
- 2) **Epidemiology of AF**
- 3) **Physiology of AF**
- 4) **Advances in medical and interventional treatment:**
  - *Reduce the risk of AF-related stroke*
  - *Minimize consequences of AF on heart function (symptoms, CHF)*
- 5) **Current recommendations for management of AF**

# Clinical Case

## History of Present Illness

67-year-old woman reports for a routine care visit. Exertional capacity stable, no new exertional dyspnea or chest discomfort. No palpitations, presyncopal symptoms, or syncopal events.

## Past Medical History

Hypertension

Hyperlipidemia

Diabetes Mellitus

## Physical Exam

Vital signs: BP 136/80 HR 96 RR 12 O2 sat 99% on RA

General: Well-appearing

Pulmonary: clear lung sounds throughout both lung fields

Cardiac: irregularly irregular S1S2, no S3, no rubs/murmurs

Abdomen: benign

Lower extremities: well-perfused, no edema

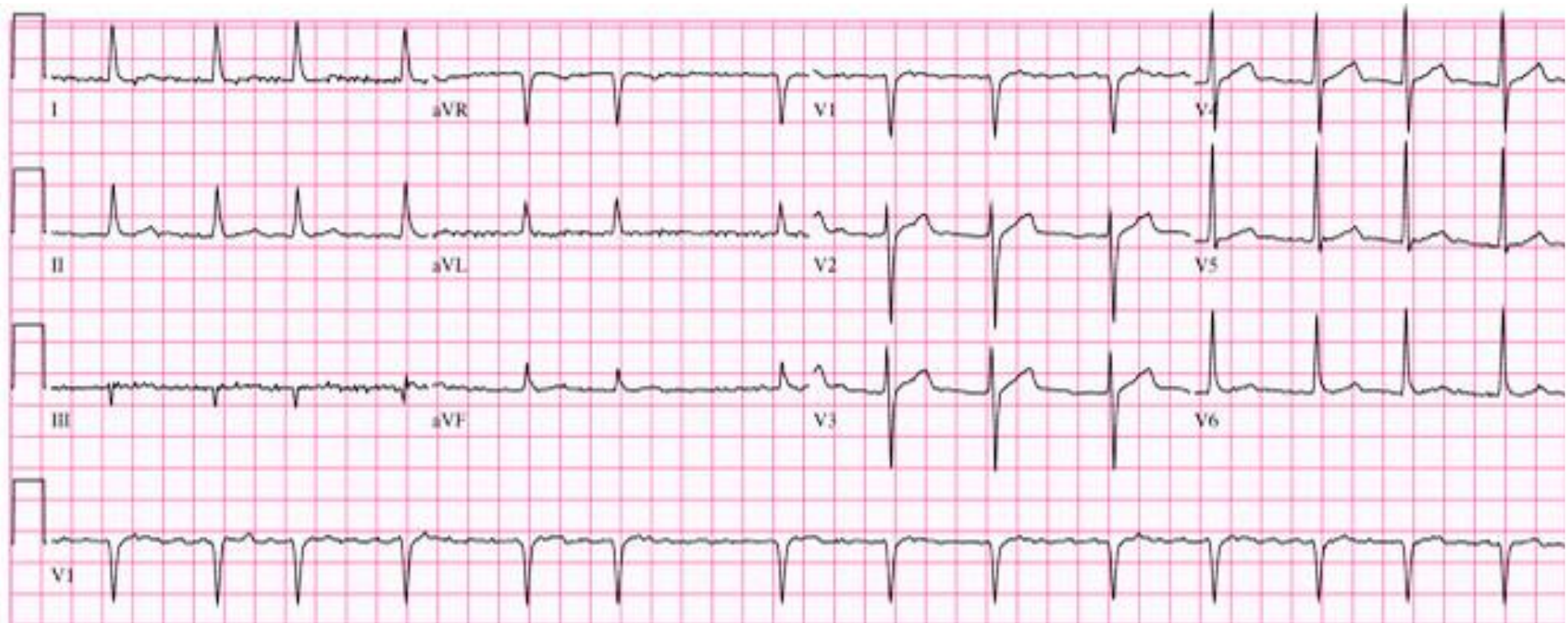


MASSACHUSETTS  
GENERAL HOSPITAL

CORRIGAN MINEHAN  
HEART CENTER

# Clinical Case

## 12-Lead ECG



# Clinical Case

## Echocardiogram

- Left ventricular ejection fraction 66% with no ventricular hypertrophy or wall motion abnormalities.
- No hemodynamically significant valvular disease.

## Exercise Stress Test

- Good exercise capacity (6 METS).
- Adequate workload (88% age-predicted maximum heart rate).
- No ECG evidence of ischemia.



MASSACHUSETTS  
GENERAL HOSPITAL

CORRIGAN MINEHAN  
HEART CENTER

# Question 1

## **Which of the following represents the most appropriate strategy for AF management in this patient?**

- 1) Initiation of oral anticoagulant and a rate control strategy with beta blocker or calcium channel blocker.
- 2) Initiation of oral anticoagulant and a rhythm control strategy (anti-arrhythmic drug or ablation procedure).
- 3) Initiate a platelet inhibitor (aspirin) and a rate control strategy with beta blocker or calcium channel blocker.
- 4) Initiate a platelet inhibitor (aspirin) and a rhythm control strategy (anti-arrhythmic drug or ablation procedure).



# Question 1

## Which of the following represents the most appropriate strategy for AF management in this patient?

- 1) Initiation of oral anticoagulant and a rate control strategy with beta blocker or calcium channel blocker.
- 2) **Initiation of oral anticoagulant and a rhythm control strategy (anti-arrhythmic drug or ablation procedure).**
- 3) Initiate a platelet inhibitor (aspirin) and a rate control strategy with beta blocker or calcium channel blocker.
- 4) Initiate a platelet inhibitor (aspirin) and a rhythm control strategy (anti-arrhythmic drug or ablation procedure).

# Outline

- 1) **Clinical Case**
- 2) **Epidemiology of AF**
- 3) **Physiology of AF**
- 4) **Advances in medical and interventional treatment:**
  - *Reduce the risk of AF-related stroke*
  - *Minimize consequences of AF on heart function (symptoms, CHF)*
- 5) **Current recommendations for management of AF**

# Prevalence of AF/AFL

## **AF is the most common arrhythmia in adults.**

- 1-2% of people in the US are affected.
- 50 million people affected worldwide in 2020.<sup>1</sup>

## **Incidence of AF increases with age.**

- 9% of people over 65 are affected.
- Prevalence expected to double between 2010 and 2030.<sup>1</sup>

*(From 5 to 12 million people in the US)*

1. Joglar et al., Circulation 2024;1:e1-e156.
2. Ball et al., Int J Cardiol 2013;167:1807-1824.

# Question 2

Which of the following are correct regarding consequences of AF:

1. AF increases the risk of death  $\geq$  3-fold and increases risk of stroke 3-fold.
2. AF increases the risk of death  $\geq$  5-fold and increases risk of stroke 1.5-fold.
3. AF increases the risk of death  $\geq$  1.5-fold and increases risk of stroke 5-fold.
4. AF increases the risk of death  $\geq$  2 fold and increases risk of stroke 4-fold.

# Question 2

Which of the following are correct regarding consequences of AF:

1. AF increases the risk of death  $\geq 3$ -fold and increases risk of stroke 3-fold.
2. AF increases the risk of death  $\geq 5$ -fold and increases risk of stroke 1.5-fold.
3. AF increases the risk of death  $\geq 1.5$ -fold and increases risk of stroke 5-fold.
4. AF increases the risk of death  $\geq 2$  fold and increases risk of stroke 4-fold.

# Impact of AF/AFL on Patient Outcomes

## **Morbidity and mortality is higher in patients with AF than in patients without AF.**

- AF is associated with a 1.5- to 2-fold increase in risk of death.<sup>1</sup>
- Worse outcomes in AF are driven by increased stroke risk.<sup>2</sup>

## **Presence of AF increases risk of stroke by 500%.**

- Almost 20% of all strokes occur in patients with AF.
- AF-related stroke mortality is 50% higher than for strokes not due to AF.<sup>3,4</sup>

## **Many people with AF are asymptomatic.**

- Stroke could be the first presenting sign of AF.

1. Joglar et al., Circulation 2024;1:e1-e156.
2. Alkhouli et al., JACC Clin EP 2018;4:618-625.
3. Lamassa et al., Stroke 2001;32:392-398.
4. Steger et al. Eur Heart J 2004;25:1734-1740.

# AF-Related Treatment Costs

## **AF is associated with higher health care utilization.**

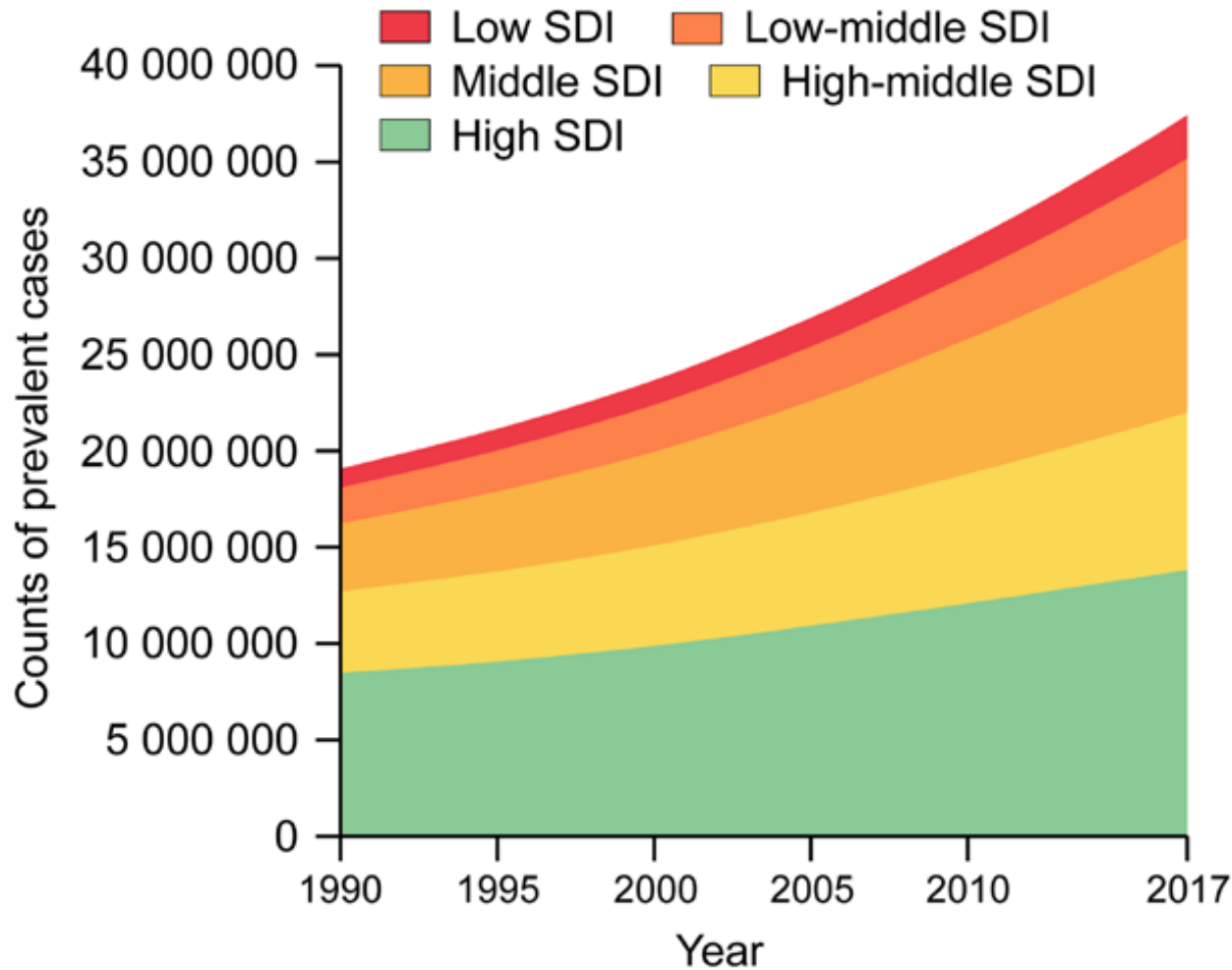
- Increased risk of ED visits and inpatient admissions [RR 2.41].<sup>1,2</sup>
- Increased health care costs per annum (\$63K vs. \$27K for non-AF).<sup>1,2</sup>

## **AF-related treatment costs exceed \$28 billion per annum.<sup>3</sup>**

1. Bode and Ptaszek, Current Cardiology Reports 2021;23:179.
2. Desmukh et al., Heart Rhythm O2 2022;3:577-586.
3. Joglar et al., Circulation 2024;1:e1-e156.



# Prevalence of AF is Higher in People with Lower Social Demographic Index



Joglar et al., Circulation 2024;1:e1-e156.

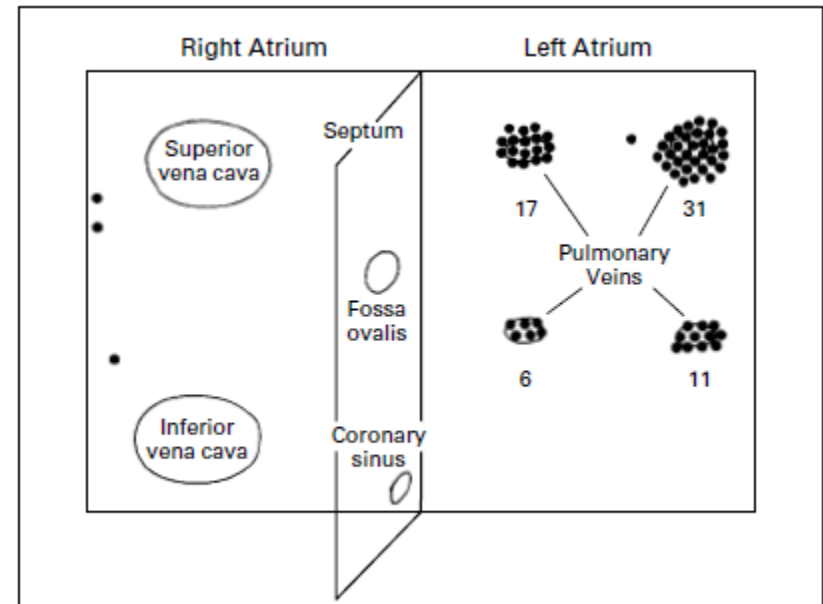
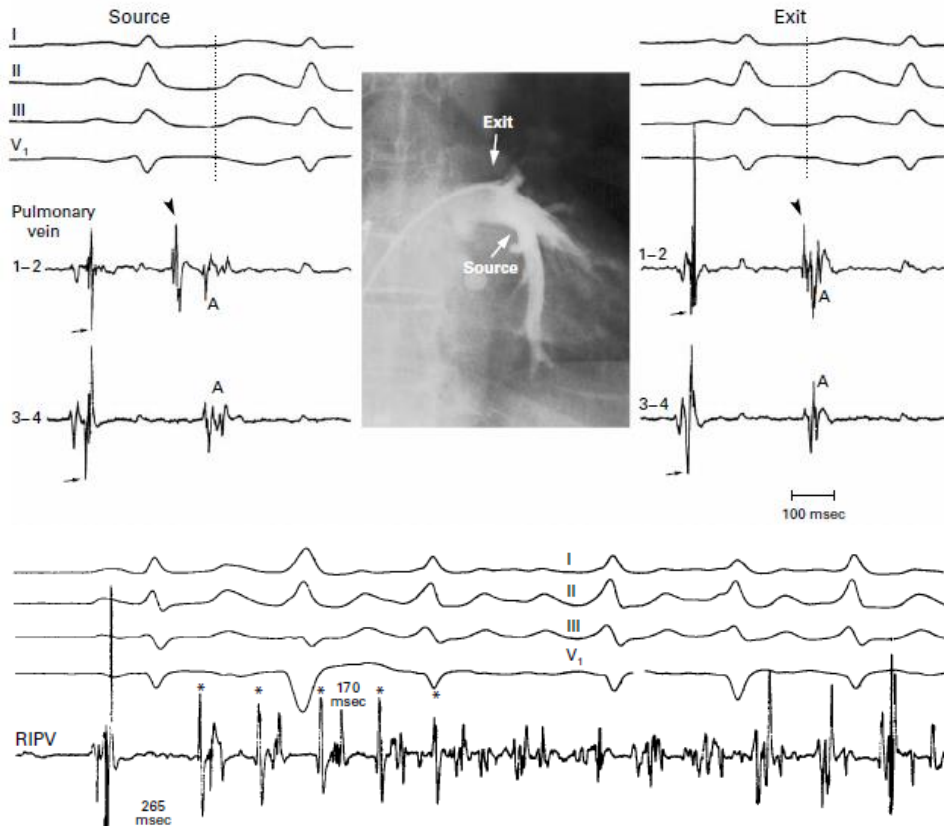


# Outline

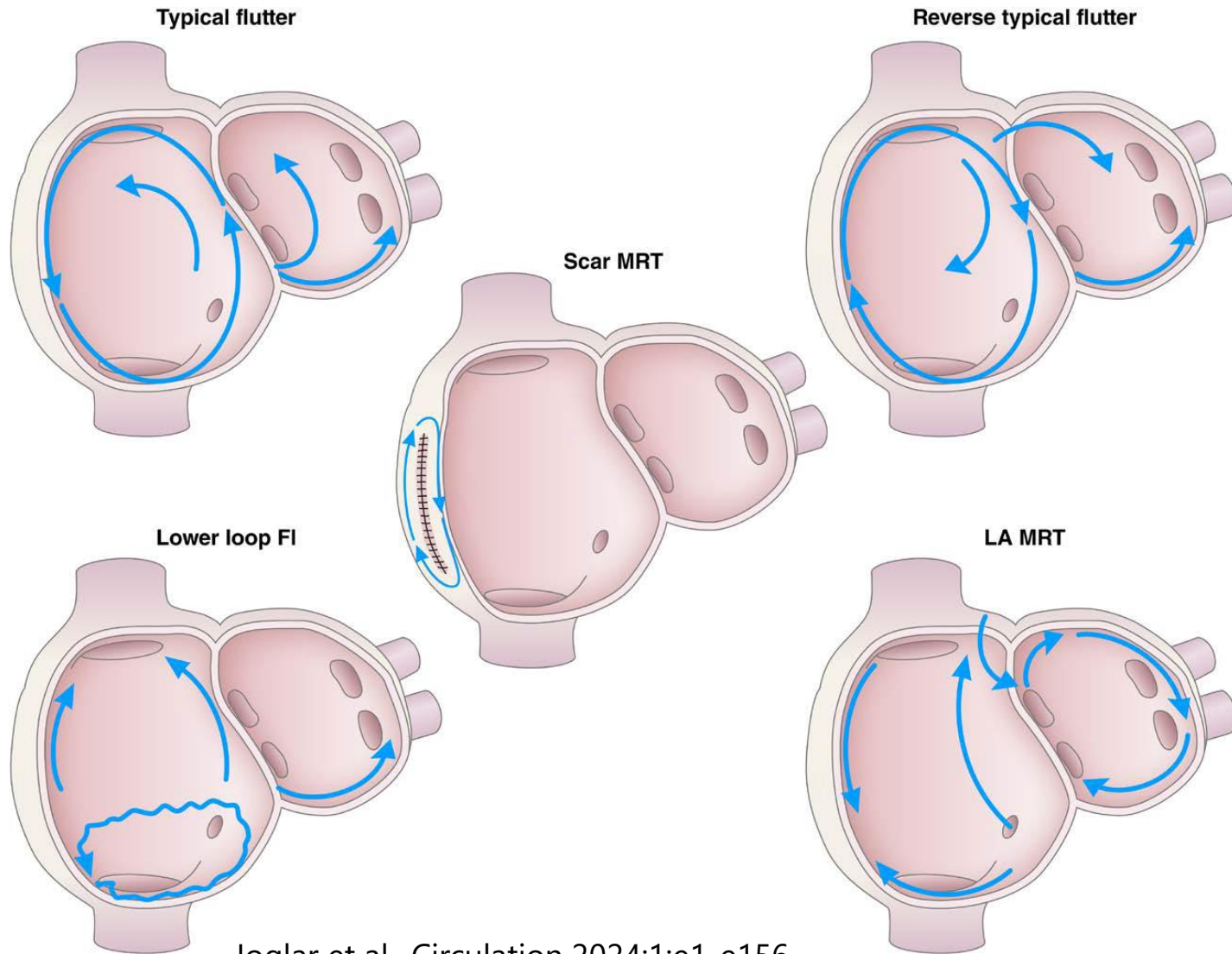
- 1) **Clinical Case**
- 2) **Epidemiology of AF**
- 3) **Physiology of AF**
- 4) **Advances in medical and interventional treatment:**
  - *Reduce the risk of AF-related stroke*
  - *Minimize consequences of AF on heart function (symptoms, CHF)*
- 5) **Current recommendations for management of AF**

# Pathophysiology of AF and AFL

AF is the result of rapid, chaotic electrical impulses emanating from multiple areas of the atria, notably the pulmonary veins.



# Types of AFL



# AF is Associated with Thrombus Formation in the Atria

## TEE With Contrast: Left Atrium



R Beigel et al., JACC Cardiovasc Imaging 2014;7:1251-1265.

TEE is the gold standard for LAA thrombus detection but cardiac CT and cardiac MR have excellent sensitivity/specificity and are viable alternatives to TEE (T Vira et al., Europace 2018;21:e1-e10).

# AF is Associated with Lower Cardiac Output and Can Precipitate Heart Failure

**Rapid and irregular heart rhythm in the context of AF can produce symptoms and can decrease quality of life.<sup>1</sup>**

**Sustained tachycardia due to AF can precipitate tachycardia-induced cardiomyopathy.<sup>2</sup>**

**AF and heart failure are both common: each makes the other more difficult to treat.<sup>3</sup>**

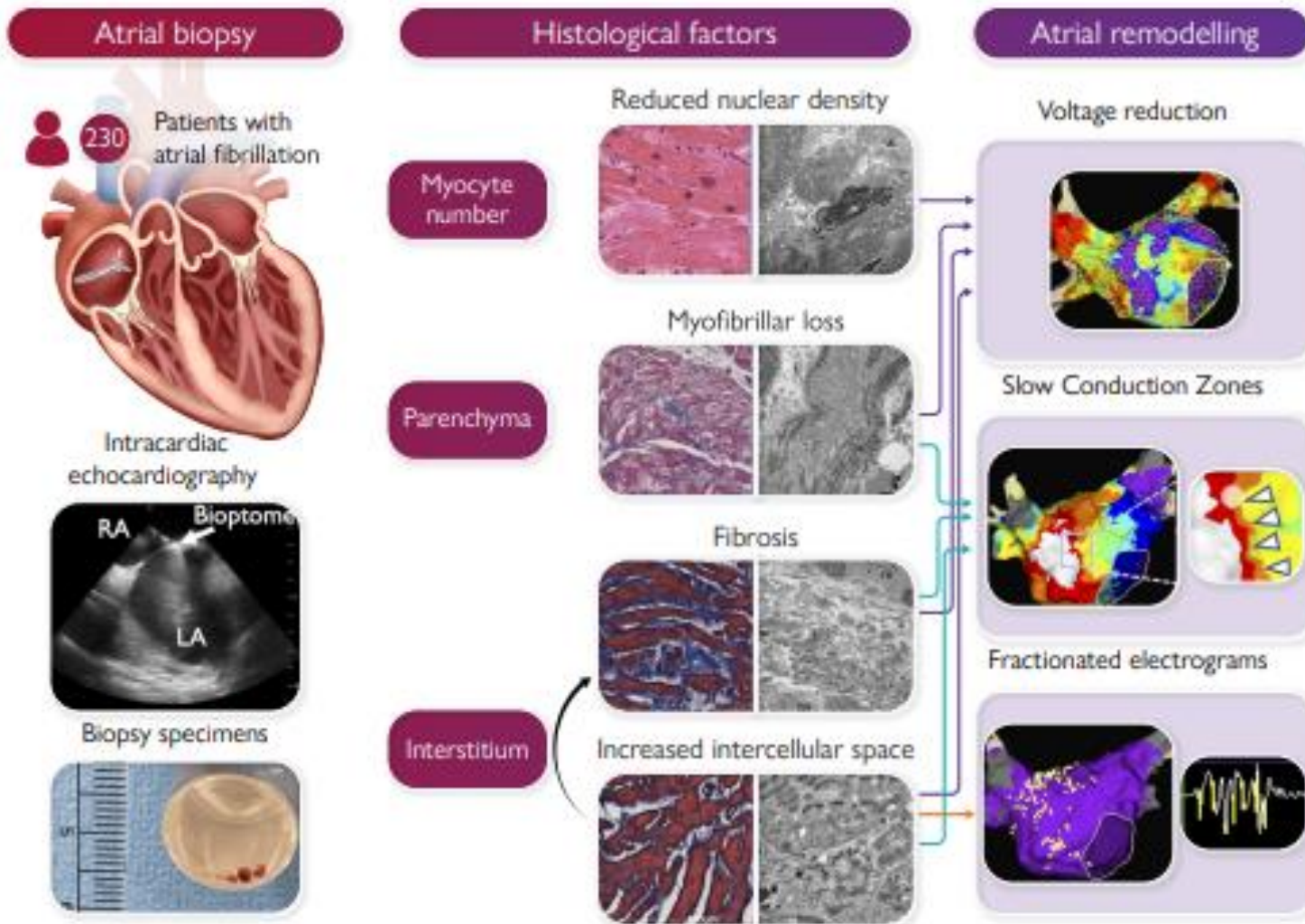
1. Ha et al., Circ Cardiovasc Qual Outcomes 2014;7:896-904.
2. Keefe et al., JACC Heart Fail 2024;12:605-615.
3. Joglar et al., Circulation 2024;1:e1-e156.

# Classification of AF

Category	Duration
<b>Paroxysmal</b>	Intermittent, terminates within $\leq 7$ days of onset
<b>Persistent</b>	Continuous for $> 7$ days and requires intervention
<b>Long-Standing Persistent</b>	Continuous for $\geq 12$ months

Joglar et al., Circulation 2024;1:e1-e156.

# AF-Related Atrial Remodeling



Takahashi et al., European Heart J 2023;44:3339-3353.

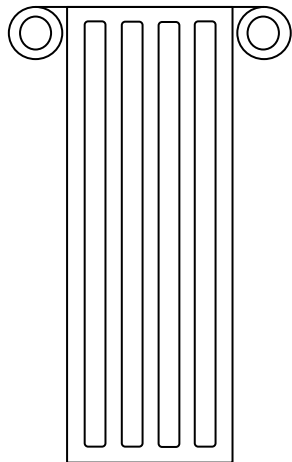
# Outline

- 1) **Clinical Case**
- 2) **Epidemiology of AF**
- 3) **Physiology of AF**
- 4) **Advances in medical and interventional treatment:**
  - *Reduce the risk of AF-related stroke*
  - *Minimize consequences of AF on heart function (symptoms, CHF)*
- 5) **Current recommendations for management of AF**

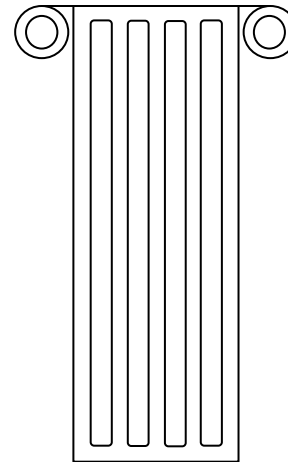


# Two “Pillars” of AF Management

**Reduce Risk of  
AF-Related  
Stroke**



**Minimize  
Hemodynamic  
Consequences of AF**



# Stroke Prevention in Non-Valvular AF

- **Oral anticoagulant (OAC) therapy is the cornerstone of stroke risk reduction.<sup>1</sup>**
- **Appropriate use of OAC can reduce the risk of AF-related stroke by ~65%.**
- **Guidelines support the use of OAC for stroke risk reduction in qualifying patients with NVAf.<sup>2,3</sup>**
  - *Risk for AF-related stroke can be estimated with metrics (e.g., CHA2DS2-VASC).*

1. Alkhouli et al., JACC 2018;71:2790-2801.

2. Kirchof et al., Eur Heart J 2016;37:2893-2962.

3. Joglar et al., Circulation 2024;149:e1-e156.

# Identifying Patients who Qualify for OAC: Calculating Risk of NVAf-Related Stroke

## CHADS-VASC score

Congestive heart failure +1

HTN +1

Age  $\geq$  65 +1

Age  $\geq$  75 +1

Diabetes +1

Stroke/TIA/thromboembolism +2

Sex (female) +1

Vascular disease (peripheral or CAD) +1

→ **Score = 1:** 0.6% CVA/year

→ **Score  $\geq$  2:**  $\geq$  2.2% CVA/year



MASSACHUSETTS  
GENERAL HOSPITAL

CORRIGAN MINEHAN  
HEART CENTER

# OAC Therapy for Patients with AF

## **Vitamin K Antagonists (e.g., warfarin)**

- Reduce stroke risk by 65%

## **DOACs (e.g., apixaban)**

- Superior stroke prevention than coumadin
- Lower bleeding risk than coumadin<sup>1</sup>

## **Anti-platelets (e.g., aspirin, clopidogrel)**

- Effectiveness debated<sup>2</sup>
- 19% stroke risk reduction (8 trials, >4,000 patients)

1. Hsu et al. 2018 Clin Pharmacol Ther 104:301-310.

2. Hsu et al. 2016 JACC 67:2913-2923.



# OAC Therapy for Patients with AF

## Vitamin K Antagonists (e.g., warfarin)

- Reduce stroke risk by 65%

## → DOACs (e.g., apixaban)

- Superior stroke prevention than coumadin
- Lower bleeding risk than coumadin<sup>1</sup>

## Anti-platelets (e.g., aspirin, clopidogrel)

- Effectiveness debated<sup>2</sup>
- 19% stroke risk reduction (8 trials, >4,000 patients)

1. Hsu et al. 2018 Clin Pharmacol Ther 104:301-310.

2. Hsu et al. 2016 JACC 67:2913-2923.



# OAC Therapy for Patients with AF

## **Outcomes for patients with AF receiving OAC are better even if they have a stroke:**

Severity and in-hospital mortality is reduced for patients receiving therapeutic coumadin or DOAC.<sup>1</sup>

## **Lack of OAC is associated with higher risk of:**

Initial stroke (OR 2.95)

Recurrent stroke (OR 2.8)

All-cause death (OR 2.75).<sup>2</sup>

1. Xian et al., JAMA 2017 ;317:1057-1067.

2. Mazurek et al., Stroke 2017;48:2198-2205.



# Question 3

Which of the following scenarios describes appropriate utilization of oral anticoagulation:

1. Use of eliquis in an 80-year-old woman with persistent AF and rheumatic mitral stenosis.
2. Use of coumadin in an 80-year-old woman with persistent AF and rheumatic mitral stenosis.
3. Use of eliquis in an 80-year-old woman with persistent AF and a mechanical aortic valve replacement.
4. Use of coumadin in an 80-year-old woman with persistent AF and a mechanical aortic valve replacement.
5. Choices 2 and 4.

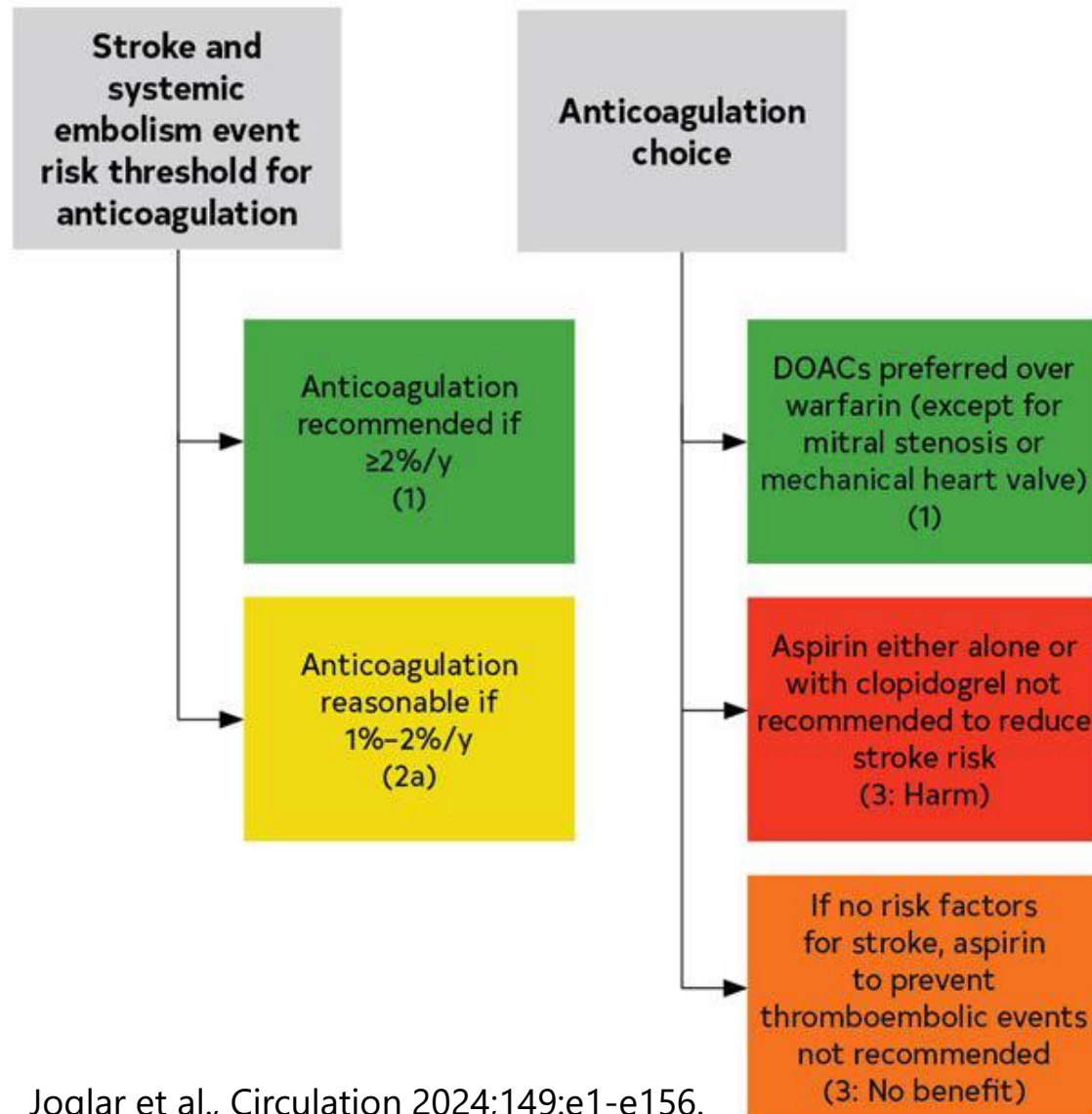
# Question 3

Which of the following scenarios describes appropriate utilization of oral anticoagulation:

1. Use of eliquis in an 80-year-old woman with persistent AF and rheumatic mitral stenosis.
2. Use of coumadin in an 80-year-old woman with persistent AF and rheumatic mitral stenosis.
3. Use of eliquis in an 80-year-old woman with persistent AF and a mechanical aortic valve replacement.
4. Use of coumadin in an 80-year-old woman with persistent AF and a mechanical aortic valve replacement.
5. **Choices 2 and 4.**

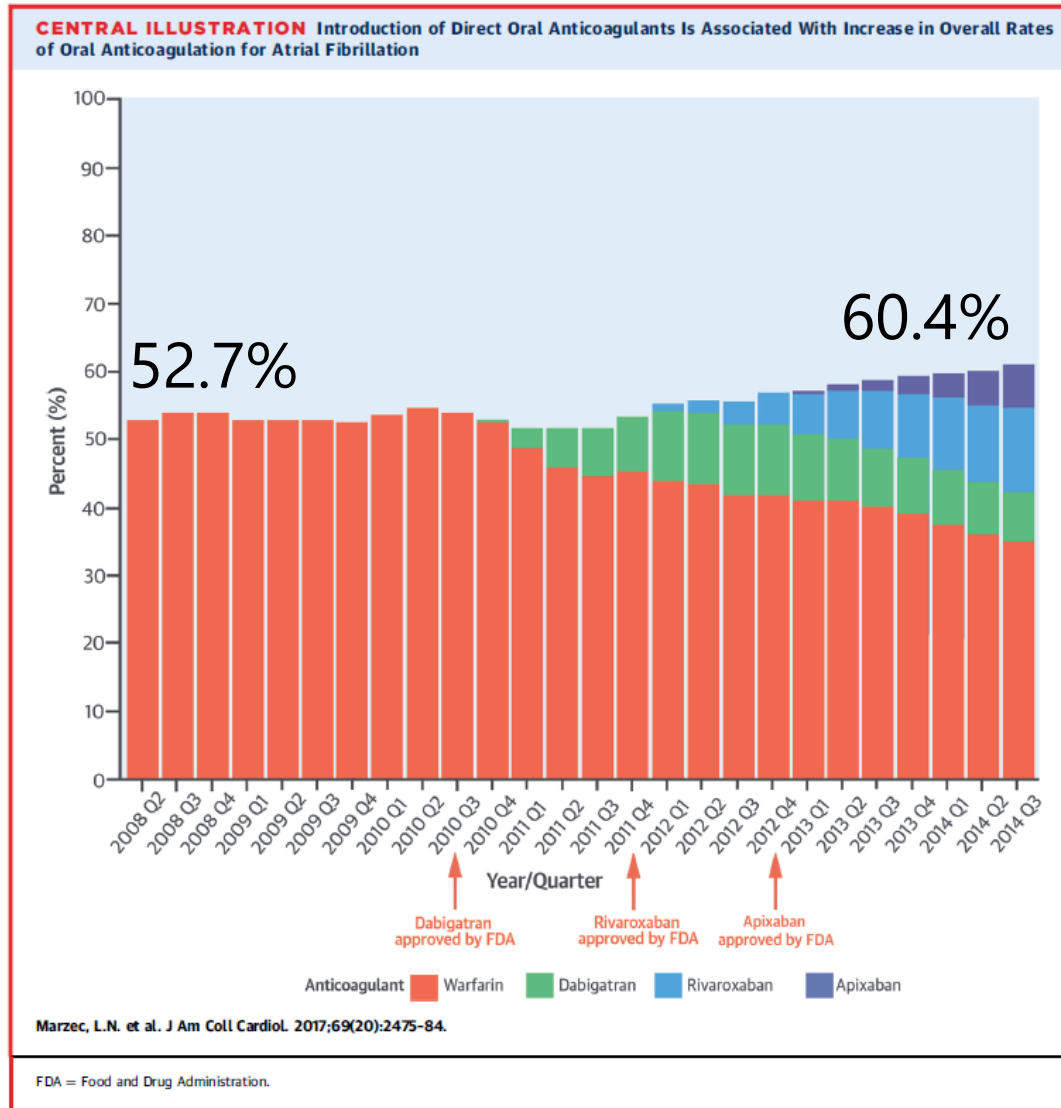


# Anticoagulation Options in AF



Joglar et al., Circulation 2024;149:e1-e156.

# Underutilization of Anticoagulation in the US



Marzec et al., JACC 2017;69:2475-2484.

# Primary Reason for Withholding Anticoagulant: Bleeding Risk

Bleeding risk associated with OAC use: HAS-BLED score

**HTN** (+1)

**A**bnormal liver fxn (+1)

**A**bnormal renal fxn (+1)

**S**troke/TIA (+1)

**B**leeding predisposition (+1)

**E**lderly: Age  $\geq$  65 (+1)

**D**rugs (anti-platelet) (+1)

**D**rugs (alcohol) (+1)



**Score 1:** 1% bleed/yr



**Score 2:** 1.9% bleed/yr



**Score 3:** 3.7% bleed/yr



**Score 4:** 8.7% bleed/yr



**Score 5:** >10% bleed/yr



MASSACHUSETTS  
GENERAL HOSPITAL

CORRIGAN MINEHAN  
HEART CENTER

# Underutilization of OAC: Stroke Risk and Bleeding Risk Rise Together

## CHADS-VASC

**C**ongestive heart failure (+1)

**H**TN (+1)

**A**ge  $\geq$  65 (+1)

**A**ge  $\geq$  75 (+1)

**D**iabetes (+1)

**S**troke/TIA (+2)

**S**ex (female) (+1)

**V**ascular disease (+1)

## HAS-BLED

**H**TN (+1)

**A**bnormal liver funct (+1)

**A**bnormal renal funct (+1)

**S**troke/TIA (+1)

**B**leeding predisposition (+1)

**E**lderly: Age  $\geq$  65 (+1)

**D**rugs (anti-platelet) (+1)

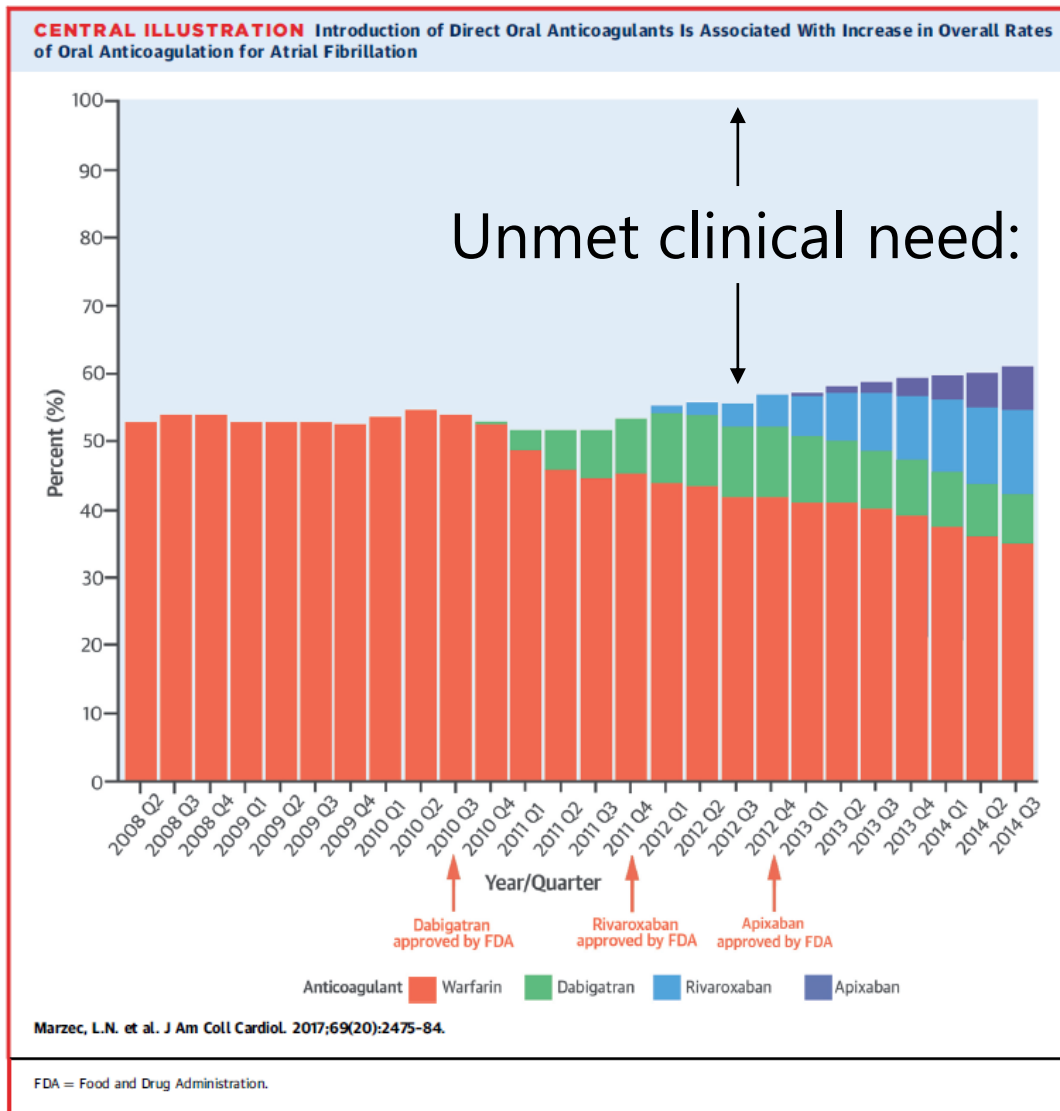
**D**rugs (alcohol) (+1)



MASSACHUSETTS  
GENERAL HOSPITAL

CORRIGAN MINEHAN  
HEART CENTER

# Underutilization of Anticoagulation in the US



# Question 4

What is the most appropriate stroke prevention strategy for: 82-year-old woman with AF and prior stroke, Eliquis 5mg twice daily not tolerated due to gait instability and repeated falls with subdural hemorrhages:

1. Aspirin 81mg daily
2. Eliquis 2.5mg twice daily
3. Surgical ligation of the left atrial appendage
4. Percutaneous left atrial appendage occlusion

# Question 4

What is the most appropriate stroke prevention strategy for: 82-year-old woman with AF and prior stroke, Eliquis 5mg twice daily not tolerated due to gait instability and repeated falls with subdural hemorrhages:

1. Aspirin 81mg daily
2. Eliquis 2.5mg twice daily
3. Surgical ligation of the left atrial appendage
4. Percutaneous left atrial appendage occlusion

# Thrombi that Cause AF-Related Stroke Usually Emanate from the Left Atrial Appendage

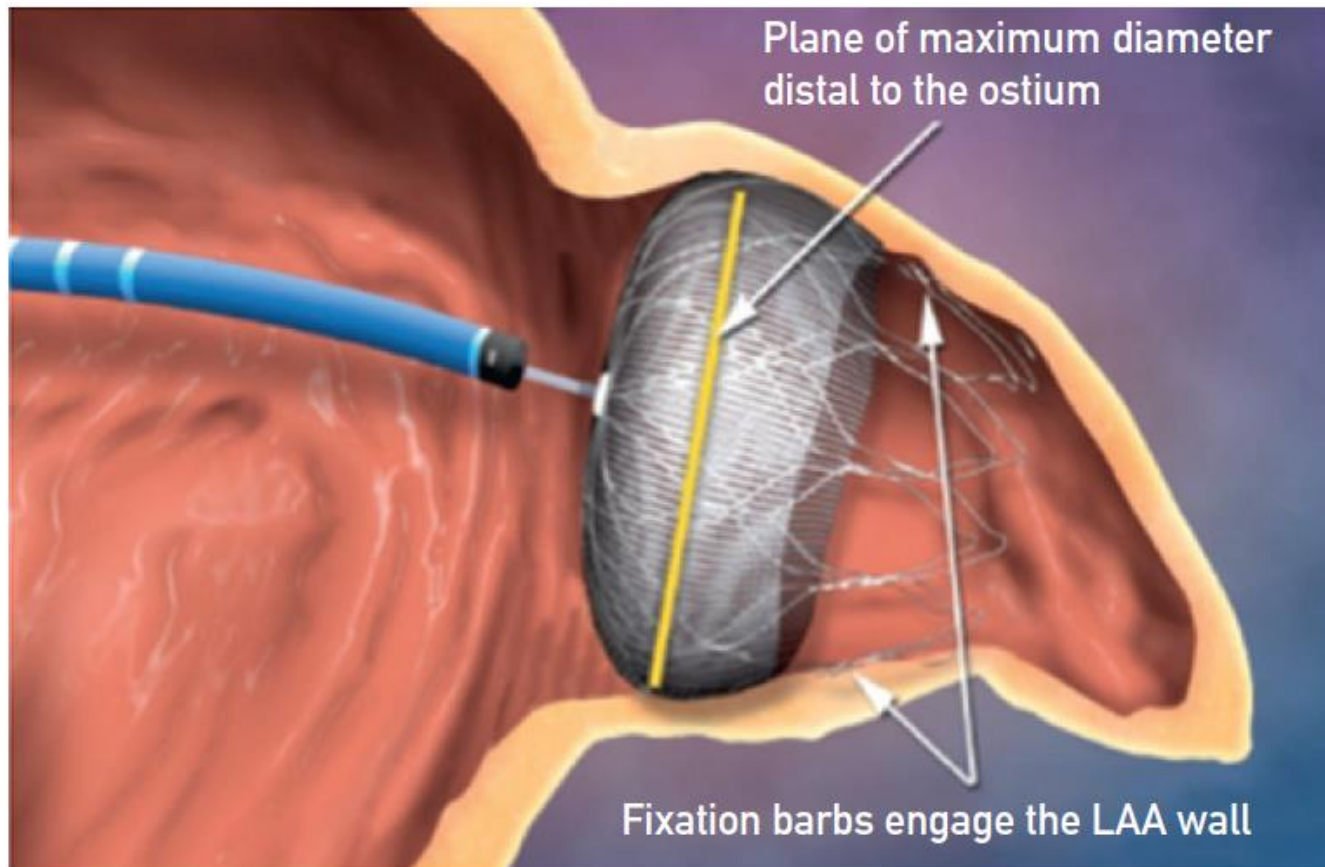
Study	# of patients	Thrombus in LAA	Thrombus in LA cavity
Stoddard et al. 1995 JACC 25:452-459	317	66	1
Manning et al. 1994 Circ 90:1-21a	233	34	1
Aberg et al. 1969 Acta Med Scand 185:373-379	506	35	12
Tsai et al. 1990 J Formosan Med Assoc 89:270-274	52	2	2
Brown et al. 1993 Int J Card Imaging 9:65-72	48	12	1
Manning et al. 1994 Circ 90:1202a	171	8	3
Klein et al. 1994 Circ 90 (Suppl 1):21a	359	19	1
Leung et al. 1994 JACC 24:755-762	272	19	0
Hart et al. 1994 Stroke 25:1337-1341	60	6	0
<b>Total</b>	<b>1,288</b>	<b>201 (91%)</b>	<b>21</b>

Blackshear JL et al., Ann Thorac Surg 1996;61:755-759

Odell JA et al., Ann Thorac Surg 1996;61:565-569



# Watchman Device Deployed in the LAA



Holmes DR et al., Mayo Clinic Proc 2019;94:864-874.

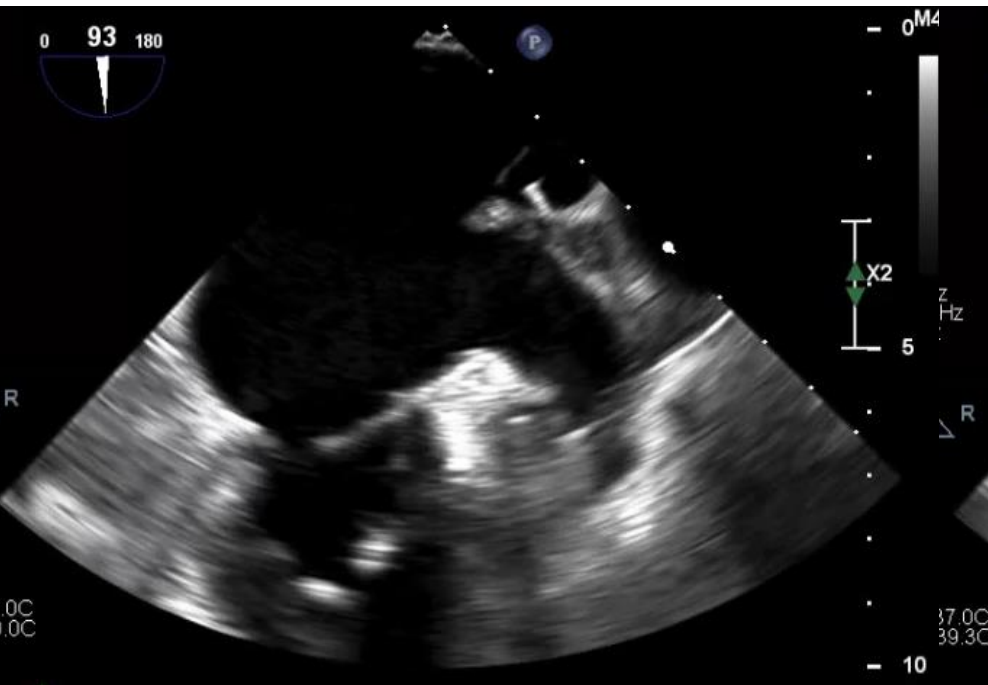


MASSACHUSETTS  
GENERAL HOSPITAL

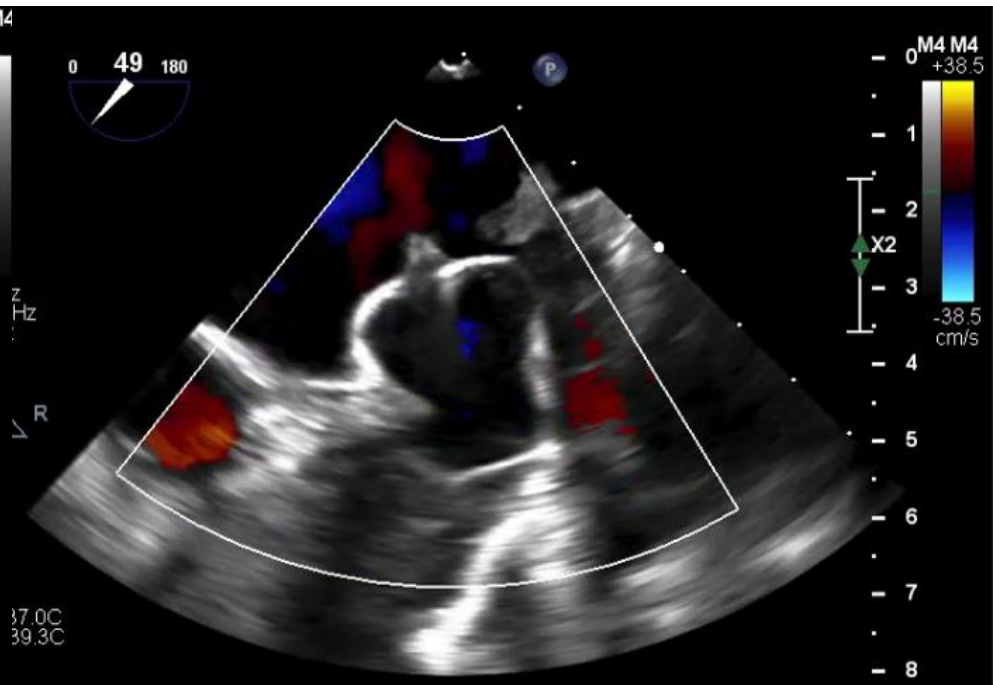
CORRIGAN MINEHAN  
HEART CENTER

# TEE Visualization of Watchman Device in the Left Atrial Appendage (TEE)

Before Watchman



After Watchman



# Watchman Clinical Trials

## Randomized Clinical Trials: Watchman vs Coumadin

- PROTECT AF (NCT00129545)<sup>1</sup>
  - Watchman non-inferior and superior to coumadin for mortality, MACE
- PREVAIL (NCT01182441)<sup>2</sup>
  - Watchman non-inferior to coumadin except for cardiovascular/unexplained death

1. Holmes DR et al., Lancet 2009;374:534-542.

2. Reddy V et al., Stroke 2018;49:1464-1470.

# Cost Effectiveness of LAA Occlusion

- Based on the pivotal trials, LAA occlusion is cost-saving compared with DOACs in 5 years and coumadin in 10 years.<sup>1</sup>
- LAA occlusion is the lowest cost therapy at 5 years (patient out-of-pocket costs).<sup>2</sup>
- LAA occlusion is the lowest cost therapy at 10 years (Medicare).<sup>1</sup>

1. Reddy et al., JACC 2015;66:2728-2739.

2. Reddy et al., JACC 2017;70:880.

# Ongoing/Future Clinical Trials: Watchman vs DOACs

Trial (identifier)	Subpopulation with NVAF	LAAO device	Control group
OPTION (NCT03795298)	1,600 patients undergoing ablation	WATCHMAN FLX	Direct OAC
CHAMPION-AF (NCT04394546)	3,000 patients eligible for long-term OAC	WATCHMAN FLX	Direct OAC
CATALYST (NCT04226547)	2,650 patients eligible for long-term OAC	Amulet	Direct OAC
OCCLUSION-AF (NCT03642509)	750 patients eligible for long-term OAC	WATCHMAN FLX, Amulet	Direct OAC
ASAP-TOO (NCT02928497)	888 patients with a contraindication to OAC	WATCHMAN 2.5	Aspirin or no medication
CLOSURE-AF (NCT03463317)	1,512 patients at high risk of bleeding or with a contraindication to OAC	Any device with a CE mark	Direct OAC or VKA
COMPARE-LAAO (NCT04676880)	609 patients with a contraindication to OAC	WATCHMAN FLX, Amulet	Antiplatelet or no medication
STROKECLOSE (NCT02830152)	750 patients with intracerebral haemorrhage in past 12 months	Amulet	No medication, OAC or antiplatelet
CLEARANCE (NCT04298723)	550 patients with intracerebral haemorrhage	WATCHMAN FLX	No medication, OAC or antiplatelet
LAA-KIDNEY (NCT05204212)	430 patients with end-stage renal disease (eGFR <15ml/min/1.73 m <sup>2</sup> )	Amulet	Best medical therapy
LAAOS-4 (NCT05963698)	4,000 patients with CHA <sub>2</sub> DS <sub>2</sub> -VAsc score of ≥4	WATCHMAN FLX plus direct OAC	Direct OAC alone

eGFR, estimated glomerular filtration rate; LAAO, left atrial appendage occlusion; NVAF, non-valvular atrial fibrillation; OAC, oral anticoagulation; VKA, vitamin K antagonist.

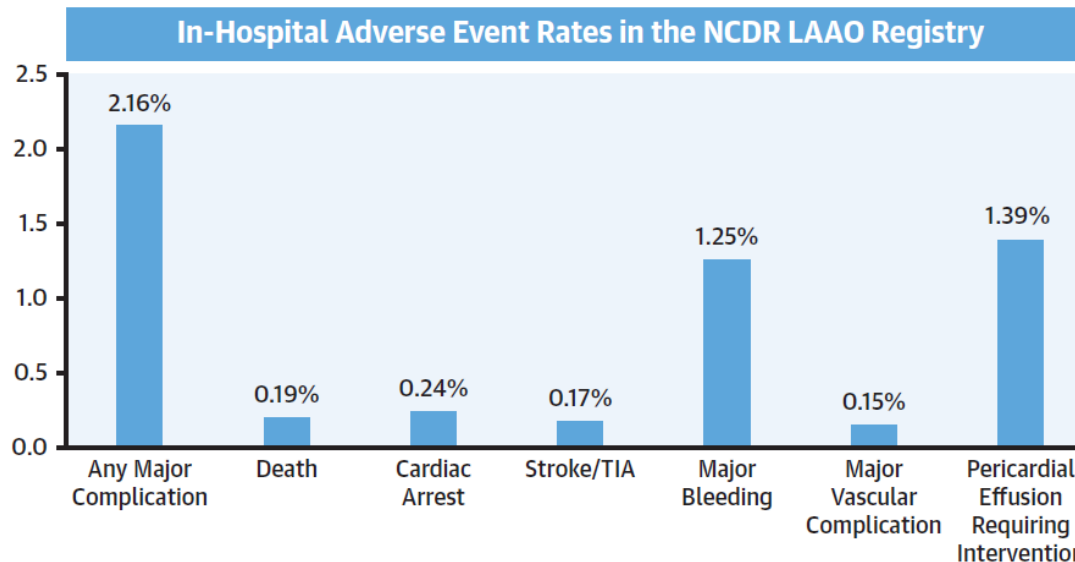
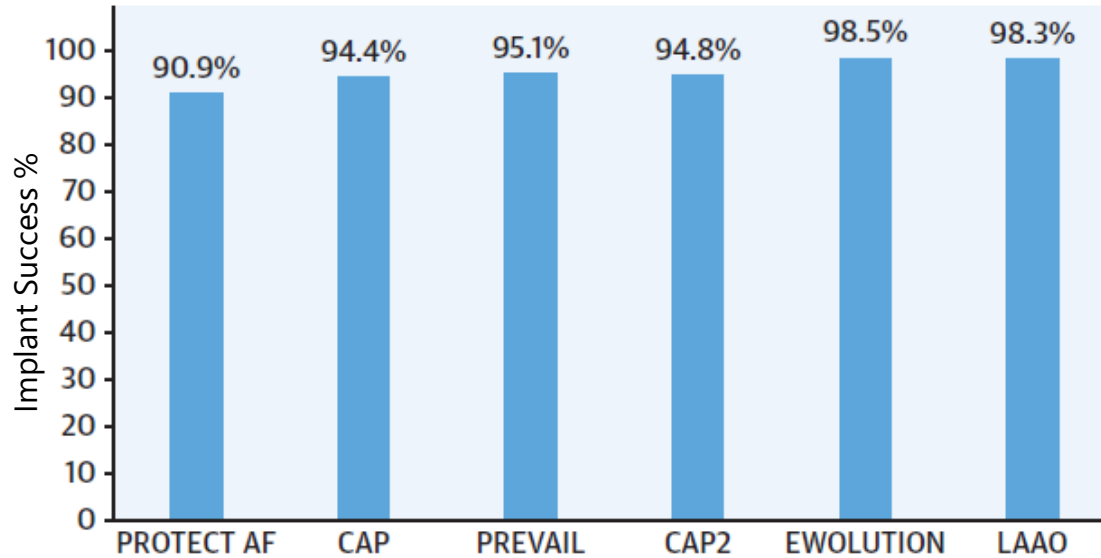
J Saw et al.,  
Nat Rev Cardiol  
2024;21:153-154



MASSACHUSETTS  
GENERAL HOSPITAL

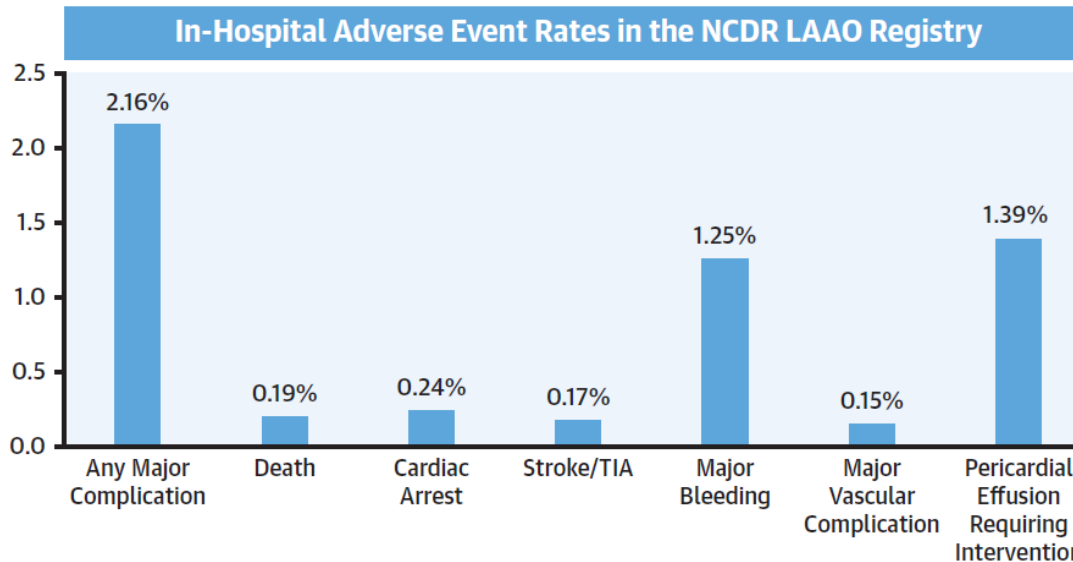
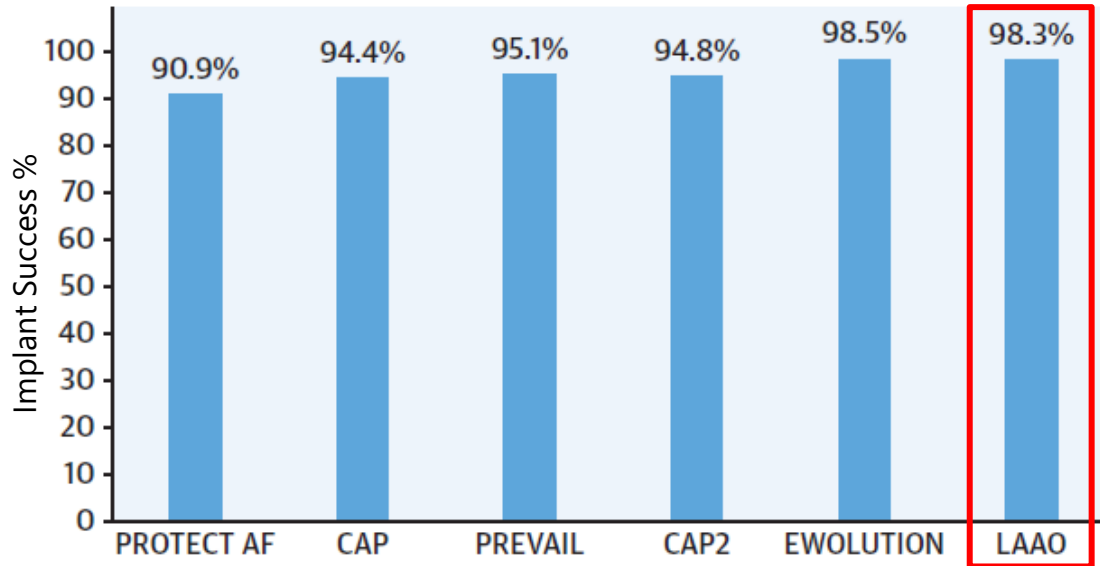
CORRIGAN MINEHAN  
HEART CENTER

# Watchman Implantation Outcomes: Clinical Trials versus Post-Approval Registry



Freeman JV et al.,  
J Am Coll Cardiol  
2020;75:1503-1518

# Watchman Implantation Outcomes: Clinical Trials versus Post-Approval Registry



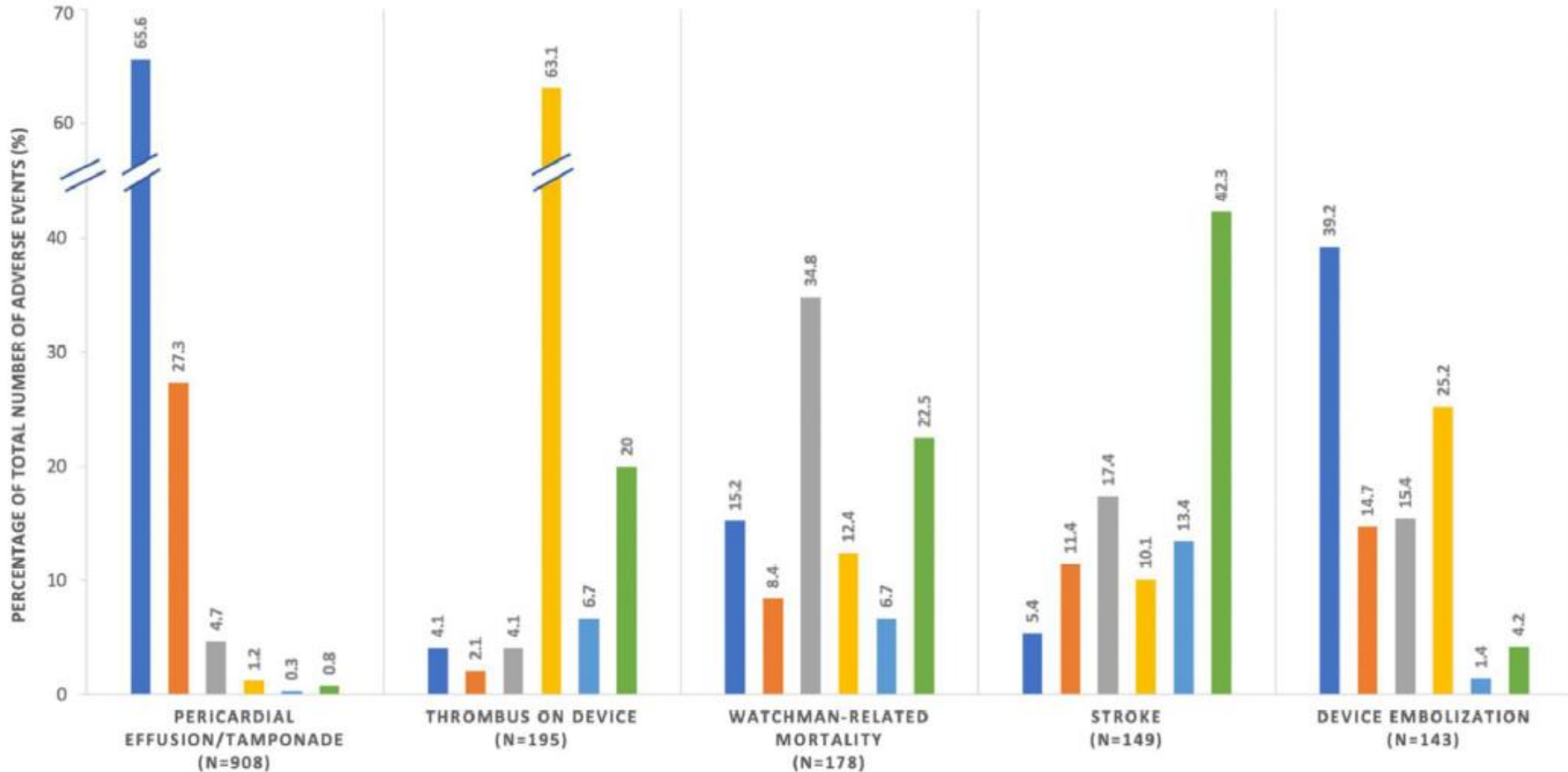
Freeman JV et al.,  
J Am Coll Cardiol  
2020;75:1503-1518

# Safety Considerations after Watchman Implantation: Analysis of the MAUDE Database

TIME COURSE OF COMMON WATCHMAN IMPLANTATION ADVERSE EVENTS (PERCENT OF TOTAL)

Manufacturer And User facility Device Experience database

■ Intraoperative ■ Same Day ■ 2-30 Days ■ 1-12 Months ■ >1 Year ■ Unspecified





# Incomplete LAA Occlusion

**The LAA orifice is typically oval, and the Watchman device is round, raising the possibility of blood leakage around the device.**

**Significant leak has been defined to be 1 to 5mm on TEE.**

**Leak >5 mm is an indication for continuation of OAC.**

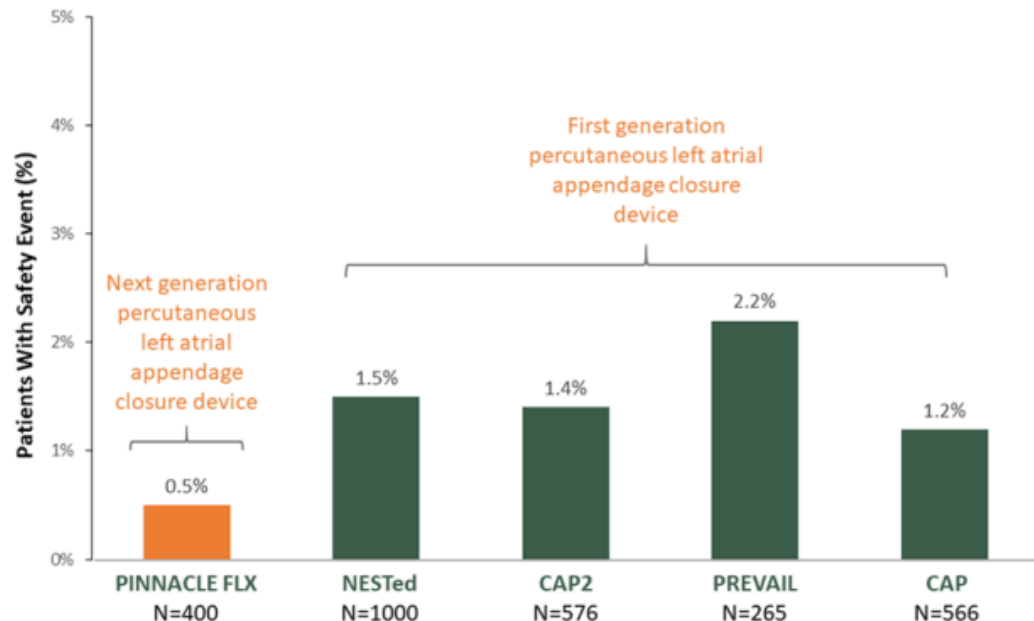
**Leak closure can also be considered in such cases.<sup>1</sup>**

# Next Generation Watchman Devices



## Watchman FLX:

- Decreased pericardial effusion and device embolism
- Improved device sealing
- Evaluated with PINNACLE FLX Trial (NCT02702271)



Kar et al., Circulation 2021;143:1754-1762.

# Percutaneous Approach to LAA Occlusion

<b>COR</b>	<b>LOE</b>	<b>Recommendations</b>
<b>2a</b>	<b>B-NR</b>	In patients with AF, a moderate to high risk of stroke (CHA2DS2-VASc score $\geq 2$ ), and a contraindication to long-term oral anticoagulation due to a nonreversible cause, percutaneous LAAO (pLAAO) is reasonable.
<b>2b</b>	<b>B-R</b>	In patients with AF and a moderate to high risk of stroke and a high risk of major bleeding on oral anticoagulation, pLAAO may be a reasonable alternative to oral anticoagulation based on patient preference, with careful consideration of procedural risk and with the understanding that the evidence for oral anticoagulation is more extensive.

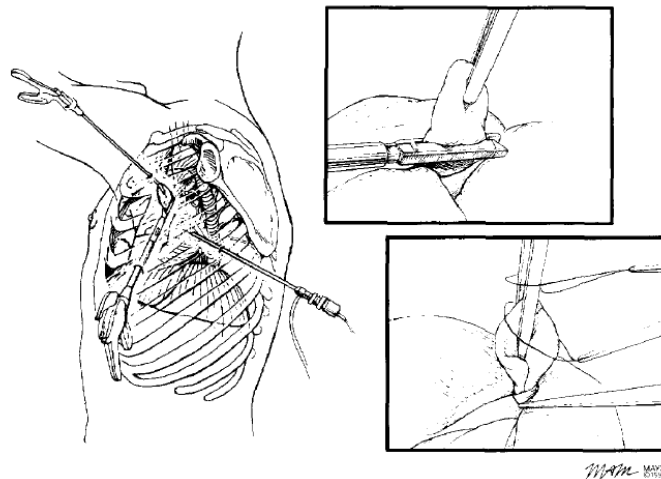
Joglar et al., Circulation 2024;149:e1-e156.

# Other LAA Occlusion Devices

- **Endocardial:**
  - Amplatzer Cardiac Plug (ACP)
  - Amulet
- **Epicardial/Surgical:**
  - AtriClip
  - Lariat

# Surgical LAA Occlusion

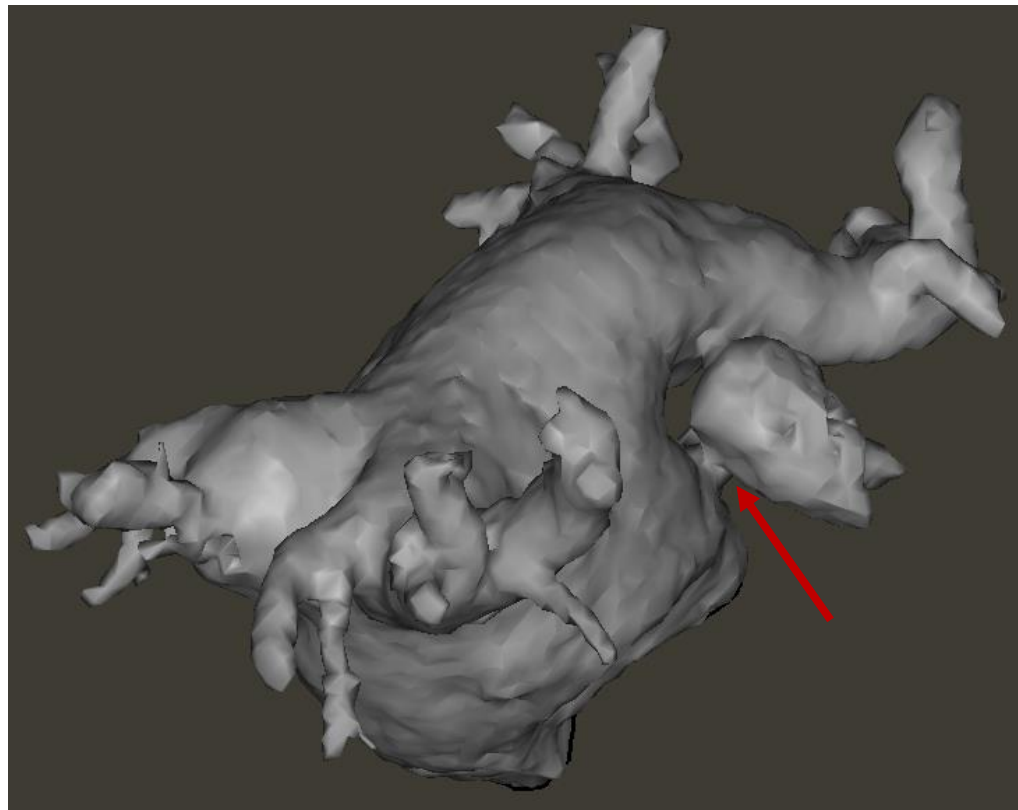
- Surgical LAA occlusion has been described and is typically performed in the context of other cardiac surgery (e.g., valve surgery, CABG).
- Techniques for closure vary, including oversew of the appendage versus true amputation.



Adapted from Odell JA et al., Ann Thoracic Surg 1996;61:565-569.

# Surgical LAA Occlusion

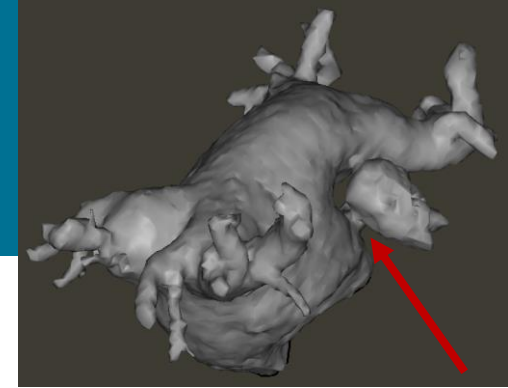
Incomplete closure is associated with thromboembolic events.



Aryana et al., Heart Rhythm 2015;12:1431-1437.

Ptaszek et al., J Innov Card Rhythm Mgmt 2013;4:1237-1241

# Surgical LAA Occlusion



COR	LOE	Recommendations
<b>1</b>	<b>A</b>	In patients with AF undergoing cardiac surgery with a CHA2DS2-VASc score $\geq 2$ or equivalent stroke risk, surgical LAA exclusion, in addition to continued anticoagulation, is indicated to reduce the risk of stroke and systemic embolism.
<b>1</b>	<b>A</b>	In patients with AF undergoing cardiac surgery and LAA exclusion, a surgical technique resulting in absence of flow across the suture line and a stump of $< 1$ cm as determined by intraoperative transesophageal echocardiography should be used.
<b>2b</b>	<b>A</b>	In patients with AF undergoing cardiac surgery with CHA2DS2-VASc score $\geq 2$ or equivalent stroke risk, the benefit of surgical LAA exclusion in the absence of continued anticoagulation to reduce the risk of stroke and systemic embolism is uncertain.

# Second “Pillar” of AF Management: Address Hemodynamic Consequences

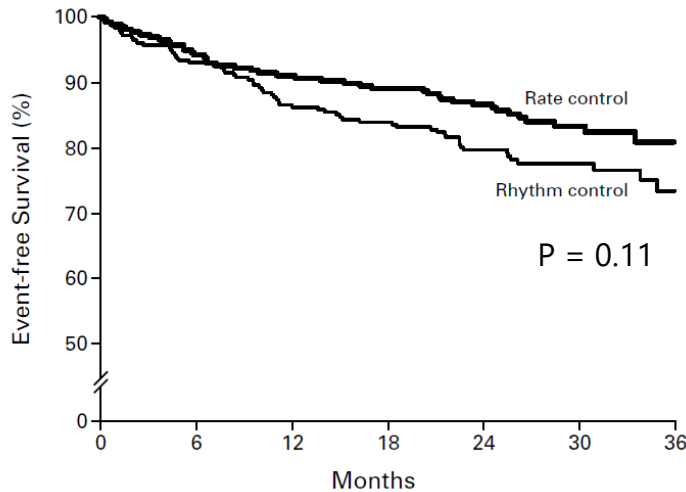
- **For each patient, the clinician must decide if a rate control strategy or a rhythm control strategy is most appropriate.**
- **In some situations, the decision to adopt a rhythm control strategy can be straightforward:**
  - *Adequate rate control cannot be achieved*
  - *Intolerable symptoms even with adequate rate control*
  - *Heart failure*
- **What about rate versus rhythm control in the general population of patients with AF?**



# Rate vs Rhythm Control: RACE and AFFIRM

The RACE and AFFIRM trials concluded rate control is non-inferior to rhythm control (anti-arrhythmic drugs and cardioversion only, ablation therapy was not included).

## RACE



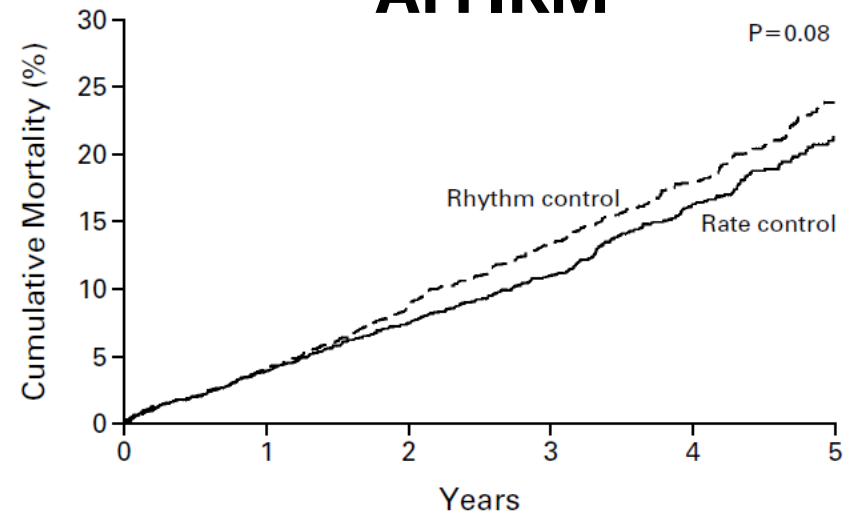
No. AT RISK	0	6	12	18	24	30	36
Rate control	256	239	232	222	212	99	25
Rhythm control	266	243	224	218	207	85	24

### NO. OF DEATHS

	0	1	2	3	4	5
Rhythm control	0	80 (4)	175 (9)	257 (13)	314 (18)	352 (24)
Rate control	0	78 (4)	148 (7)	210 (11)	275 (16)	306 (21)

IC Van Gelder et al., NEJM 2002;347:1834-1840.

## AFFIRM



AFFIRM Investigators, NEJM 2002;347:1825-1833



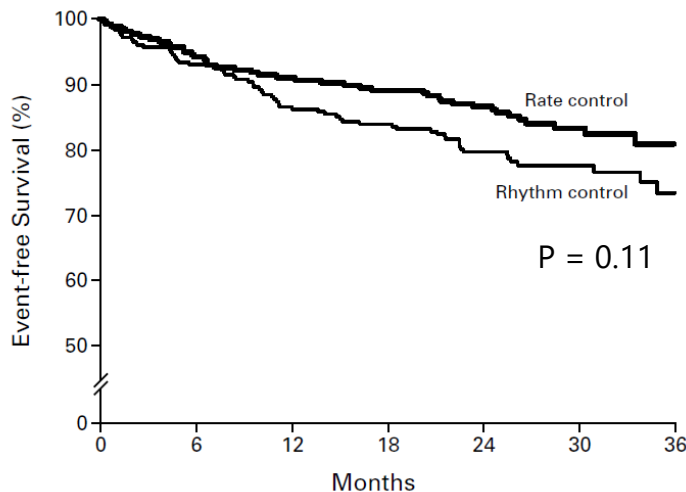
MASSACHUSETTS  
GENERAL HOSPITAL

CORRIGAN MINEHAN  
HEART CENTER

# Rate vs Rhythm Control: RACE and AFFIRM

The RACE and AFFIRM trials concluded rate control is non-inferior to rhythm control

## RACE



No. AT RISK	0	6	12	18	24	30	36
Rate control	256	239	232	222	212	99	25
Rhythm control	266	243	224	218	207	85	24

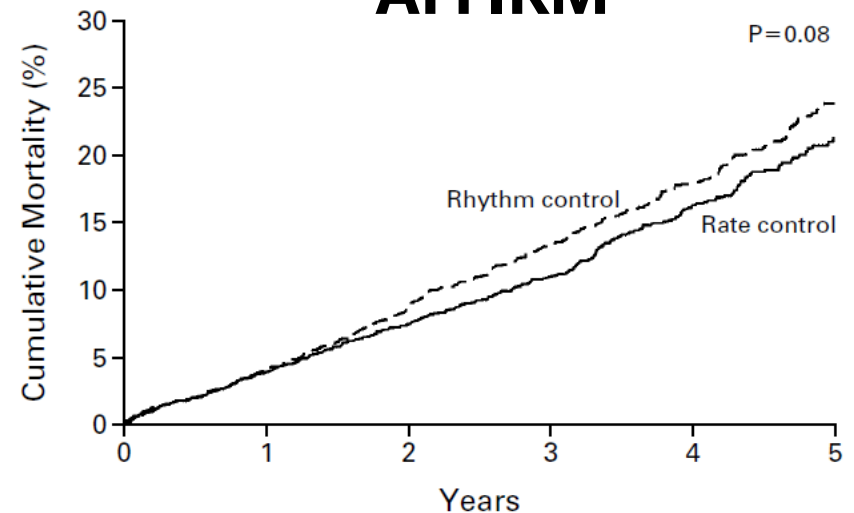
### NO. OF DEATHS

	0	1	2	3	4	5
Rhythm control	0	80 (4)	175 (9)	257 (13)	314 (18)	352 (24)
Rate control	0	78 (4)	148 (7)	210 (11)	275 (16)	306 (21)

IC Van Gelder et al., NEJM 2002;347:1834-1840.

From four weeks before until four weeks after electrical cardioversion, all patients received acenocoumarol or fenprocoumon (target international normalized ratio [INR], 2.5 to 3.5). If sinus rhythm was present at one month, the oral anticoagulant could be stopped or changed to aspirin (80 to 100 mg daily). Aspirin was also allowed

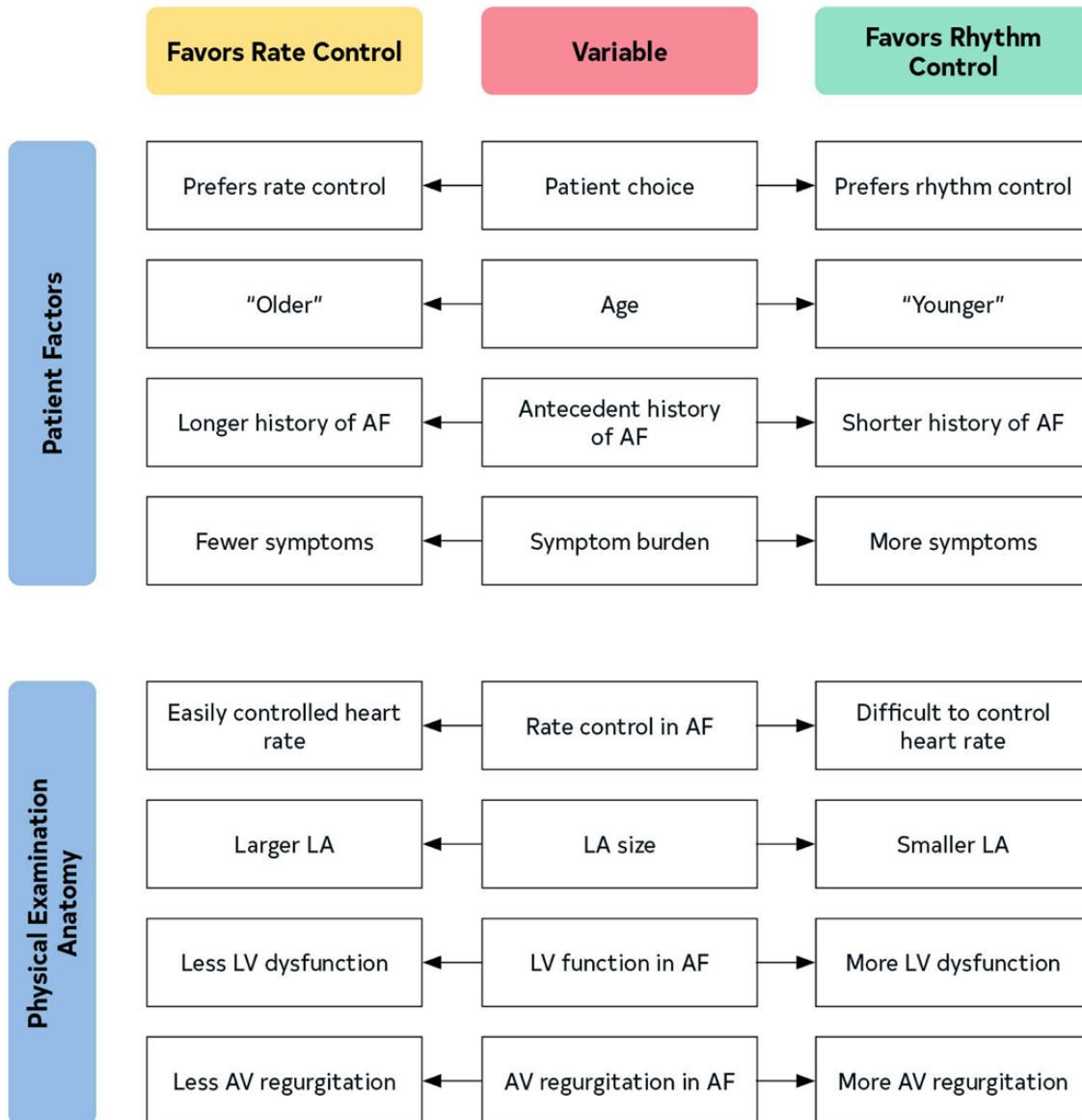
## AFFIRM



AFFIRM Investigators, NEJM 2002;347:1825-1833

international normalized ratio (INR) of 2.0 to 3.0. In the rhythm-control group, continuous anticoagulation was encouraged but could be stopped at the physician's discretion if sinus rhythm had

# Rate versus Rhythm Control



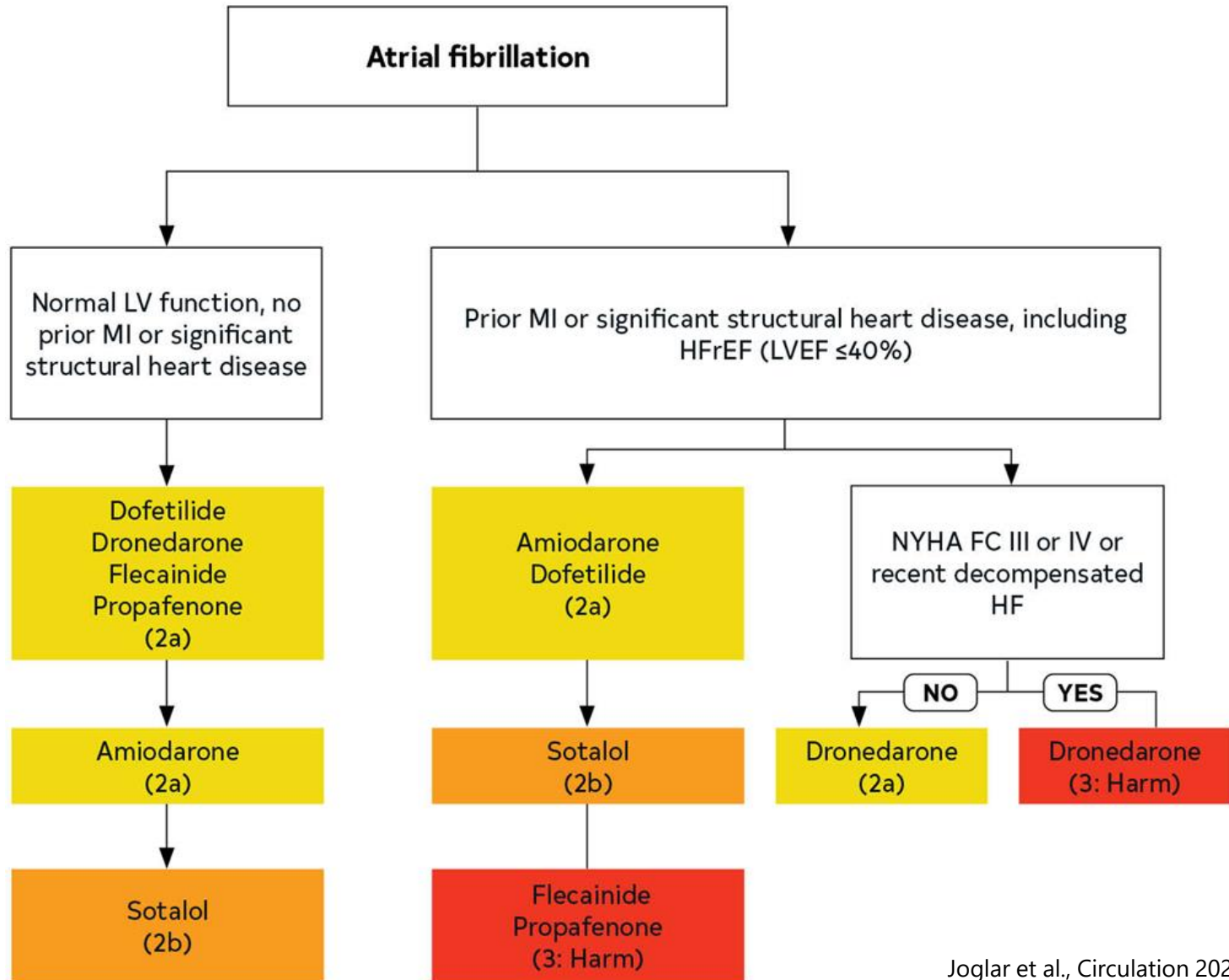
Joglar et al.,  
Circulation 2024;149:e1-e156.



MASSACHUSETTS  
GENERAL HOSPITAL

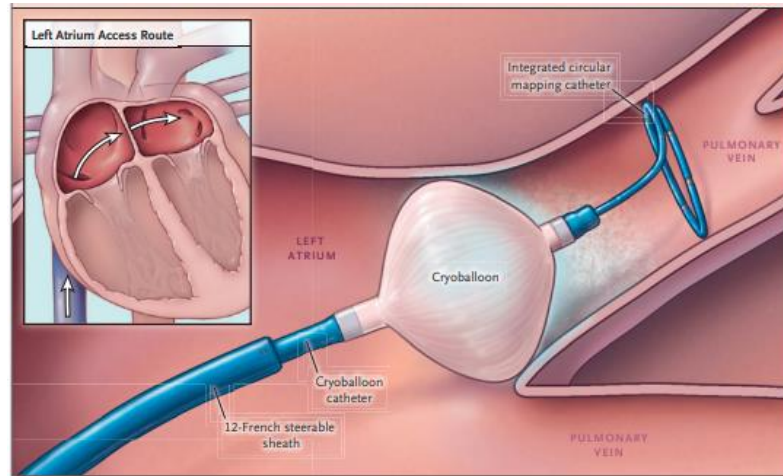
CORRIGAN MINEHAN  
HEART CENTER

# Rhythm Control with Drug Therapy

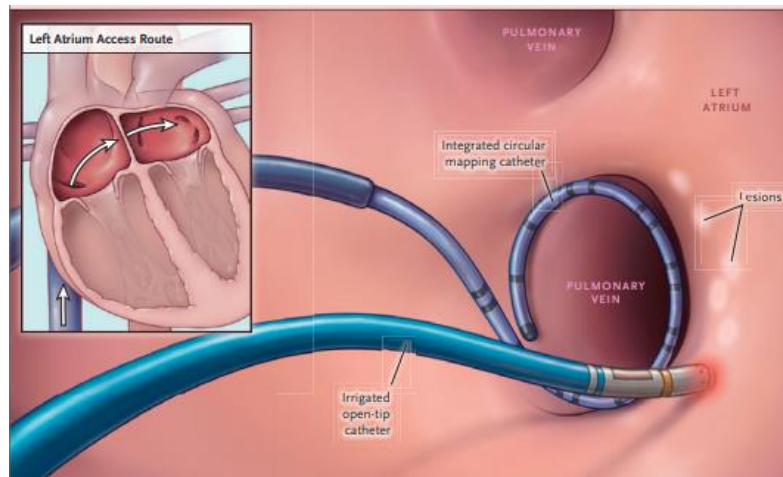


# Rhythm Control with Catheter Ablation

Cryoablation



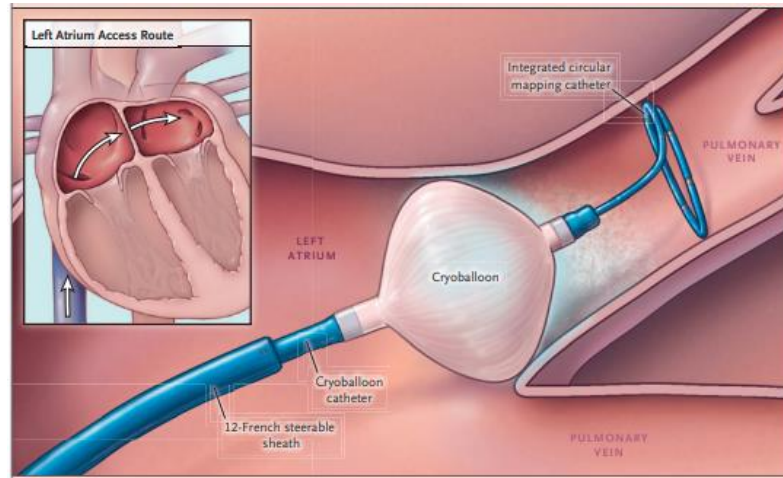
Radiofrequency Ablation



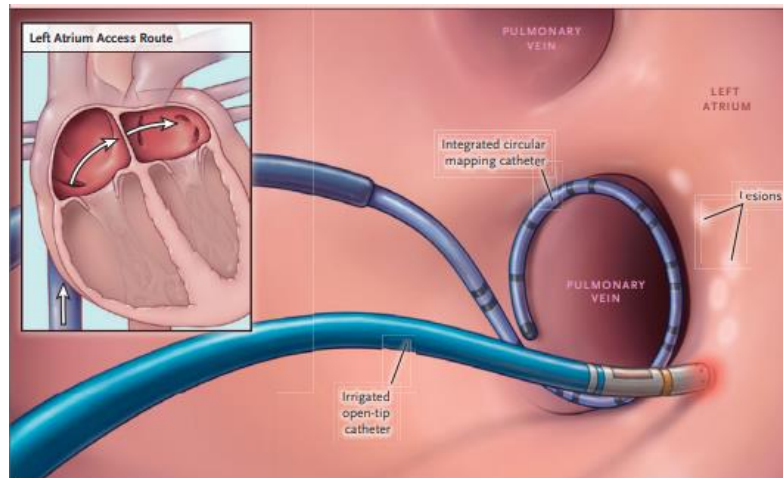
Kuck et al., NEJM 2016;374:2235-2245

# Rhythm Control with Catheter Ablation

Cryoablation



Radiofrequency Ablation

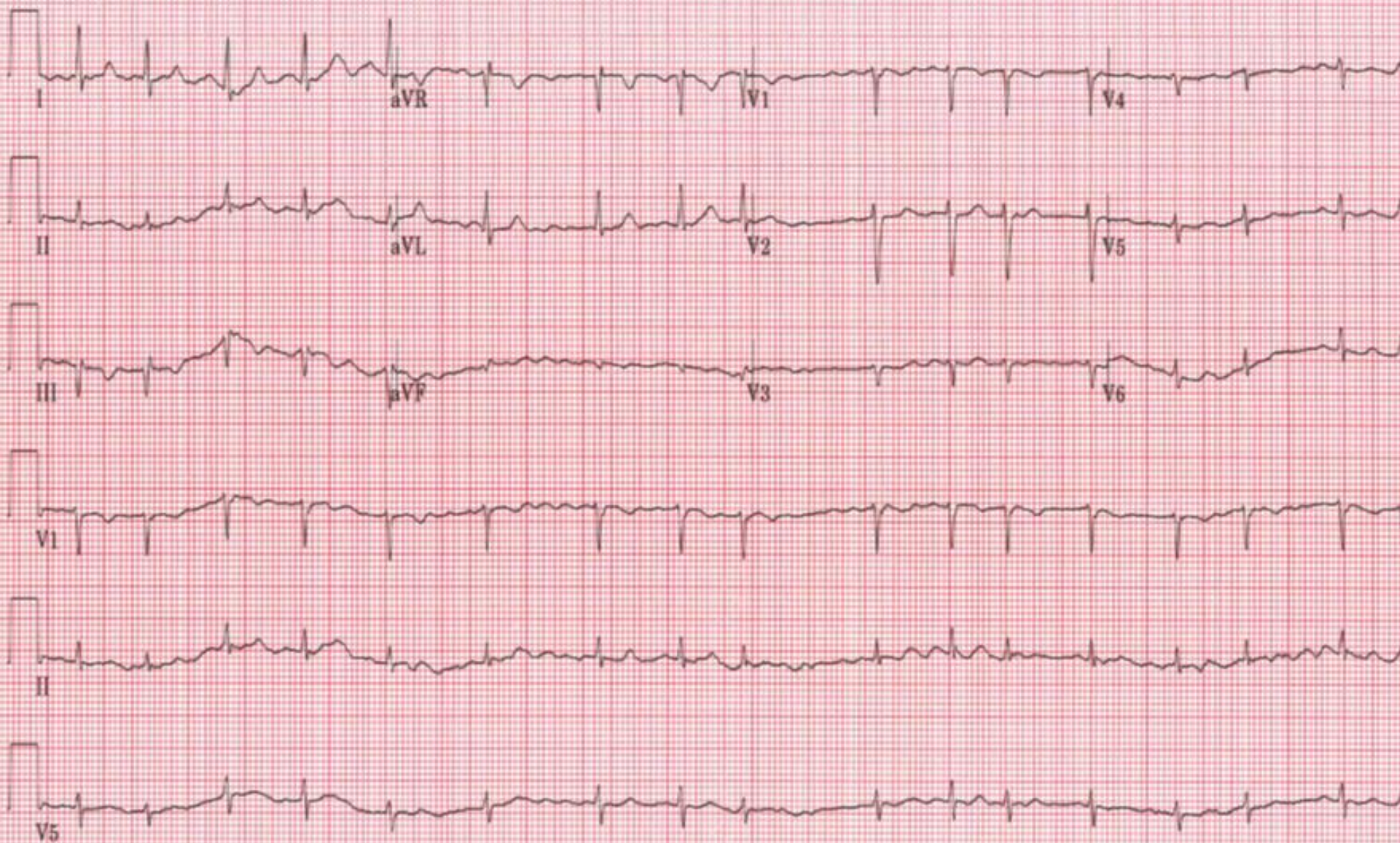


Ablation creates thermal injury which prevents electrical conduction

Kuck et al., NEJM 2016;374:2235-2245

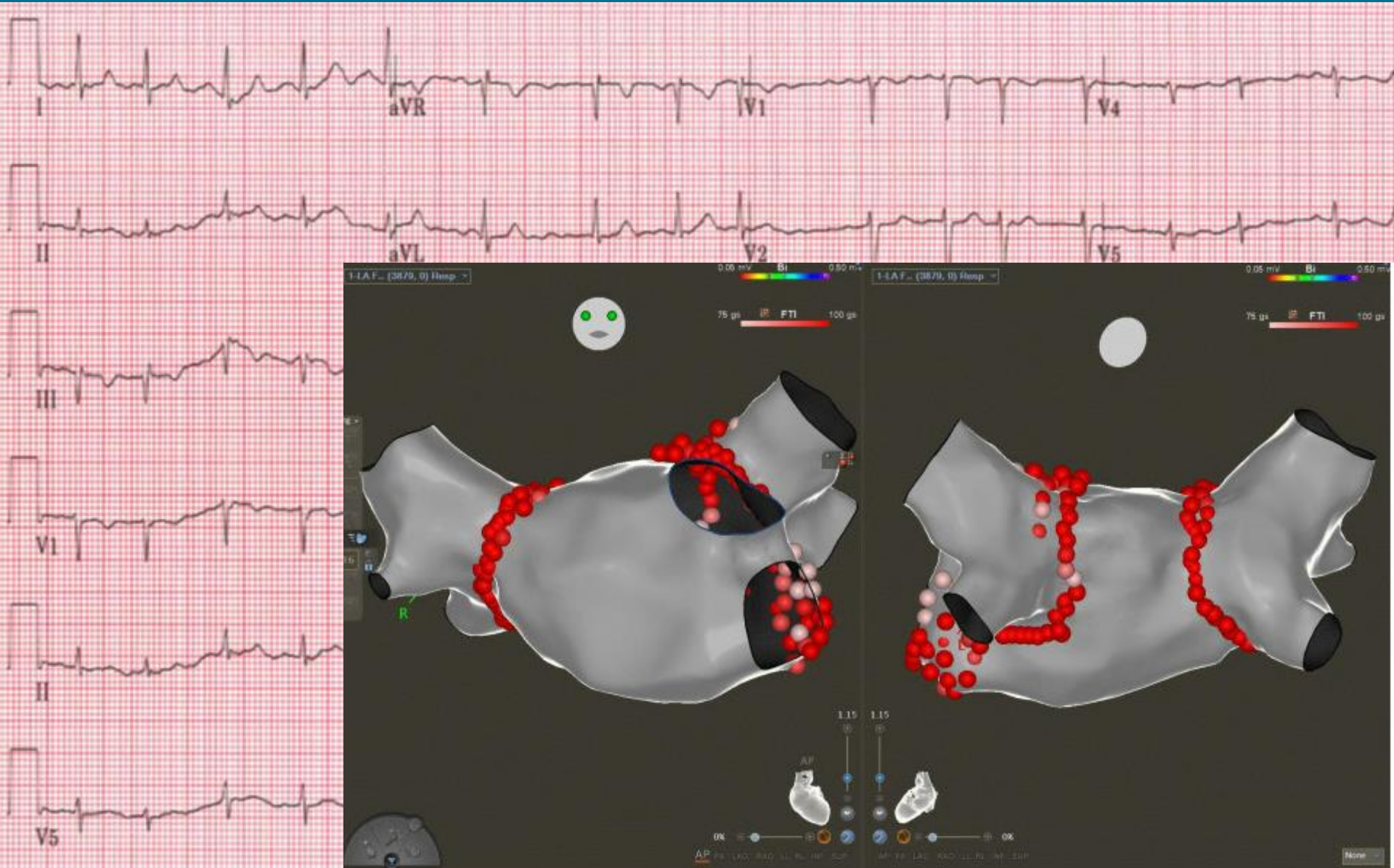


# Atrial Fibrillation



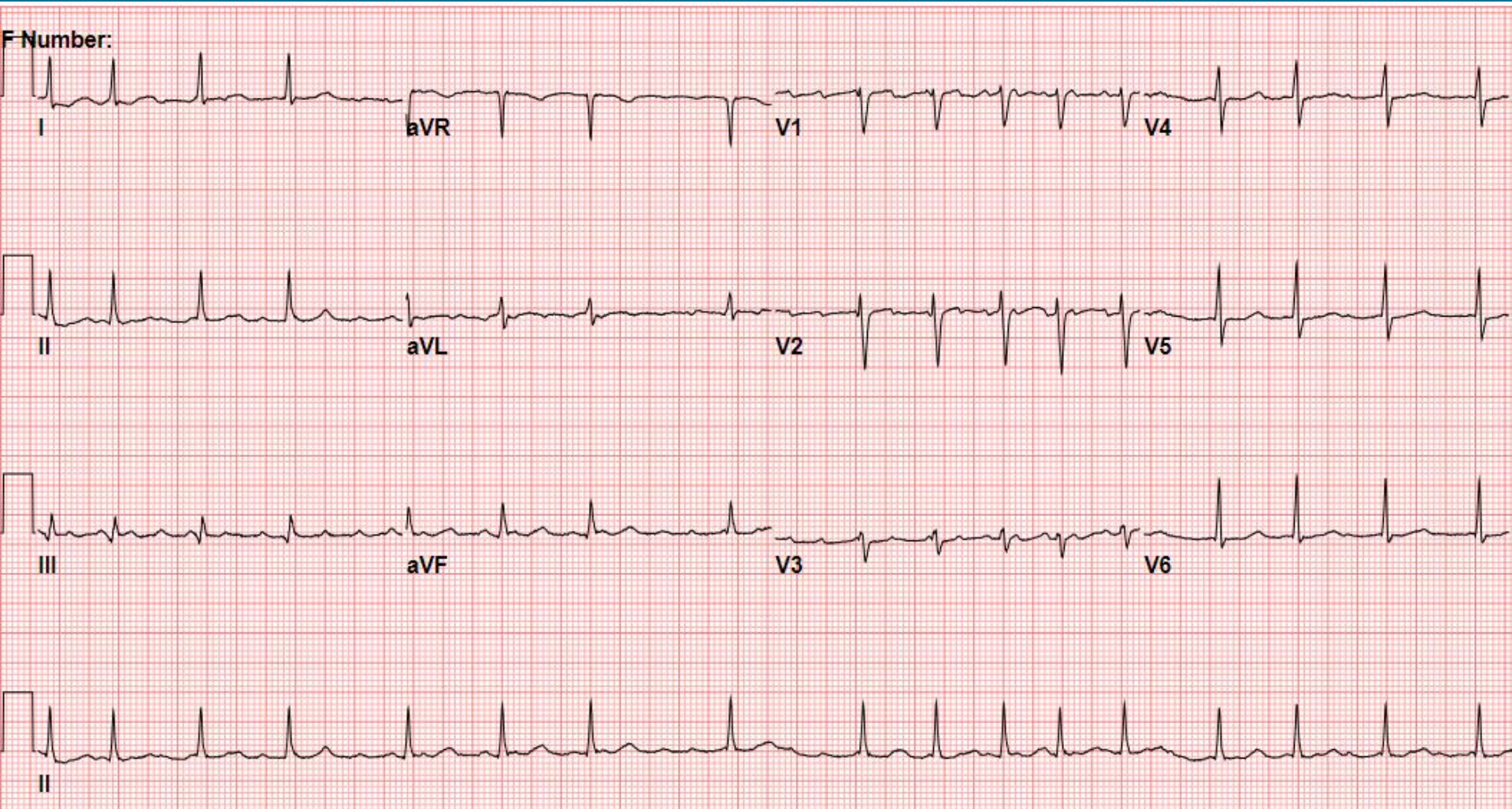


# Atrial Fibrillation





# Atypical Atrial Flutter

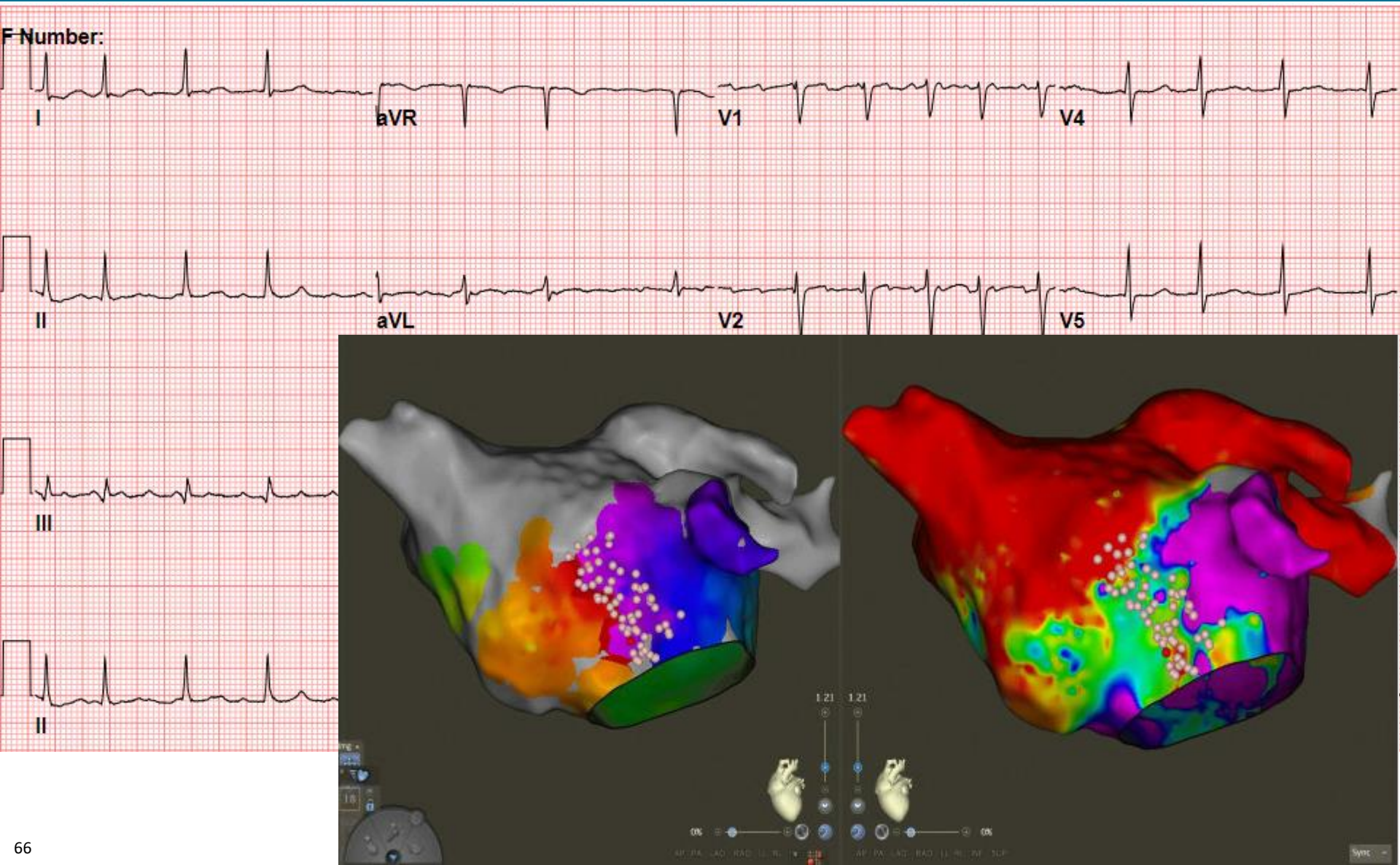


MASSACHUSETTS  
GENERAL HOSPITAL

CORRIGAN MINEHAN  
HEART CENTER

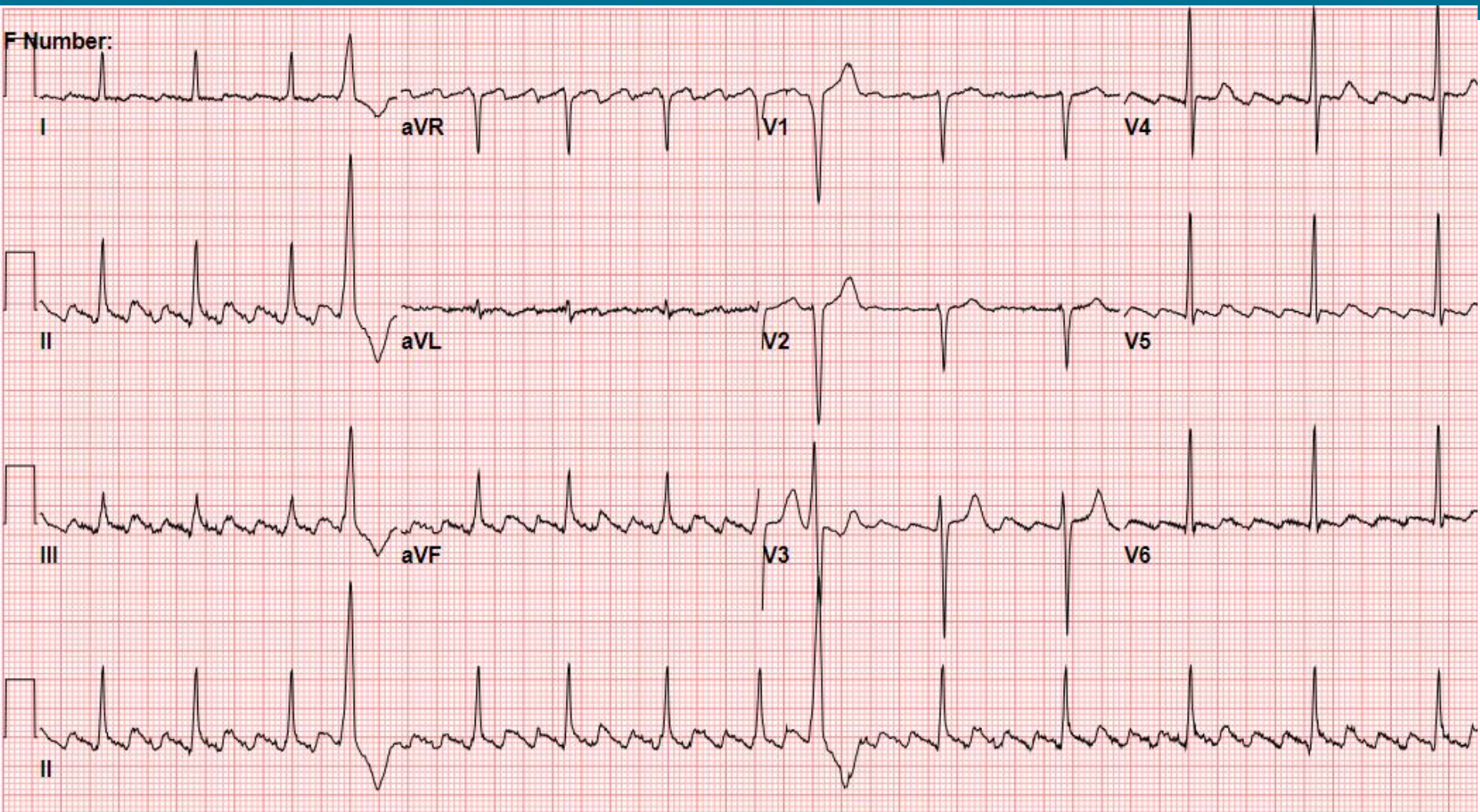


# Atypical Atrial Flutter





# Typical Atrial Flutter

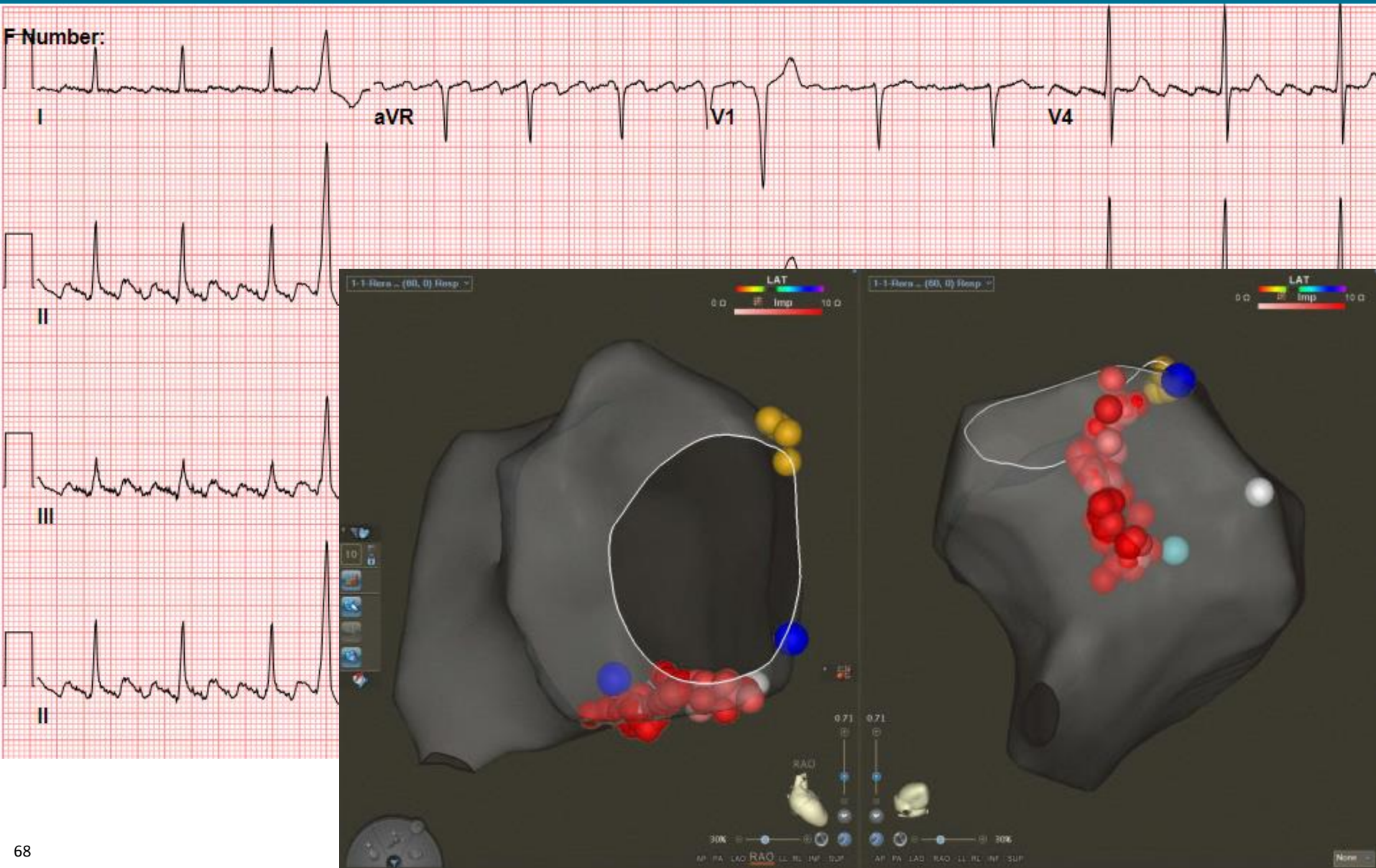


MASSACHUSETTS  
GENERAL HOSPITAL

CORRIGAN MINEHAN  
HEART CENTER



# Typical Atrial Flutter



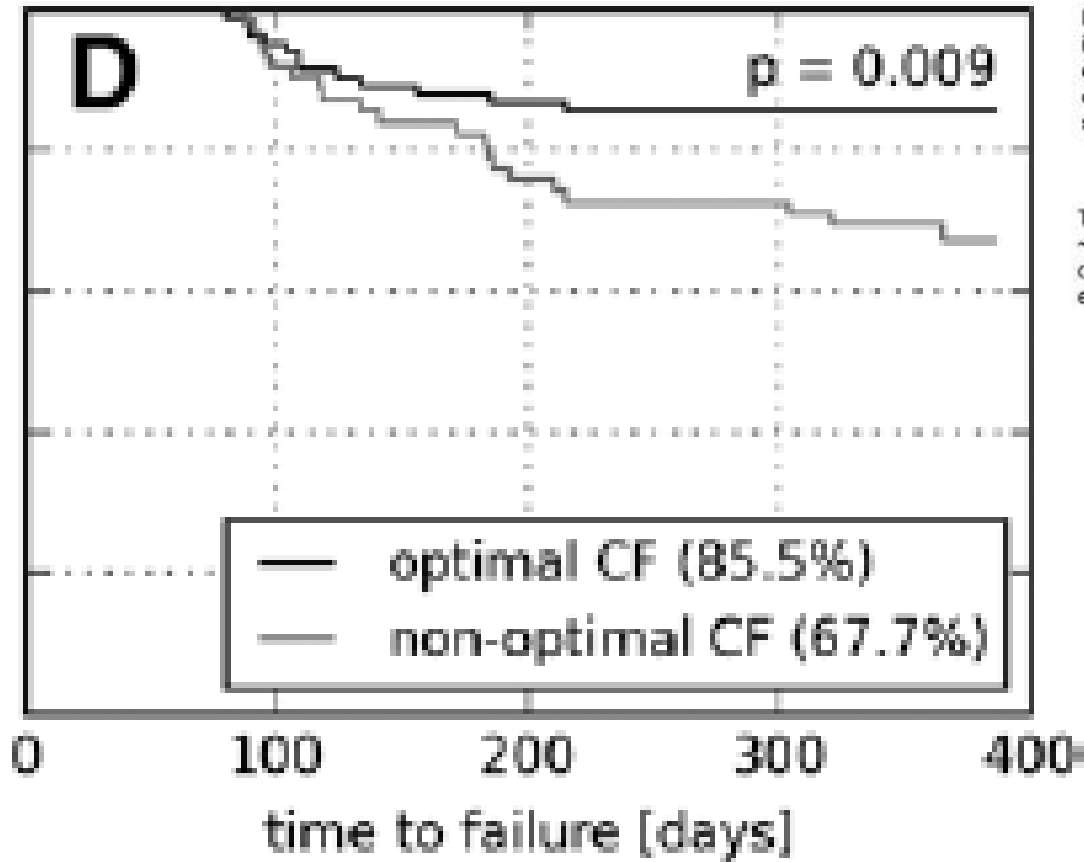
# Outcomes for Ablation with RF Circa 2009

2-year AF-free survival after RFA for Paroxysmal AF: **70%**

2-year AF-free survival after RFA for Persistent AF: **60%**

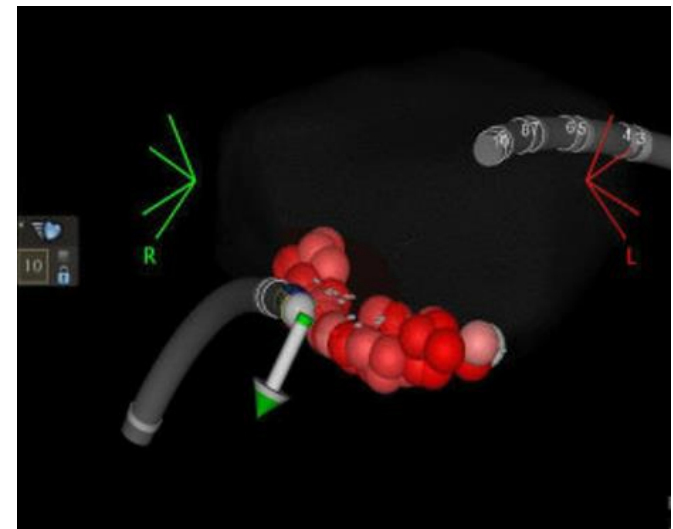
2-year AFL-free survival after CTI RFA for typical AFL: **90%**

# Improvements in Ablation Catheter and Mapping System Technology are Associated with Improved Ablation Outcomes



Flexible, irrigated tip electrode with contact force sensor

Thermocouple ~ 0.4 mm from distal end of tip electrode



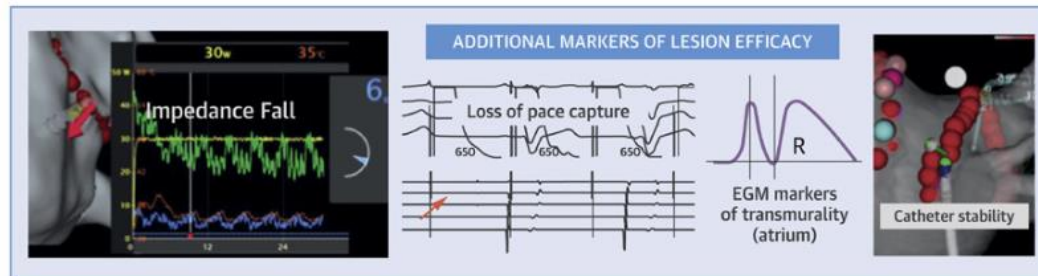
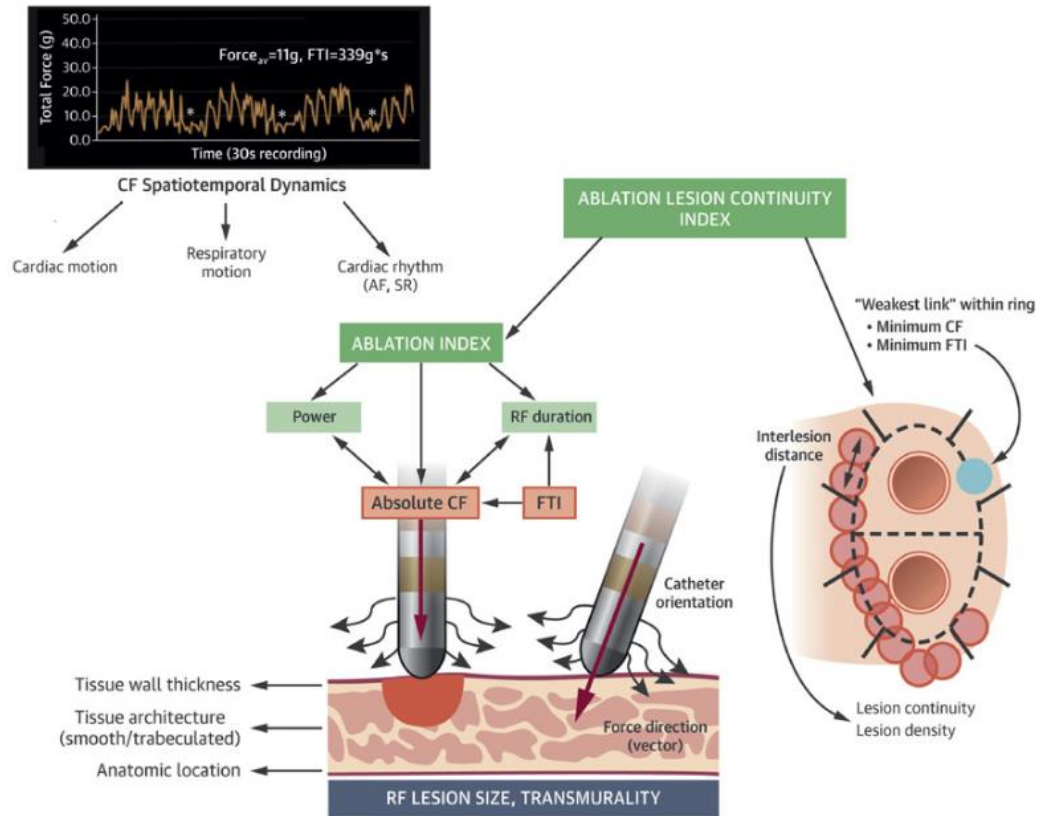
VY Reddy et al., Circulation 2015;132:905-915  
L Ptaszek et al., Heart Rhythm O2 2023;4:42-50



MASSACHUSETTS  
GENERAL HOSPITAL

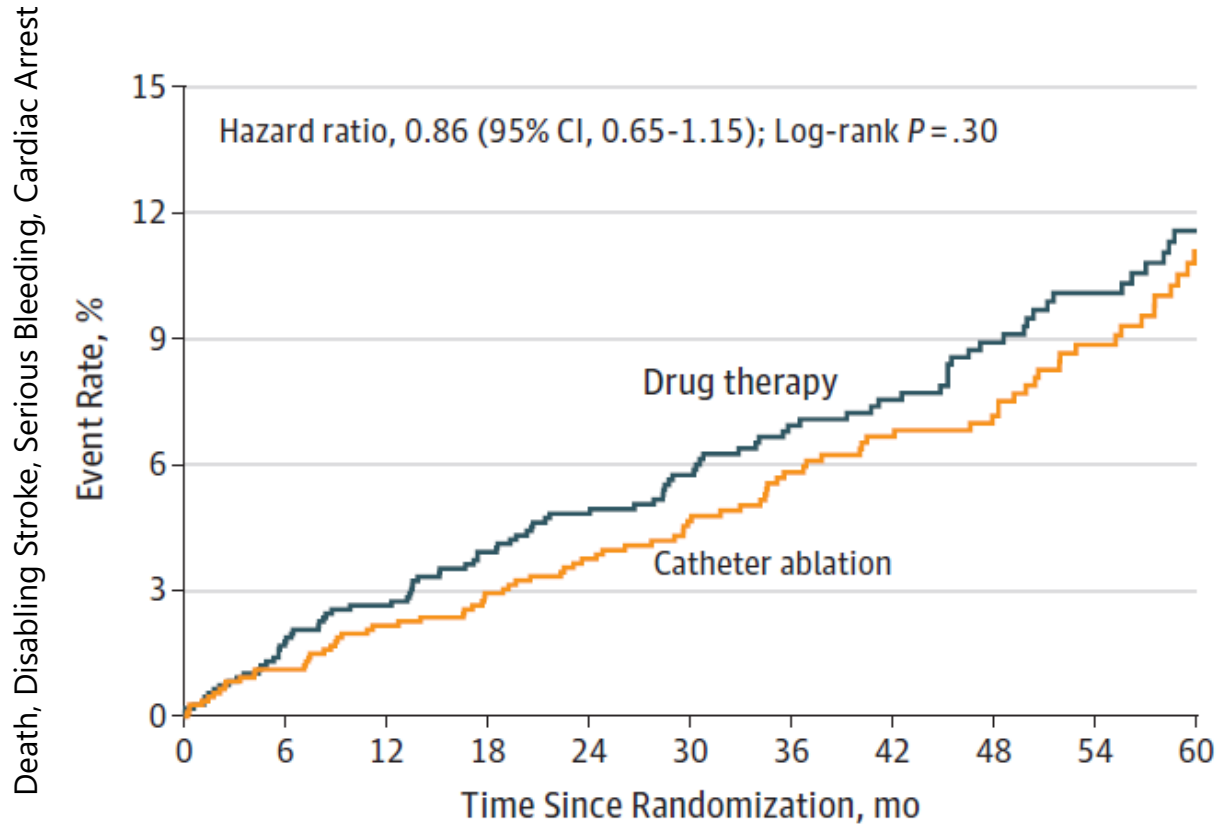
CORRIGAN MINEHAN  
HEART CENTER

# Impact of Contact Force Sensor-Equipped Catheters on Ablation Outcomes



Ariyaratna, N. et al. J Am Coll Cardiol EP. 2018;4(6):707-23.

# Outcomes for RF Ablation vs Medications: CABANA Trial

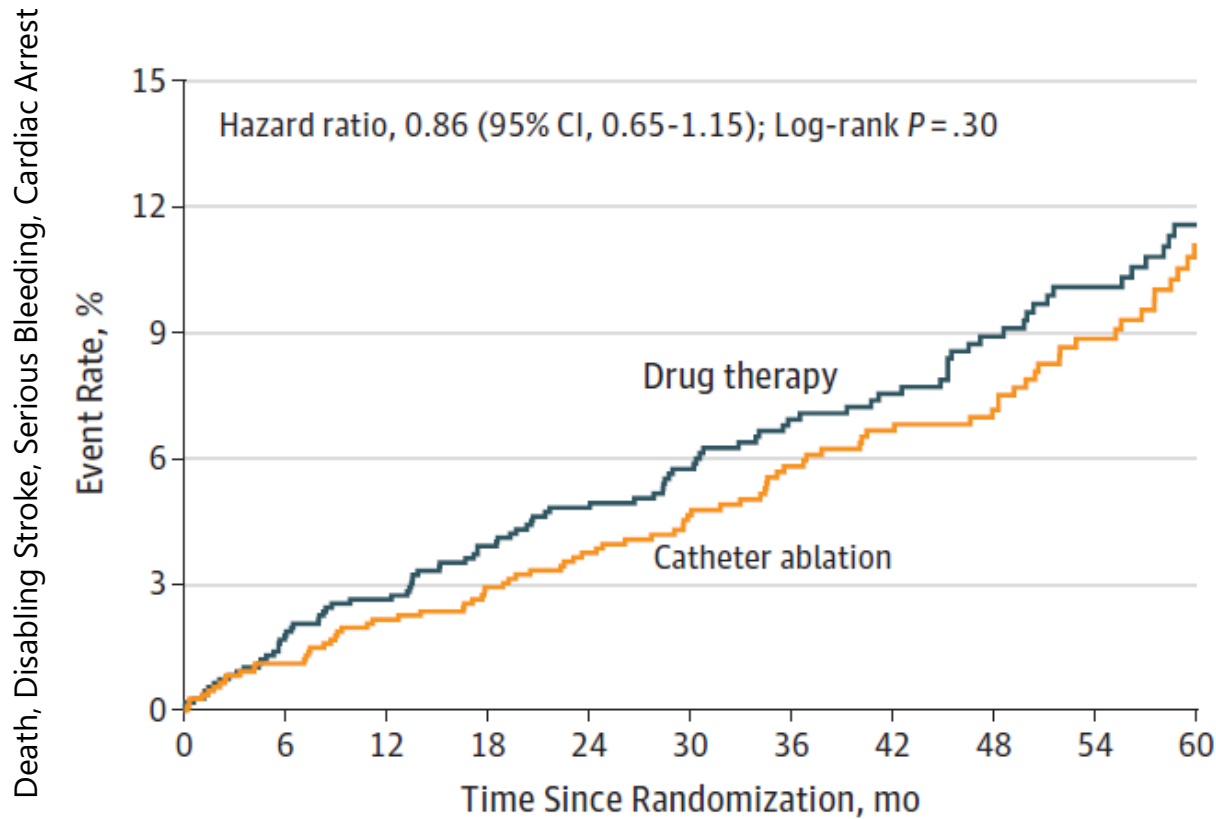


No. at risk	0	6	12	18	24	30	36	42	48	54	60
Drug therapy	1096	1036	1006	970	880	763	652	578	499	418	312
Catheter ablation	1108	1045	1021	996	915	793	700	614	535	432	309

DL Packer et al., JAMA 2019;321:1261-1274



# Outcomes for RF Ablation vs Medications: CABANA Trial



Ablation produced sig. lower burden of AF than AADs.

Ablation produced sig. lower hospitalization rate than AADs.

**Possible underestimation of treatment effect:**

Lower event rate than anticipated.

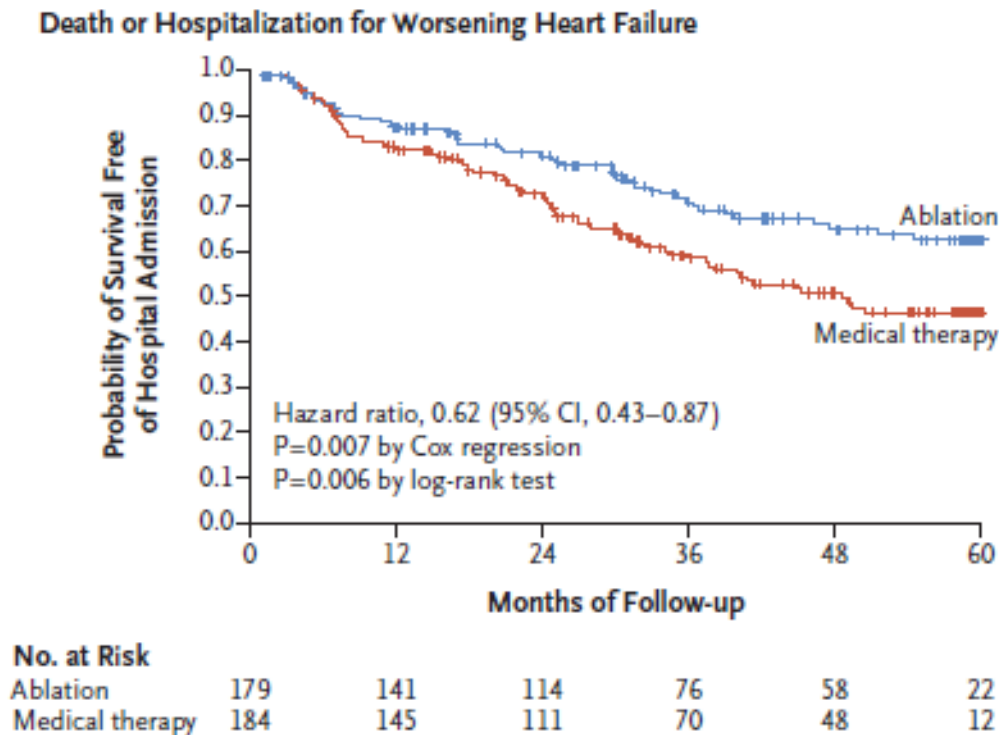
Treatment crossover

No. at risk	0	6	12	18	24	30	36	42	48	54	60
Drug therapy	1096	1036	1006	970	880	763	652	578	499	418	312
Catheter ablation	1108	1045	1021	996	915	793	700	614	535	432	309

DL Packer et al., JAMA 2019;321:1261-1274

# CASTLE-AF Trial

- Randomized trial comparing ablation vs. AADs in patients with CHF (LVEF 35%).
- Composite outcome: all-cause death, HF hospitalization.



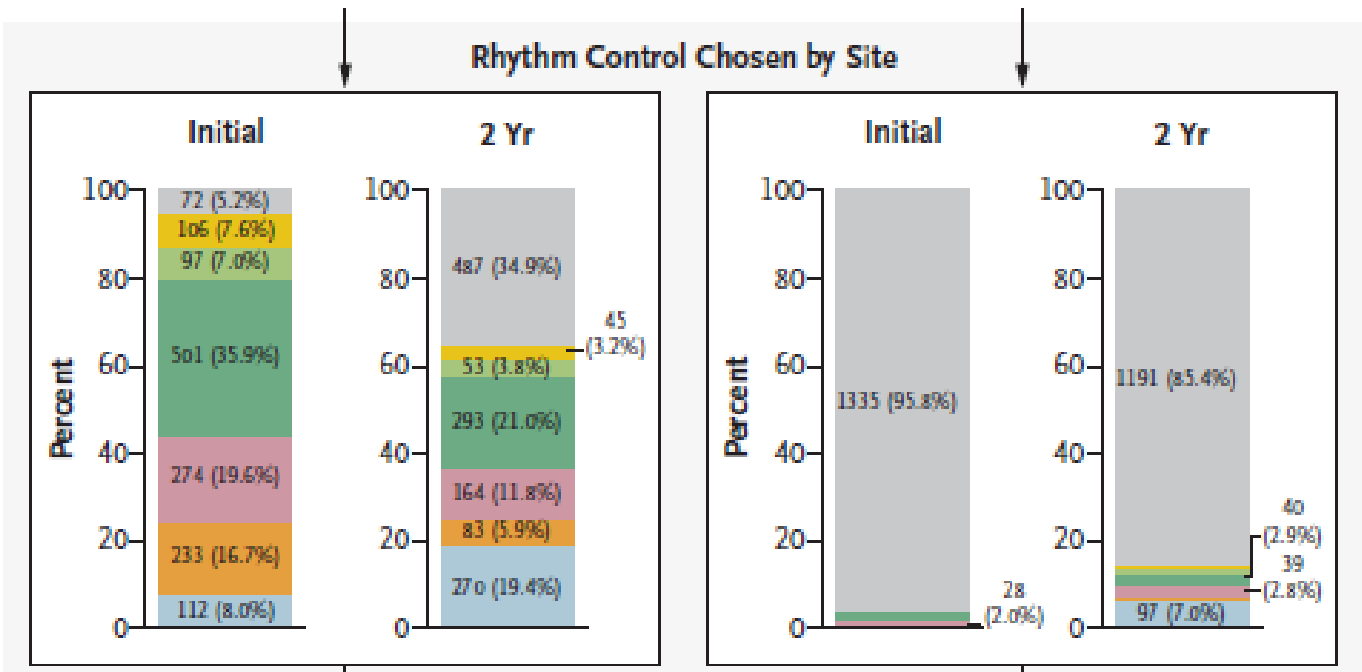
Marrouche et al., New Engl J Med 2018;378:417-427.

# Early Rhythm Control Therapy for AF: EAST-AFNET 4 Trial

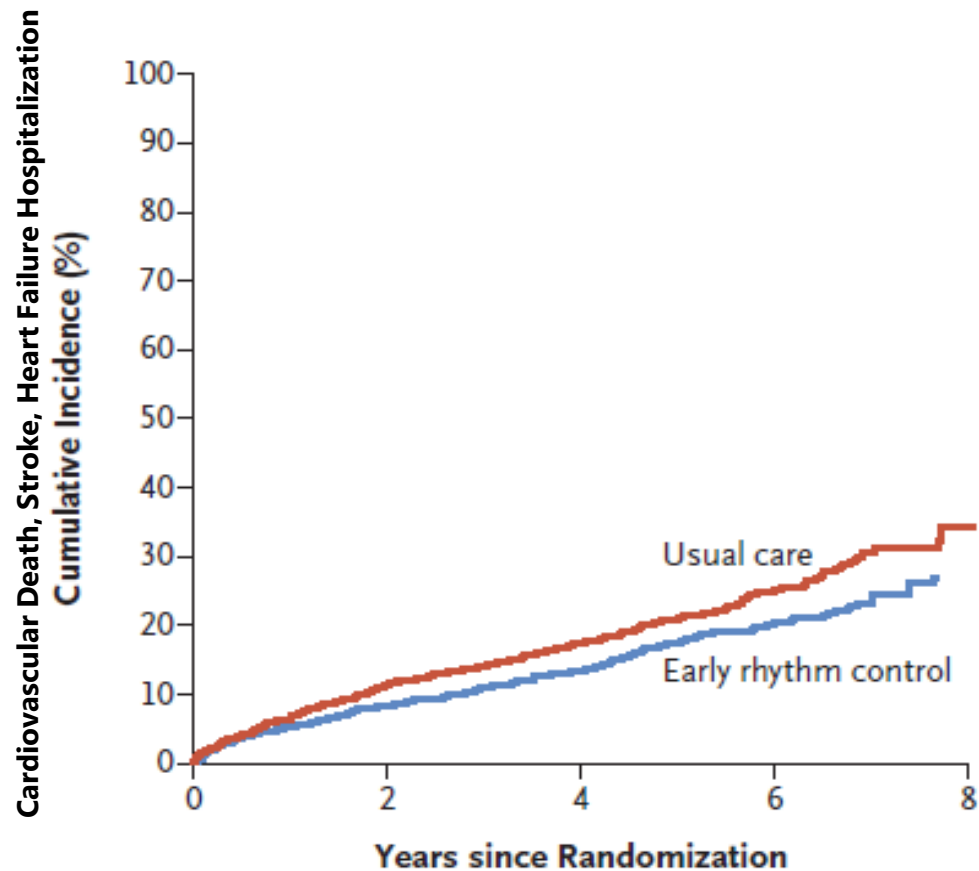
Patients with early AF (<1 yr)

Early rhythm control

Usual care



# Early Rhythm Control Therapy for AF: EAST-AFNET 4 Trial



## No. at Risk

Usual care	1394	1169	888	405	34
Early rhythm control	1395	1193	913	404	26

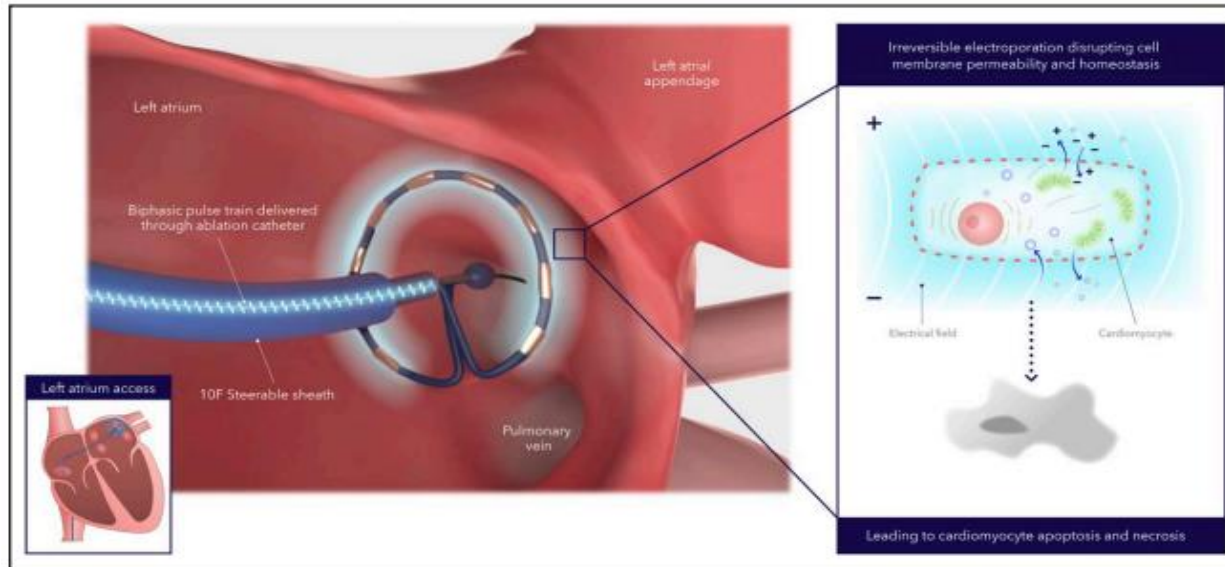
Kirchhof et al., New Engl J Med 2020;383:1305-1316.



MASSACHUSETTS  
GENERAL HOSPITAL

CORRIGAN MINEHAN  
HEART CENTER

# Pulsed Field Ablation Technique

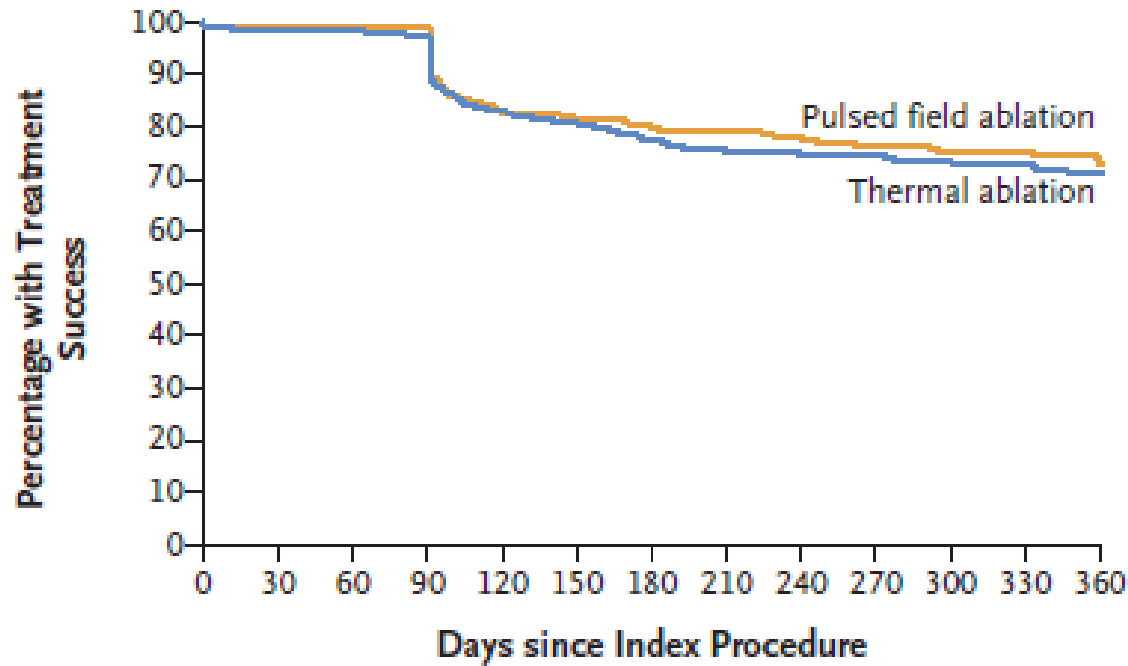


Verma et al., Circulation 2023;47:1422-1432.



Reddy VY et al., J Am Coll Cardiol 2019;74:315-326

# PFA Non-Inferior to RFA: ADVENT Trial



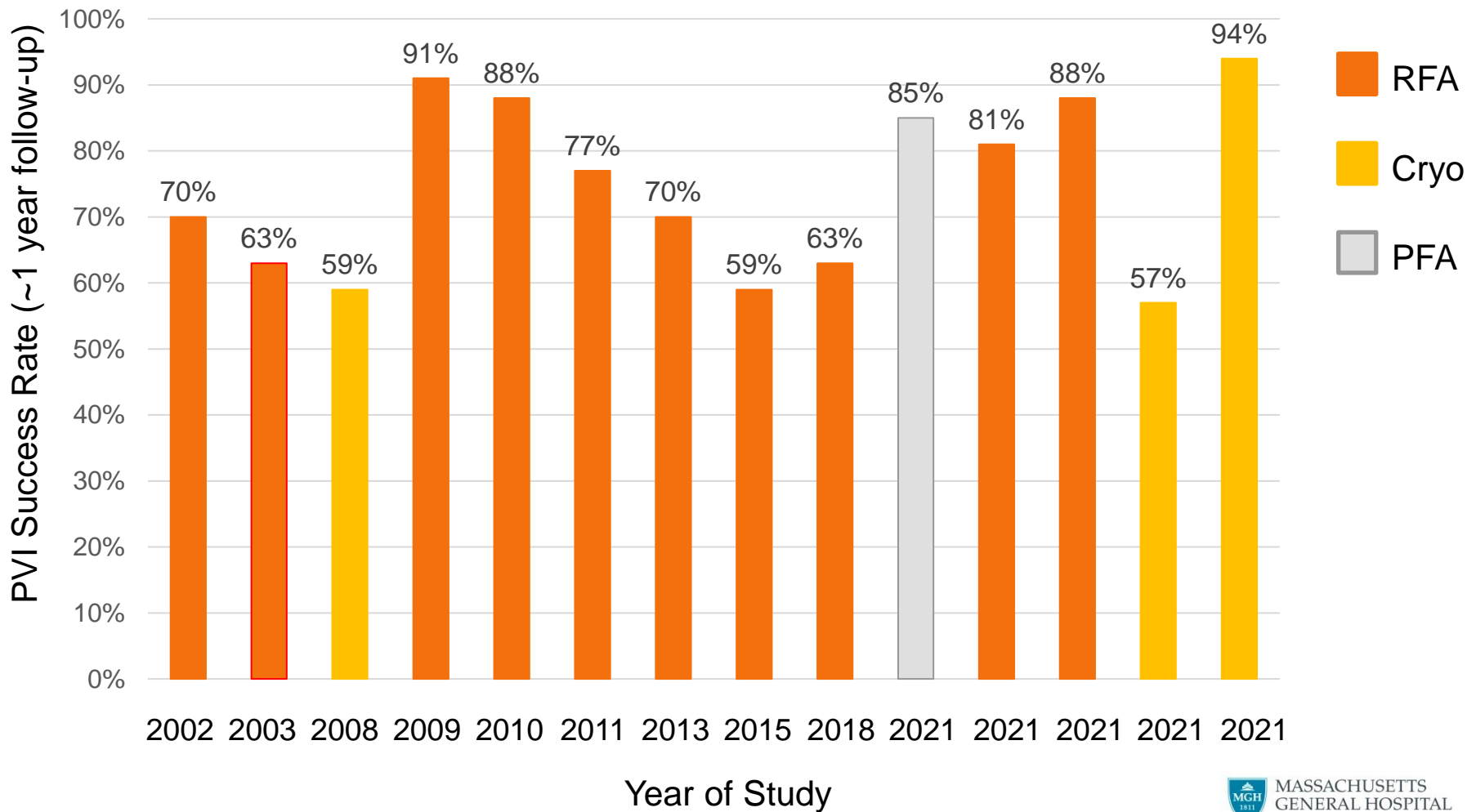
## No. at Risk

Pulsed field ablation	301	298	238	228	176
Thermal ablation	296	292	228	219	150

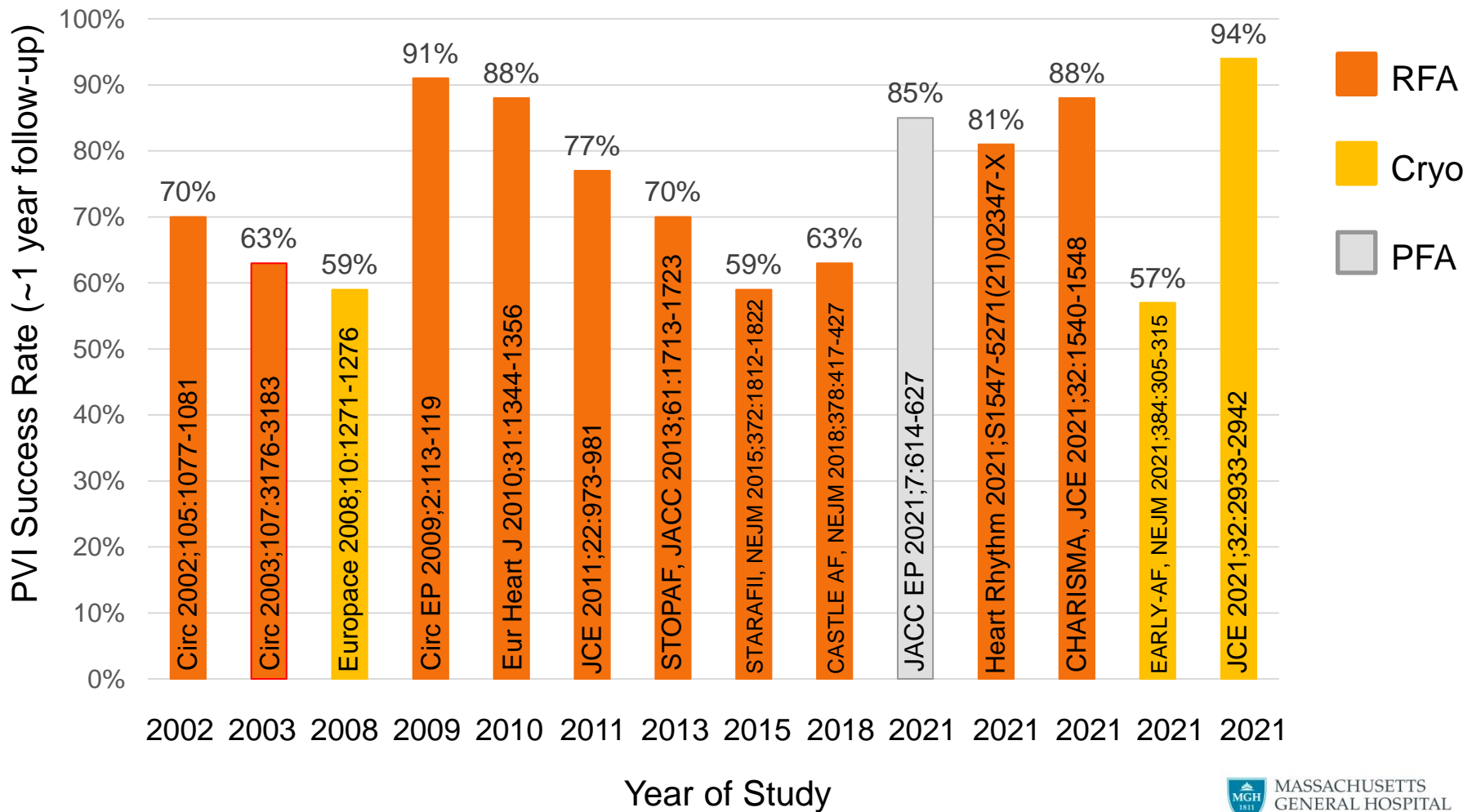
## Treatment Success (%)

Pulsed field ablation	99.3	99.0	79.7	76.4	73.1
Thermal ablation	98.7	97.3	77.5	74.5	71.3

# Variability in PVI Success Rate Persists Despite Advances in Mapping and Ablation Technology



# Variability in PVI Success Rate Persists Despite Advances in Mapping and Ablation Technology

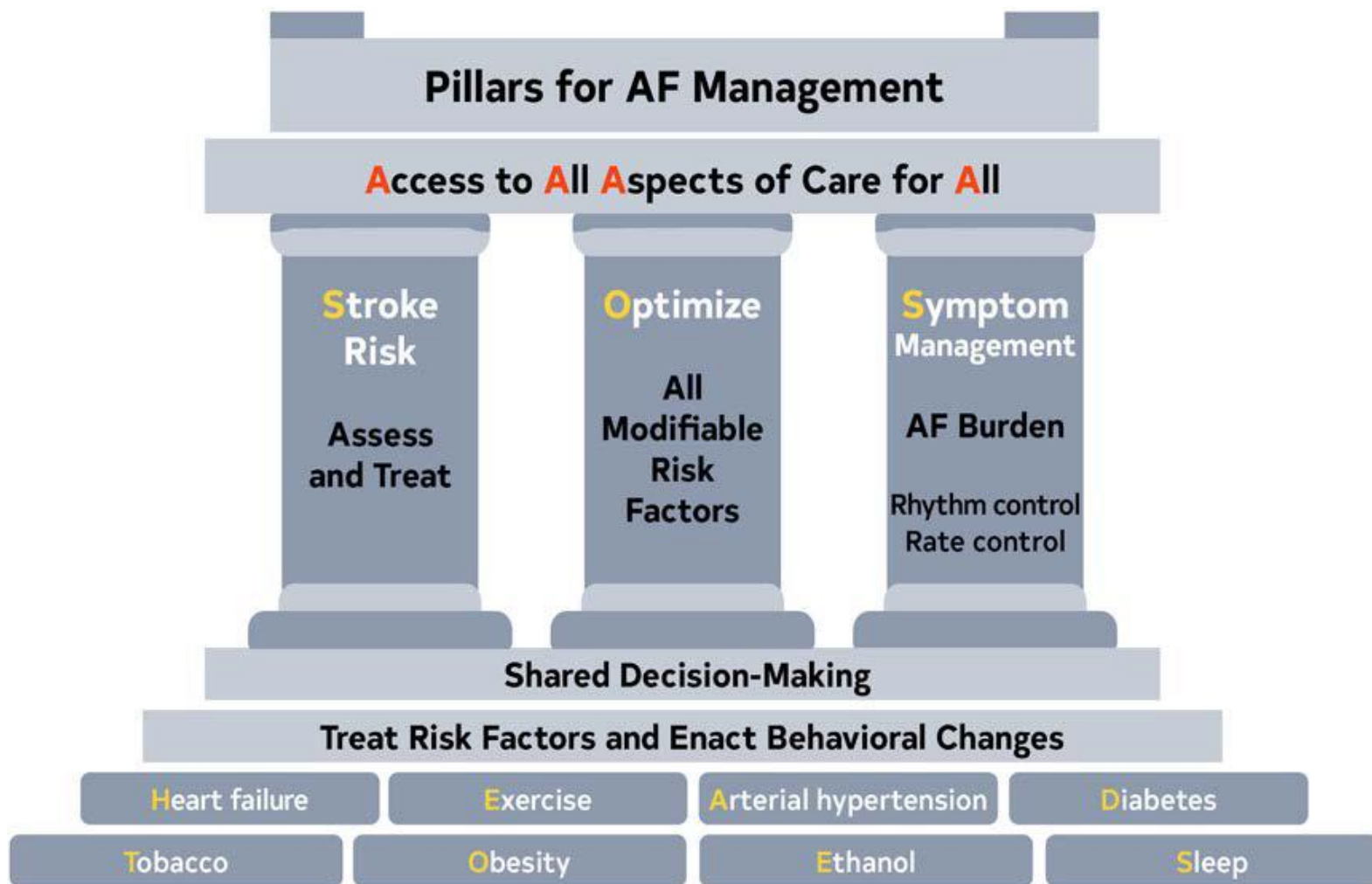




# Outline

- 1) **Clinical Case**
- 2) **Epidemiology of AF/AFL**
- 3) **Physiology of AF/AFL**
- 4) **Advances in medical and interventional treatment:**
  - *Reduce the risk of AF/AFL-related stroke*
  - *Minimize consequences of AF/AFL on heart function (symptoms, CHF)*
- 5) **Current recommendations for management of AF/AFL**

# Summary



MASSACHUSETTS  
GENERAL HOSPITAL

CORRIGAN MINEHAN  
HEART CENTER

# Summary

- 1) Utilize a stroke risk score (e.g., CHA2DS2-VASC) to assess indication for initiation of OAC.
- 2) Utilize a bleeding score (e.g., HAS-BLED) to assess risk of bleeding with OAC. For patients who not good candidates for OAC consider LAA closure device implantation.
- 3) Consider catheter ablation for patients in whom AADs are not effective, contraindicated, not tolerated, or not preferred.
- 4) Catheter ablation can be considered as first-line therapy to improve symptoms or reduce progression of paroxysmal to persistent AF.

# Summary

- 5) Early rhythm control is associated with a greater likelihood of maintaining sinus rhythm in the long term and minimizing AF burden and reducing progression of the disease.
- 6) Lifestyle and risk factor modification is a pillar of AF management.
  - Weight loss for those with body mass  $>27$  kg/m<sup>2</sup>
  - Tobacco cessation
  - Minimization or elimination of alcohol consumption
  - Screening for sleep-disordered breathing
  - Moderate-to-vigorous exercise 210 min/wk

Thank you



MASSACHUSETTS  
GENERAL HOSPITAL