

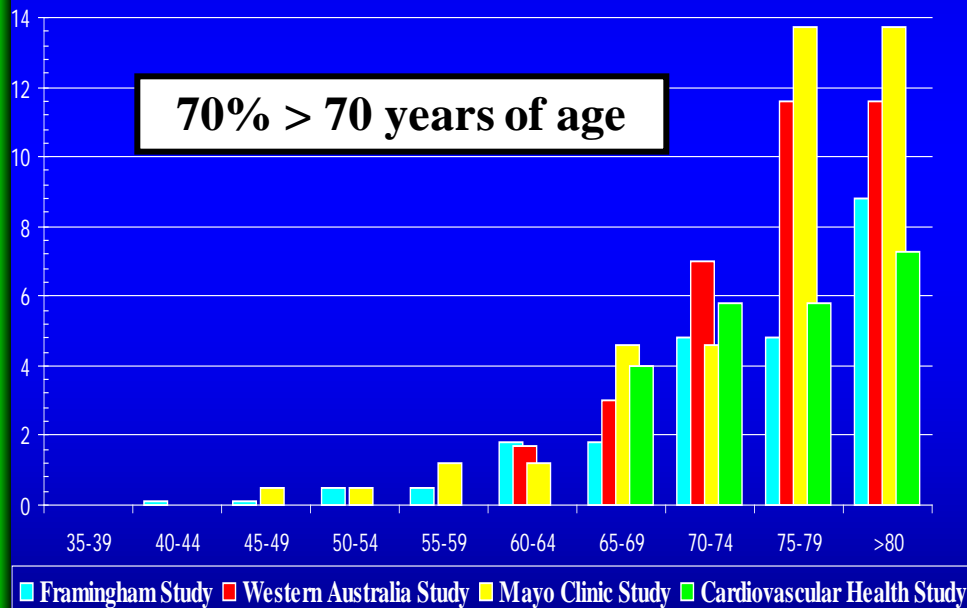
# *Atrial fibrillation and advanced age*

**Prof. Fiorenzo Gaita**

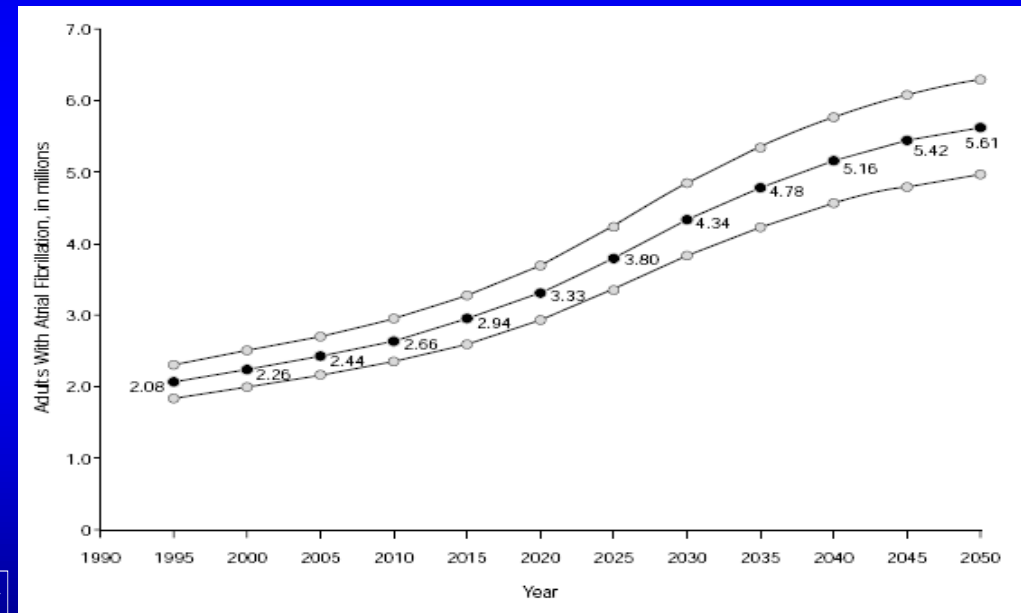
**Director of the Cardiology School  
University of Turin, Italy**

# Prevalence of AF in the general population

## Prevalence and age distribution in patients with atrial fibrillation

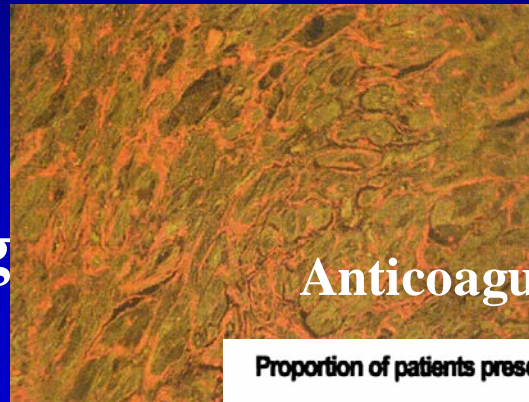
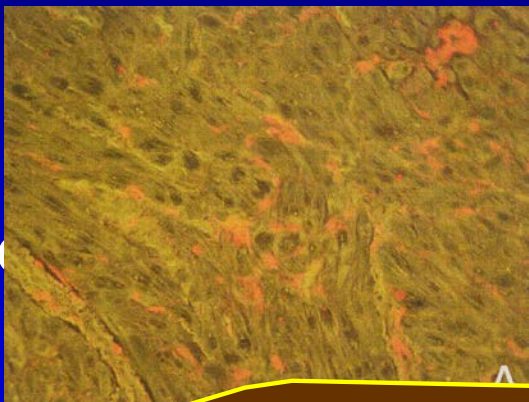


## Projected number of adult with AF in the USA between 1995-2050



# Elderly AF patients are different from Younger

- Atrial substrate modifications related to senescence



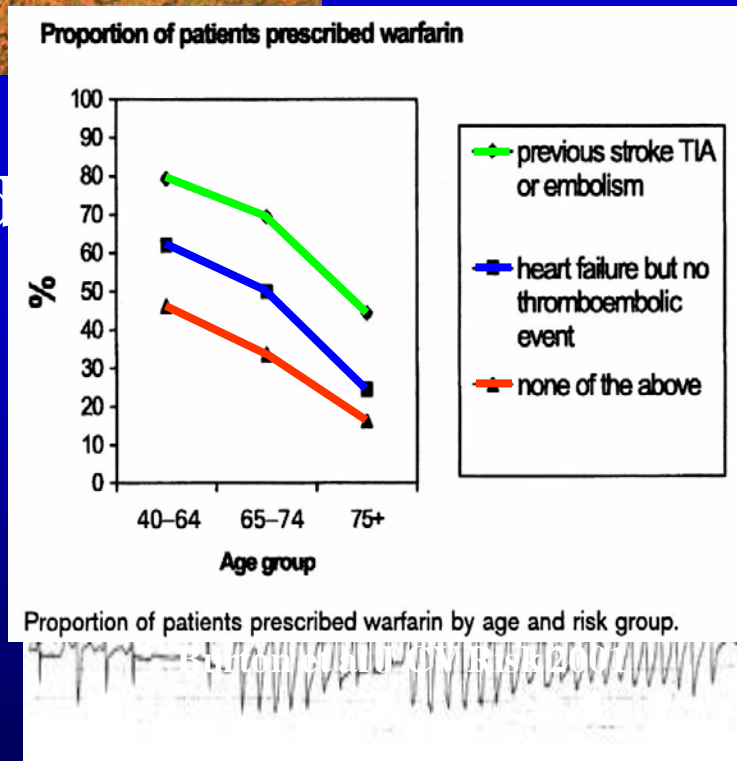
- Problem with imaging

Anticoagulation Paradox

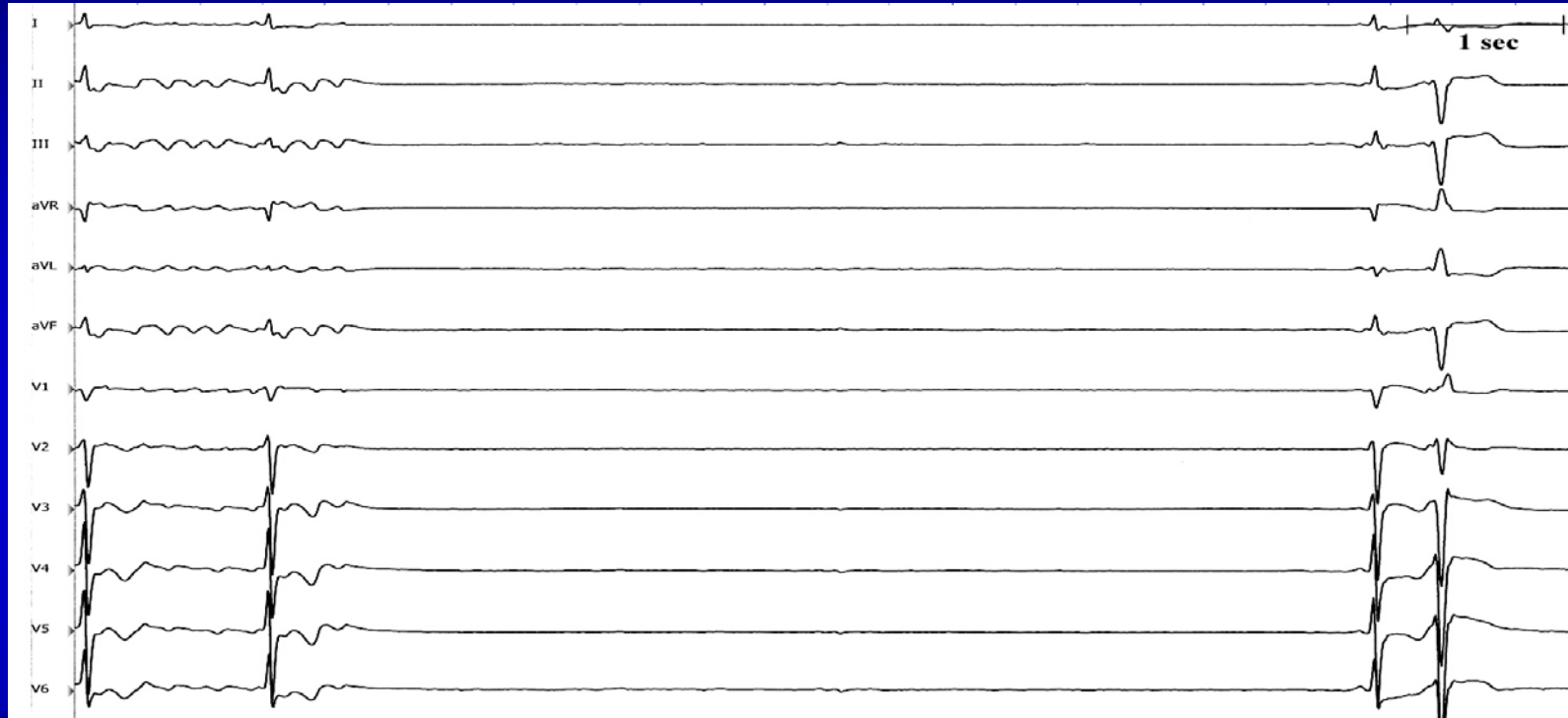
- Antiarrhythmic therapy is more difficult

hemorrhagic complications

Altered liver and renal function,  
Prevention  
Ischemic stroke  
electrolyte abnormalities  
pharmacological interactions,  
poor compliance → pro-arrhythmias



- More comorbidities, associated diseases particularly SSS

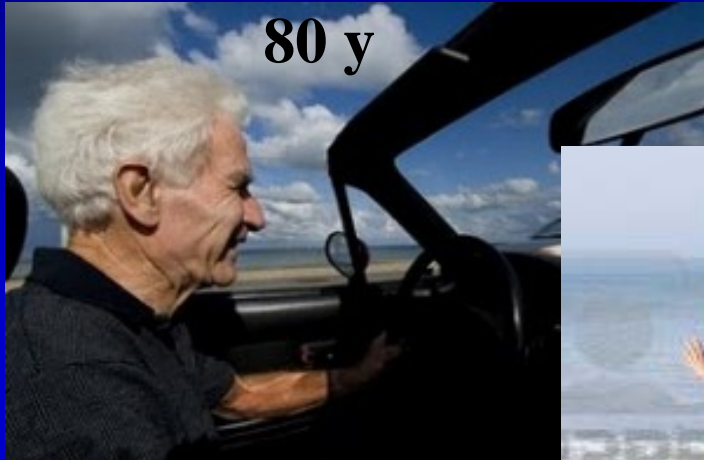


**Pathologic recovery time of sinus node  
(increased by ADDs therapy)**



# Elderly AF patients are different from Younger

*...but, above all, they are different from each others*





Created by Jacques Magka

18th WORLD CONGRESS  
in Cardiac Electrophysiology  
& Cardiac Techniques

JUNE 13 > 16 | 2012  
NICE • FRANCE



Final program



www.cardiostim.com

Reed Expositions

In collaboration with



11:00-12:30

Room 1.4 - Level 1 / Méditerranée

SESSION 108

APPROACHES TO ATRIAL FIBRILLATION AND  
STROKE PREVENTION: NEW TECHNIQUES,  
NEW TECHNOLOGY, NEW ANTICOAGULANTS  
AND NEW DRUGS FOR THE ELDERLY

Co-Chair:

Gaita F. - Asti, ITA / Munger T.T. - Rochester, USA

11:00 ■ State of the Art Lecture:

AF in the Elderly: Drugs, Ablation and Devices

Calkins H. - Baltimore, USA

108/1

11:20 ■ Case 1: 86 yr W, HTN, Diabetic w/ 2 recurrent

TIAs in setting of PAF, GI bleeds/fall

Friedman P. - Rochester, USA

108/2

11:40 ■ Case 2: 90 yr M, persistent AF, HTN, CHF, and

COPD

Shen W. - Rochester, USA

108/3

12:00 ■ Case 3: 85 yr retired surgeon runs marathon,

sinus bradycardia and symptomatic PAF

Hocini M. - Bordeaux, FRA

108/4

## Clinical Case

**Retired Surgeon, 85 yr,  
marathon runner**

- Symptomatic Paroxysmal AF with rapid ventricular rate
- Sinus bradycardia
- Hypertension
- Impaired renal function

*What could you do?*

a. AADs + OAC

b. Rate control therapy + OAC

c. AADs + LA appendage closure

d. Rate control therapy + LAA closure

e. PM+AADs + OAC

f. PM +AV node abl+ OAC

g. PM + AVnode abl + LA appendage closure

h. Afib ablation + strictly SR monitorization (Loop Recorder)

i. Afib ablation + LA appendage closure

# CHA<sub>2</sub>DS<sub>2</sub>VASc

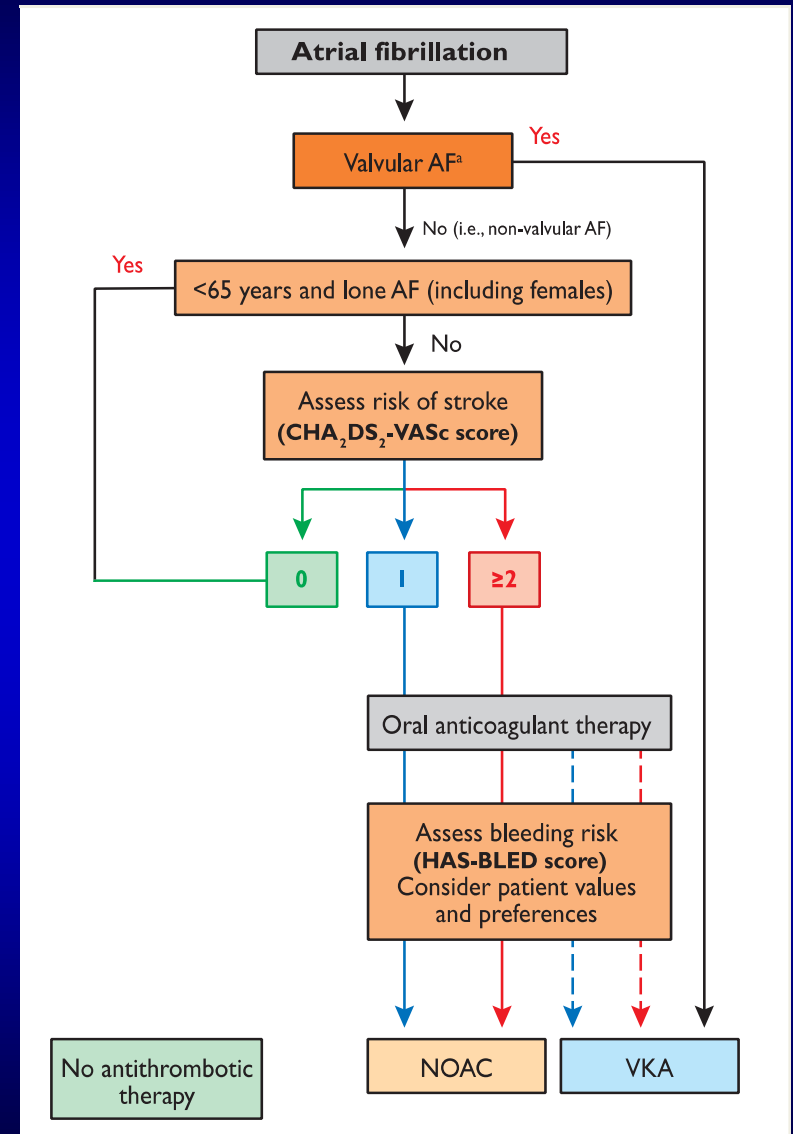
## RISK FACTORS SCORE

<b>Congestive HF/ LV dysfunction</b>	1
<b>Hypertension</b>	1
<b>Age ≥ 75</b>	2* <b>major risk factor</b>
<b>Diabetes mellitus</b>	1
<b>Stroke/TIA/ thromboembolism</b>	2* <b>major risk factor</b>
<b>Vascular disease</b>	1

## CHA<sub>2</sub>DS<sub>2</sub>VASC 3

TE risk ranging from 3.2- 6%/y

# Choice of anticoagulant





# CHA<sub>2</sub>DS<sub>2</sub>VASc

RISK FACTORS	SCORE
Congestive HF/ LV dysfunction	1
Hypertension	1
Age ≥ 75	2*major risk factor
Diabetes mellitus	1
Stroke/TIA/ thromboembolism	2* major risk factor
Vascular disease	1
Age 65-74	1
Sex category (f)	1

# HAS-BLED

RISK FACTORS	SCORE
<b>H</b> ypertension	<b>1</b>
<b>A</b> bnormal renal and liver function (1 point each)	<b>1 or 2</b>
<b>S</b> troke	1
<b>B</b> leeding	1
<b>L</b> abile INRs	1
<b>E</b> lderly (age > 65 y)	<b>1</b>

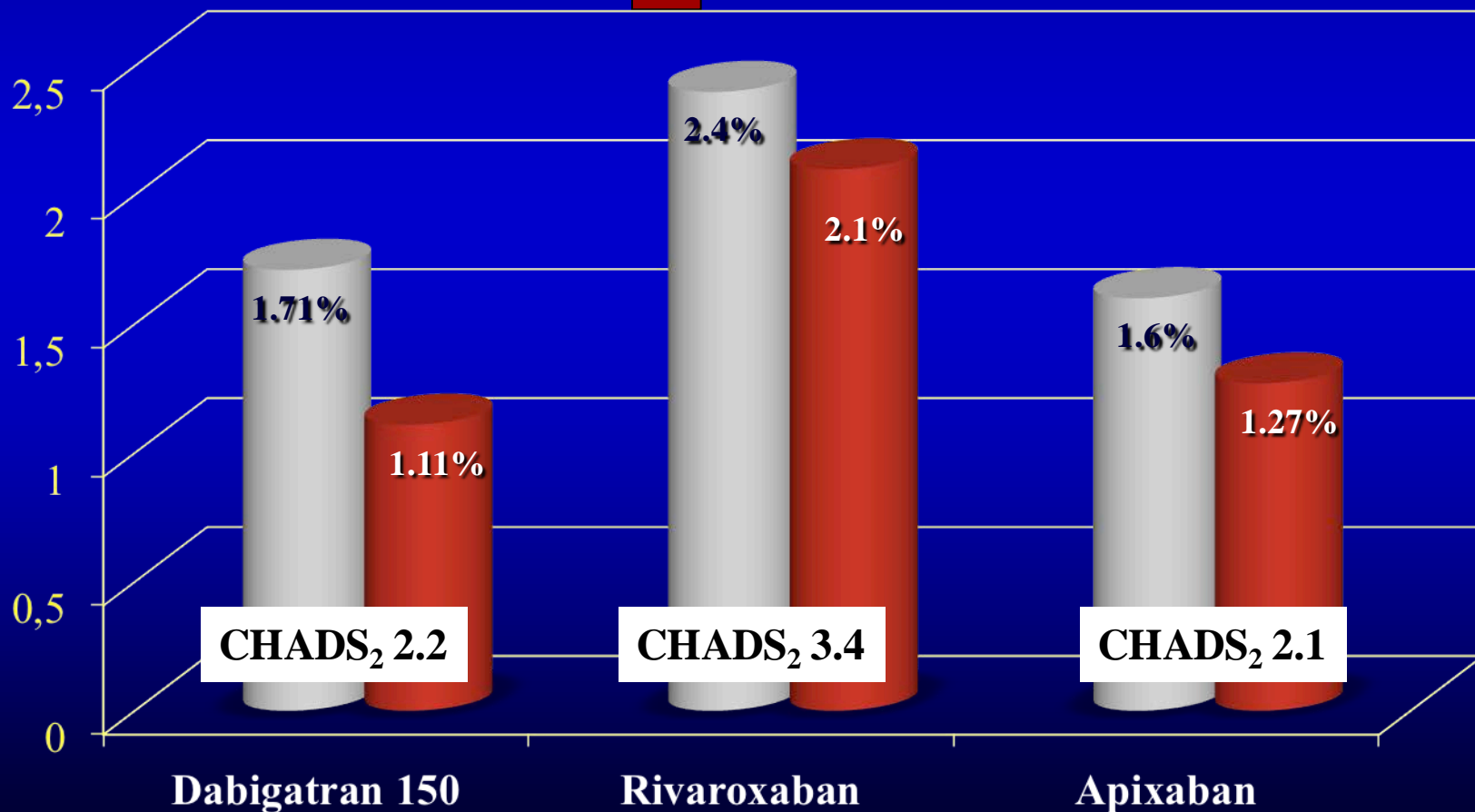
**HASBLED 3**  
**Hemorrhagic risk 3.8%/y**

# Annual Thromboembolic Risk

in pts with non valvular AF

Treated with Warfarin or New Oral Anticoagulants

■ % Stroke/year in Warfarin      ■ % Stroke/year in New Oral Anticoagulants

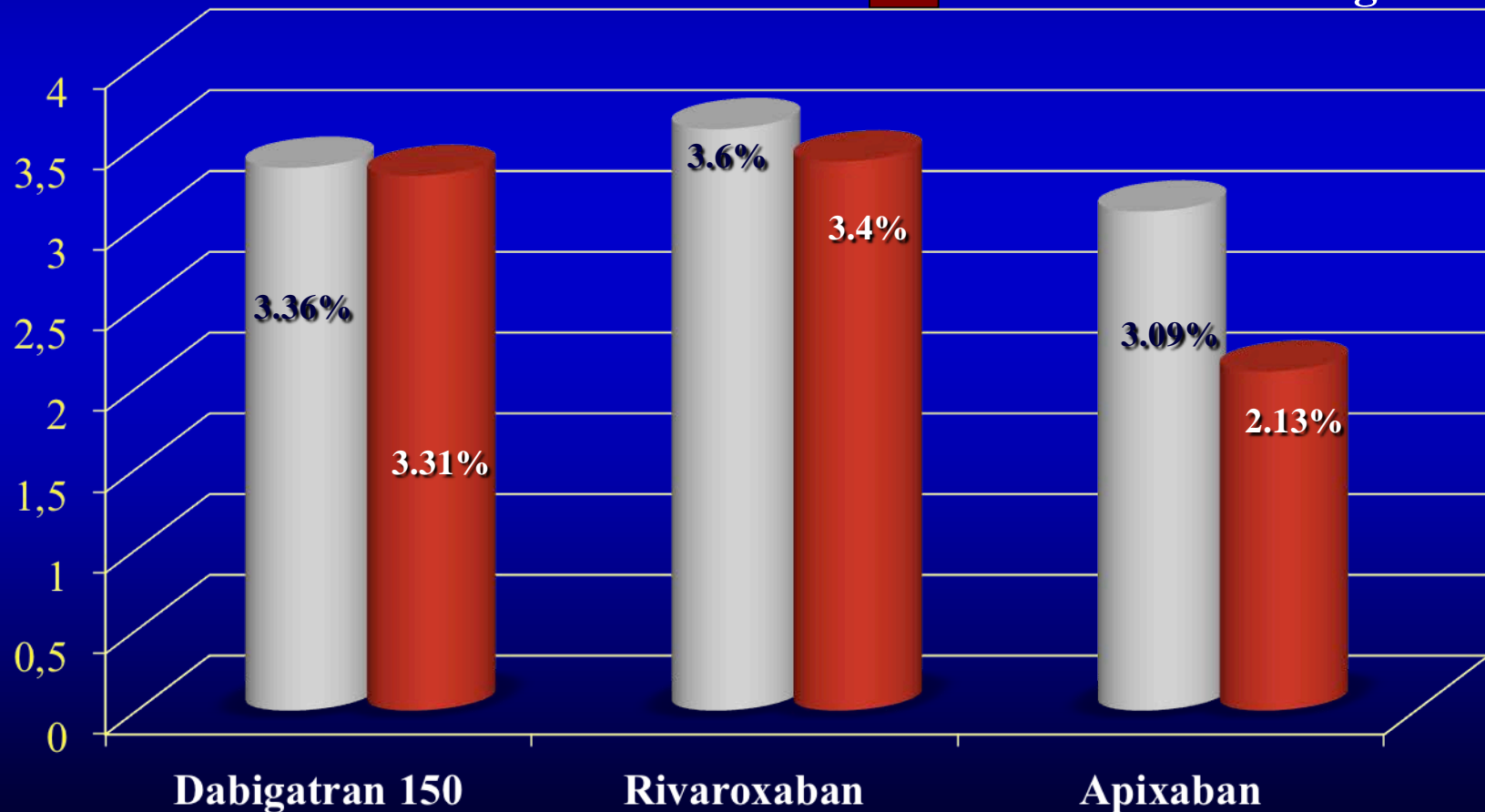


# Annual Hemorrhagic Complications

in pts with non valvular AF

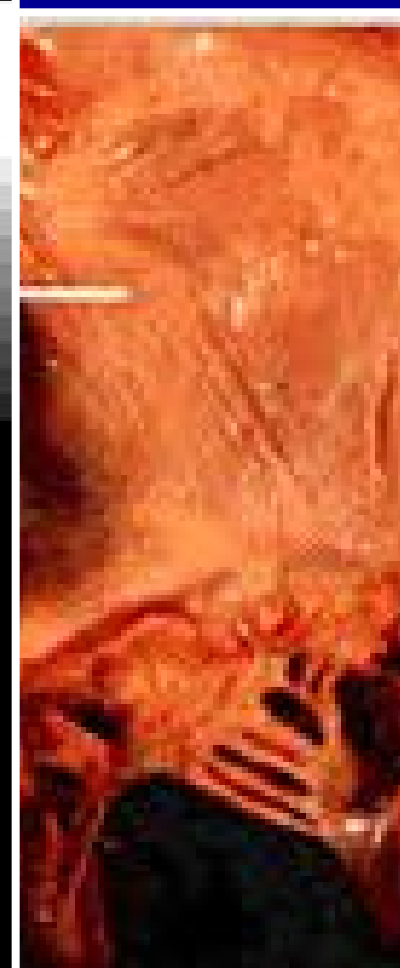
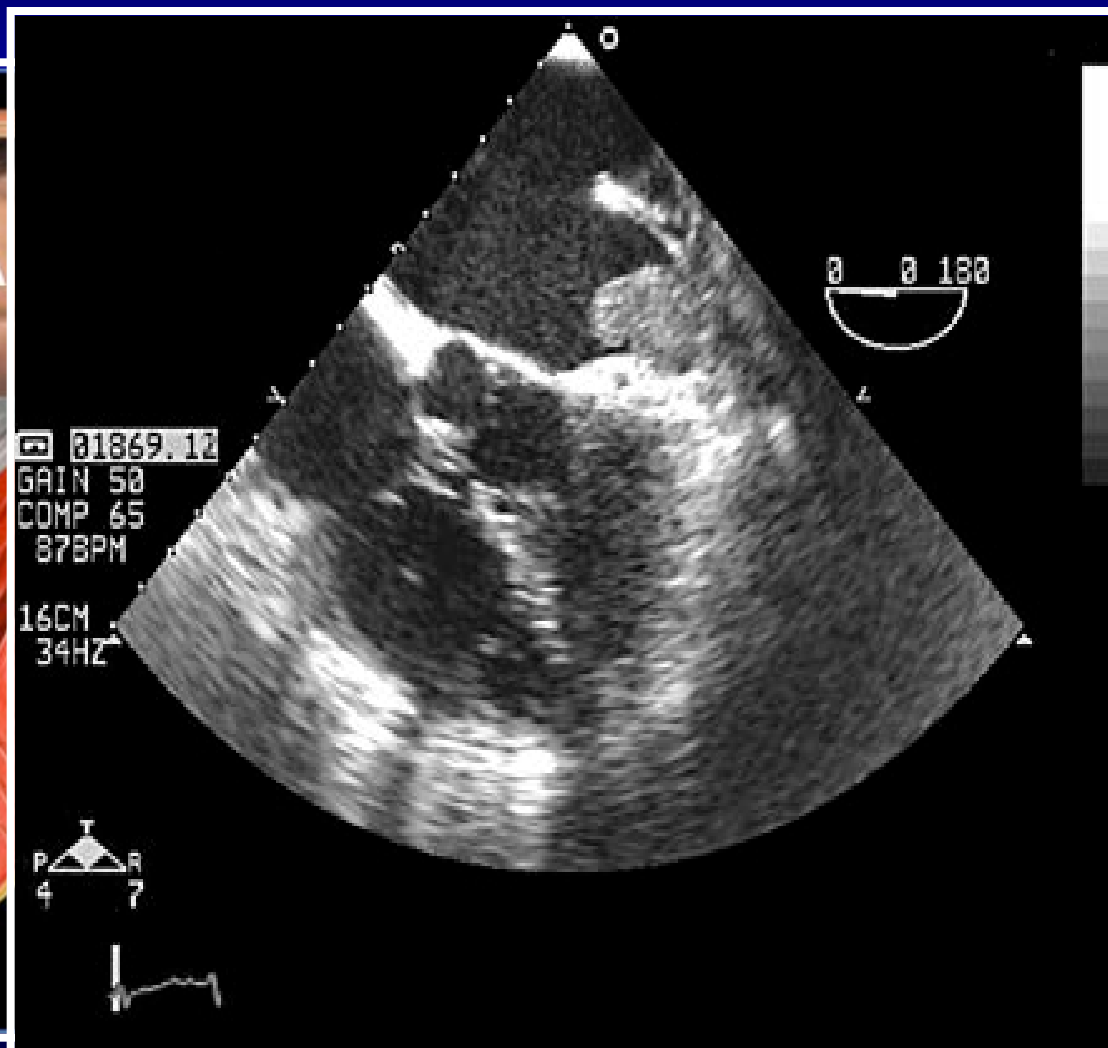
Treated with Warfarin or New Oral Anticoagulants

■ % Major Bleeding/year in Warfarin      ■ % Major Bleedings/year in New Oral Anticoagulants

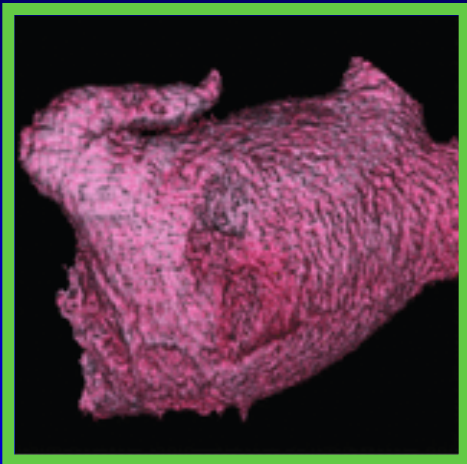


# LAA and thromboembolic risk

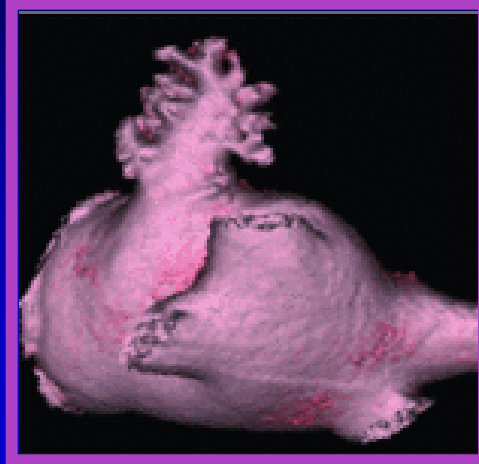
Incidence of thrombus in LAA reaches up to 91%



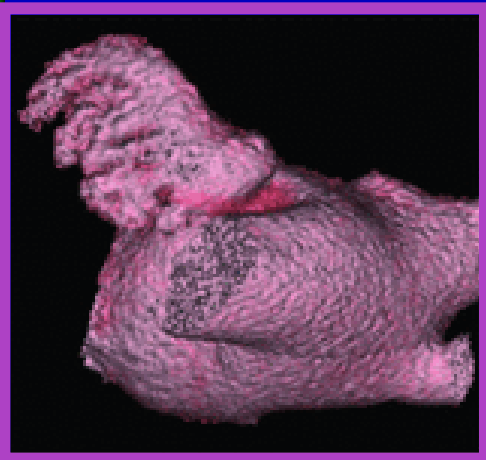
Chickenwing type LAA,  
451 (48%) pts



Cactus type LAA,  
278 (30%) pts



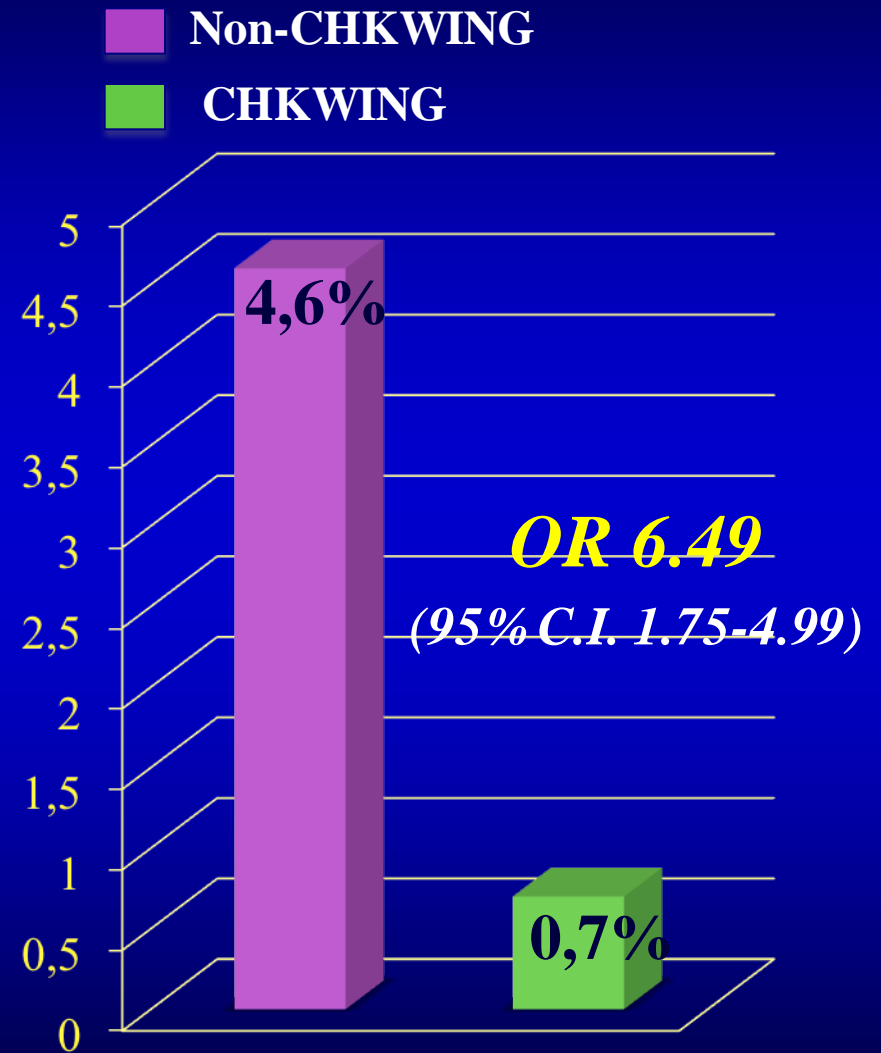
Windsock type LAA,  
179 (19%) pts



Cauliflower type LAA,  
24 (3%) pts

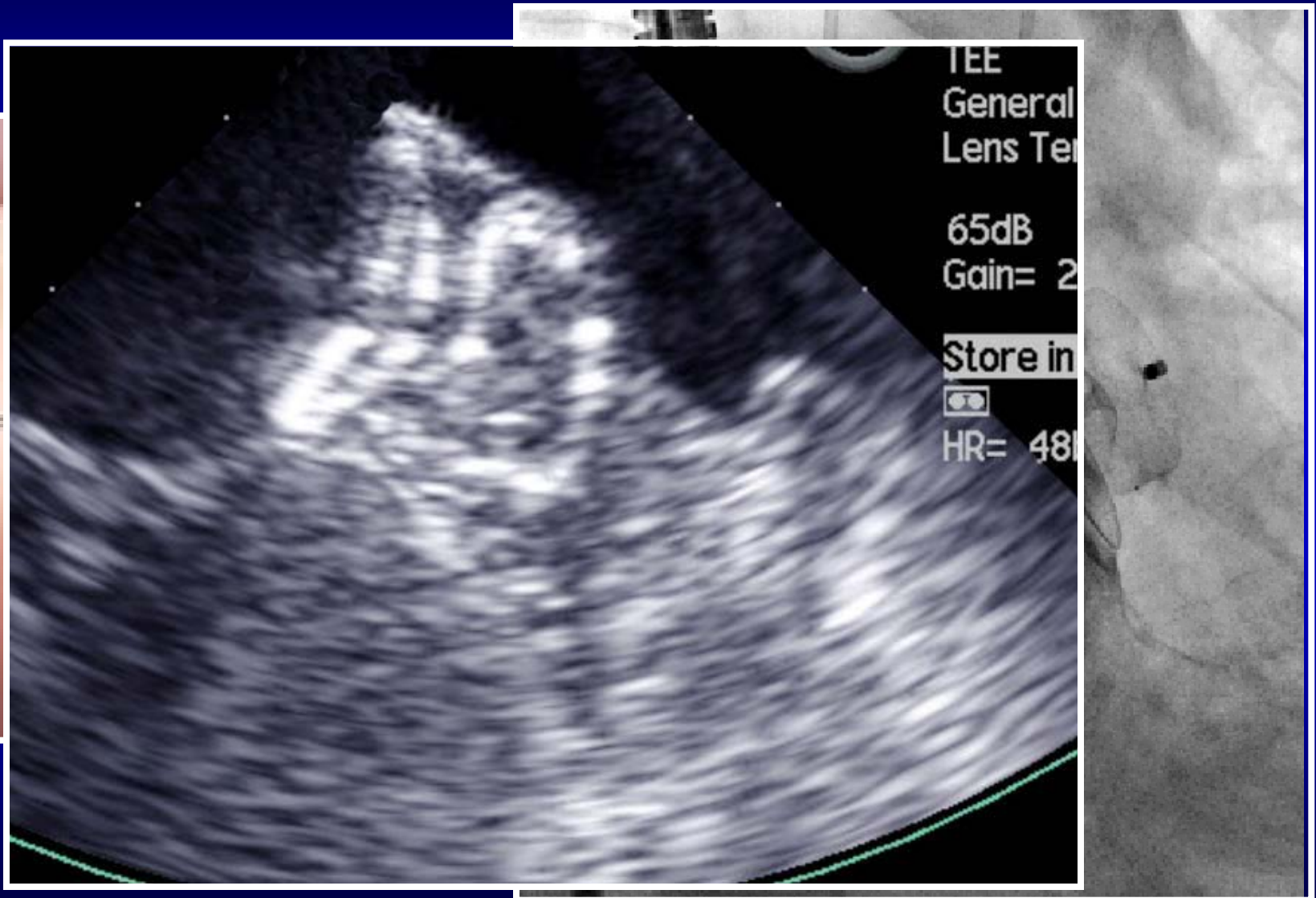


## LA appendage morphology and thromboembolic risk

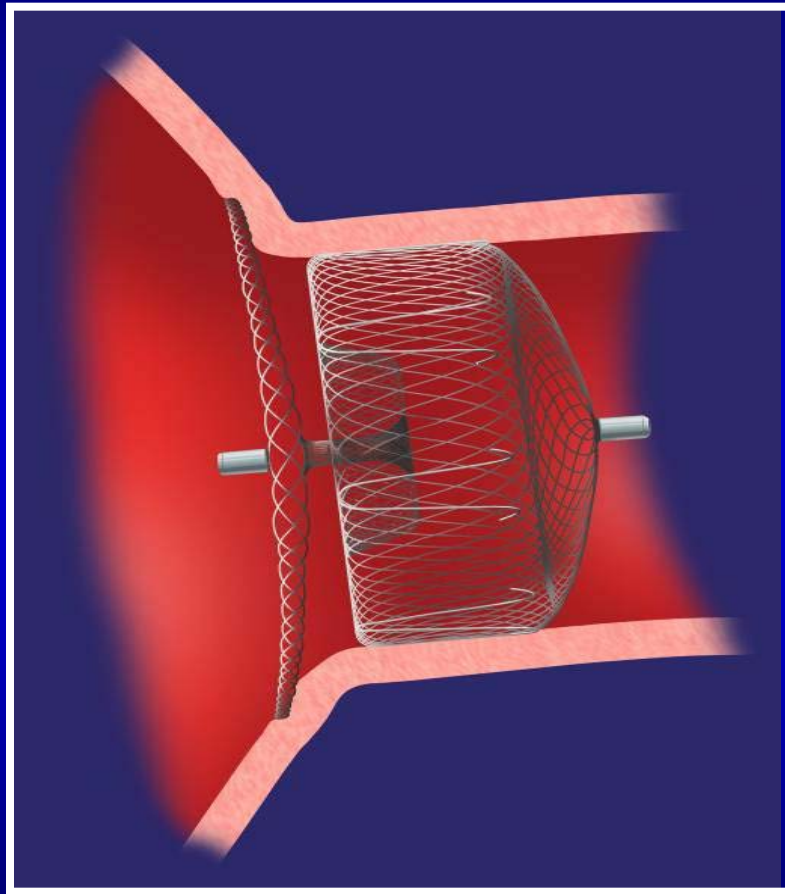




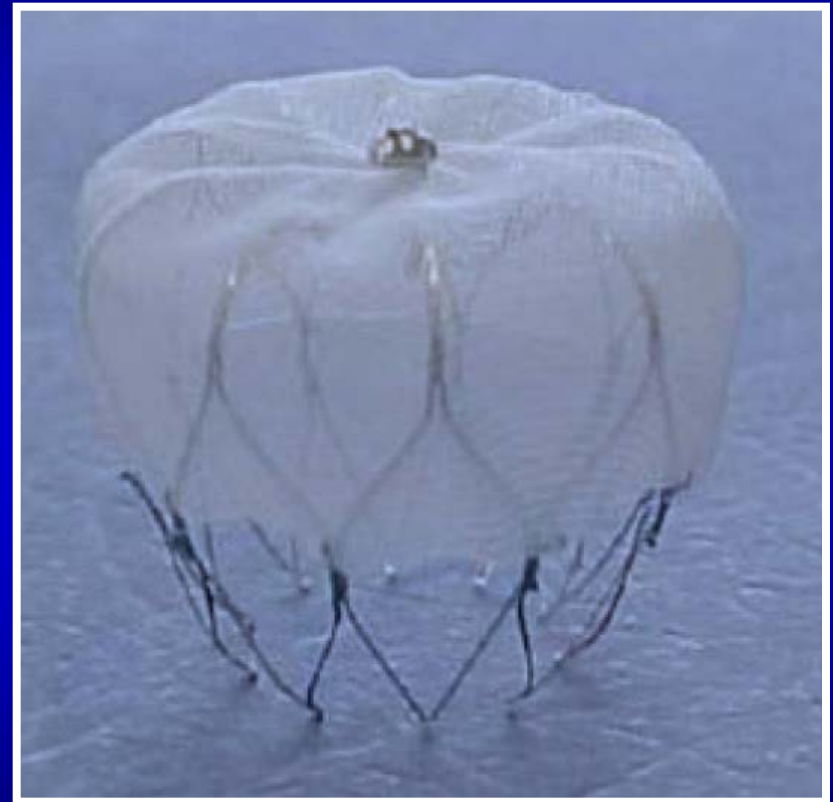
# *LA appendage closure*



# LA appendage Closure devices



**Amplatzer**

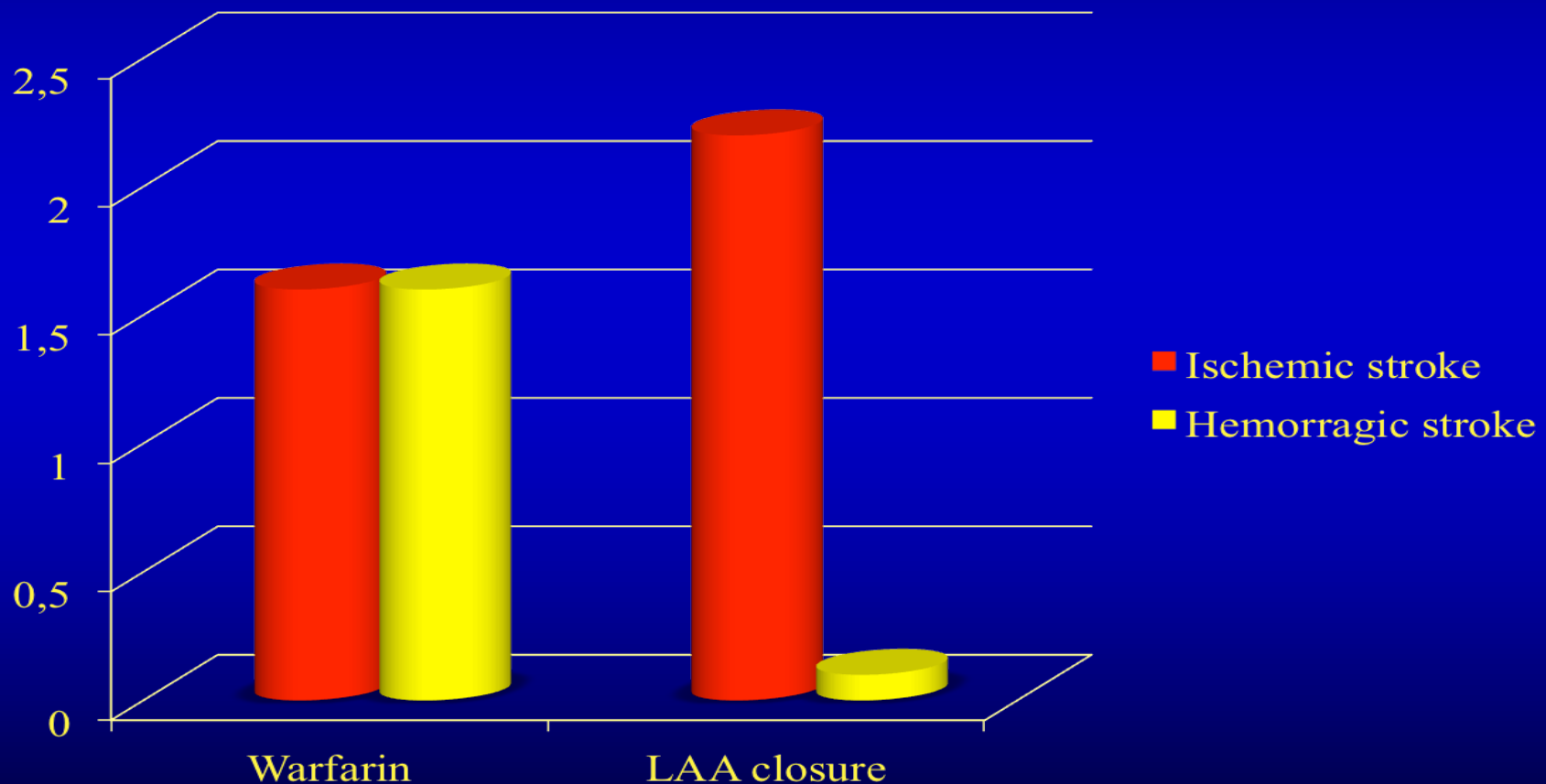


**Watchman**

# PROTECT-AF: 707 pts

M 70%, 72 y, 36% permanent AF, 38% previous stroke/TIA

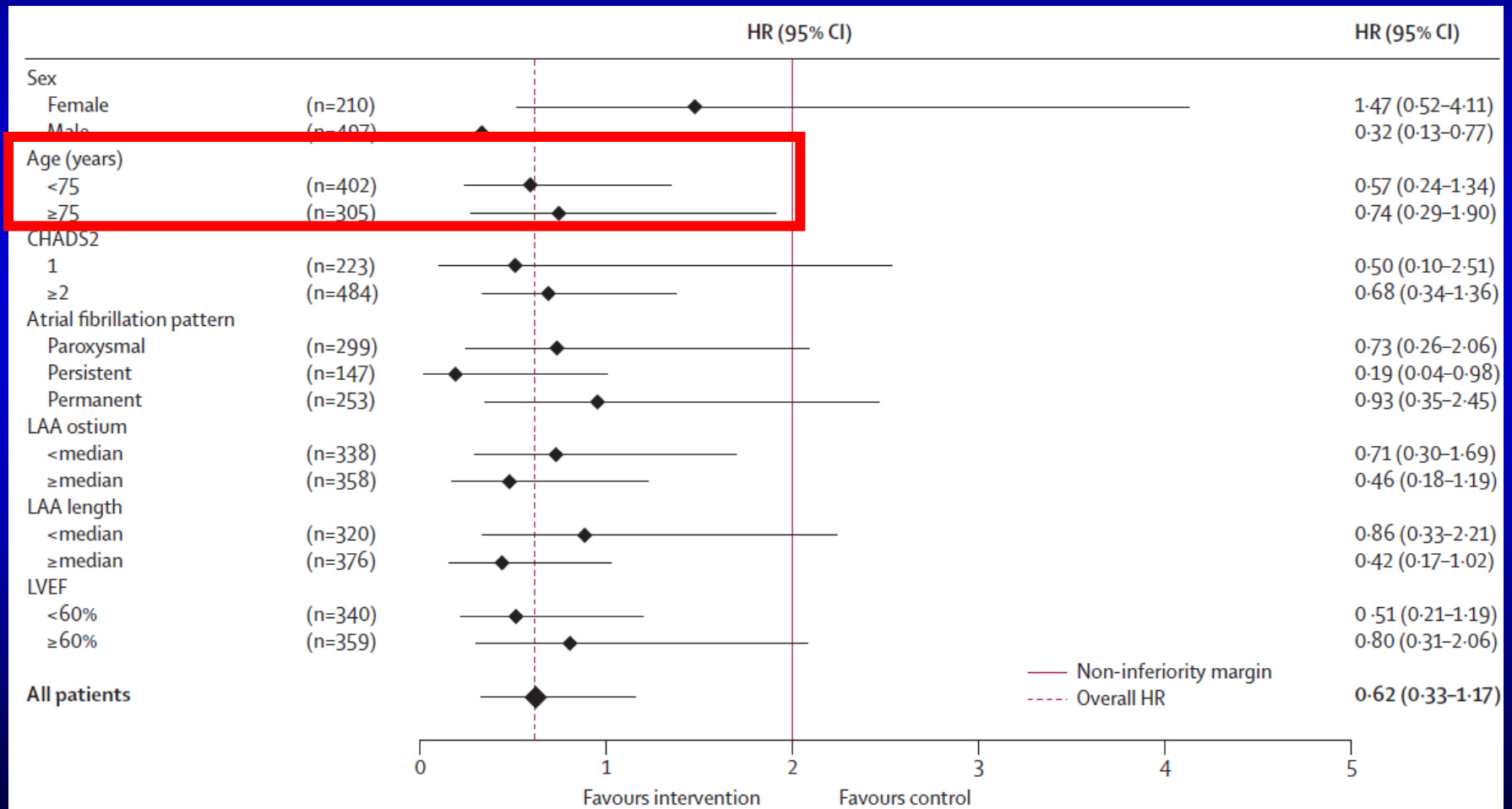
**463 percutaneous LAA closure vs 244 Warfarin (INR 2-3)**



# PROTECT-AF: 707 pts

M 70%, 72 y, 36% permanent AF, 38% previous stroke/TIA

463 percutaneous LAA closure vs 244 Warfarin (INR 2-3)



# Updated ESC guidelines

## Recommendations for LAA closure/occlusion/excision

Recommendations	Class <sup>a</sup>	Level <sup>b</sup>	Ref <sup>c</sup>
Interventional, percutaneous LAA closure may be considered in patients with a high stroke risk and contraindications for long-term oral anticoagulation.	IIb	B	115, 118
Surgical excision of the LAA may be considered in patients undergoing open heart surgery.	IIb	C	



# AFFIRM: Total Mortality (at 5 years)

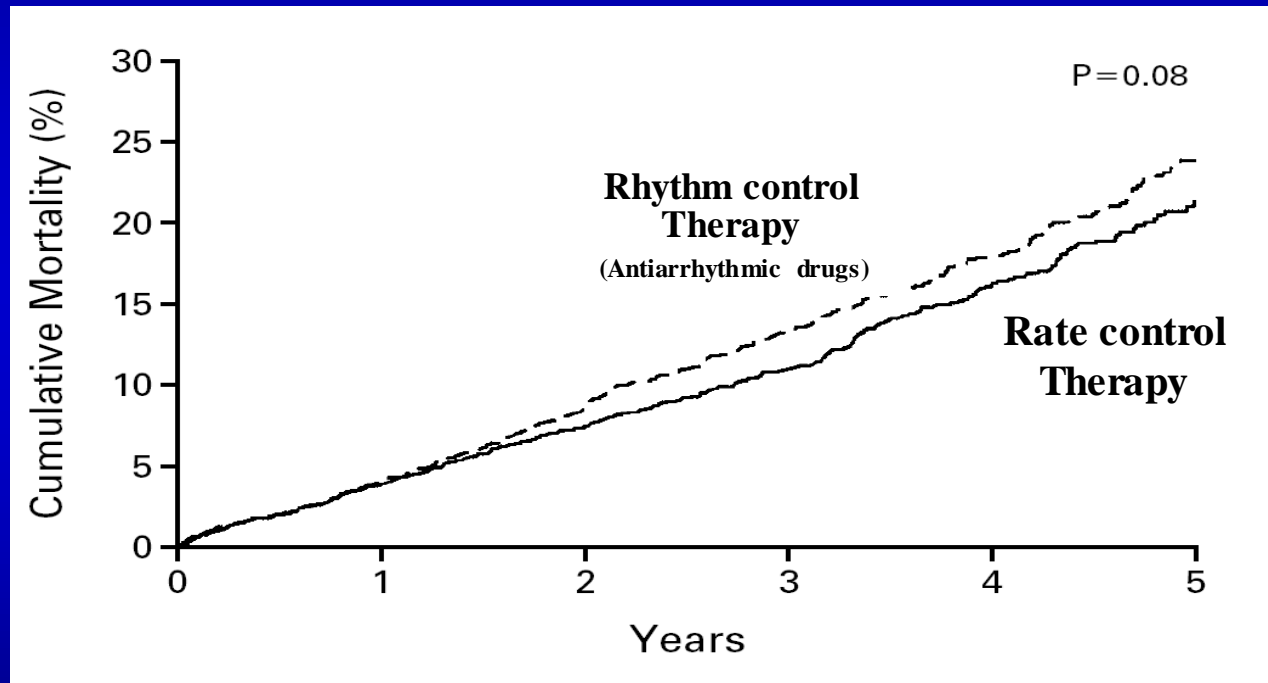
Rate control *vs* Rhythm control

Rate control therapy *vs* Rhythm control therapy

**4060 pts,**  
**Age 69.7±9 years**  
**528 pts (13%) > 80 y**

- 70.8% Hypertension
- 38.2% Ischemic
- ↓ EF 26%
- ↑ Left atrium 64,7%

Mean FU: 3,5 y



	NO. OF DEATHS					
		number (percent)				
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)

# AF therapy in the Elderly

## RATE CONTROL THERAPY

```
graph LR; A[RATE CONTROL THERAPY] --> B[No difference in term of mortality, compared to SR maintenance]; A --> C[Easily achieved in persistent AF]; A --> D["β-blockers simpler to manage than AADs"]; A --> E[Avoids proarrhythmic effects of AADs];
```

No difference in term of mortality,  
compared to SR maintenance

Easily achieved in persistent AF

$\beta$ -blockers simpler to manage than  
AADs

Avoids proarrhythmic  
effects of AADs

*..but our patient had Paroxysmal AF and Sinus Bradycardia...*

# AFFIRM:

“On treatment” analysis in a subgroup of 2796 pts

Covariates associated to survival:

Covariate	p	HR
Sinus Rhythm	<0.0001	0.53
Warfarin	<0.0001	0.50
AADs	0.0005	1.49
Digoxin	0.0007	1.42

**RS e Warfarin**  
↓ of about 50%  
risk of death

**AADs and digoxin**  
↑ of about 50%  
risk of death

# **New Therapies for SR maintenance**

**2010:**

**Dronedarone**



# ATHENA TRIAL

**4500 patients with paroxysmal (75%) / persistent AF (25%)**

**Inclusion criteria:**

**age  $\geq 75$  years**

**or  $\geq 70$  years with  $\geq 1$  risk factor:**

**hypertension; diabetes; prior stroke/TIA;**

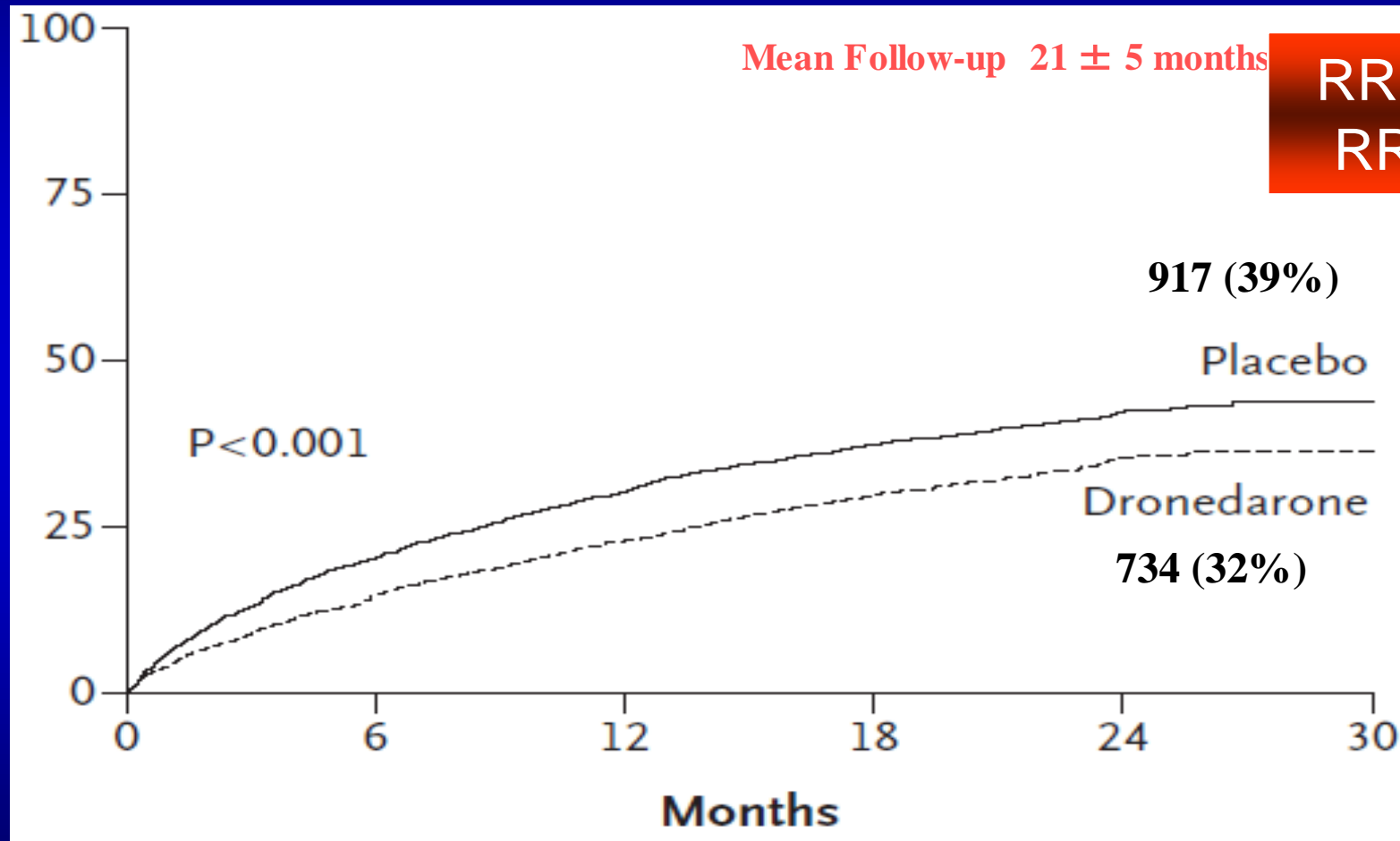
**LA  $\geq 50$  mm; LVEF  $\leq 0.40$**

**Dronedaronone 400 mg BID vs Placebo**

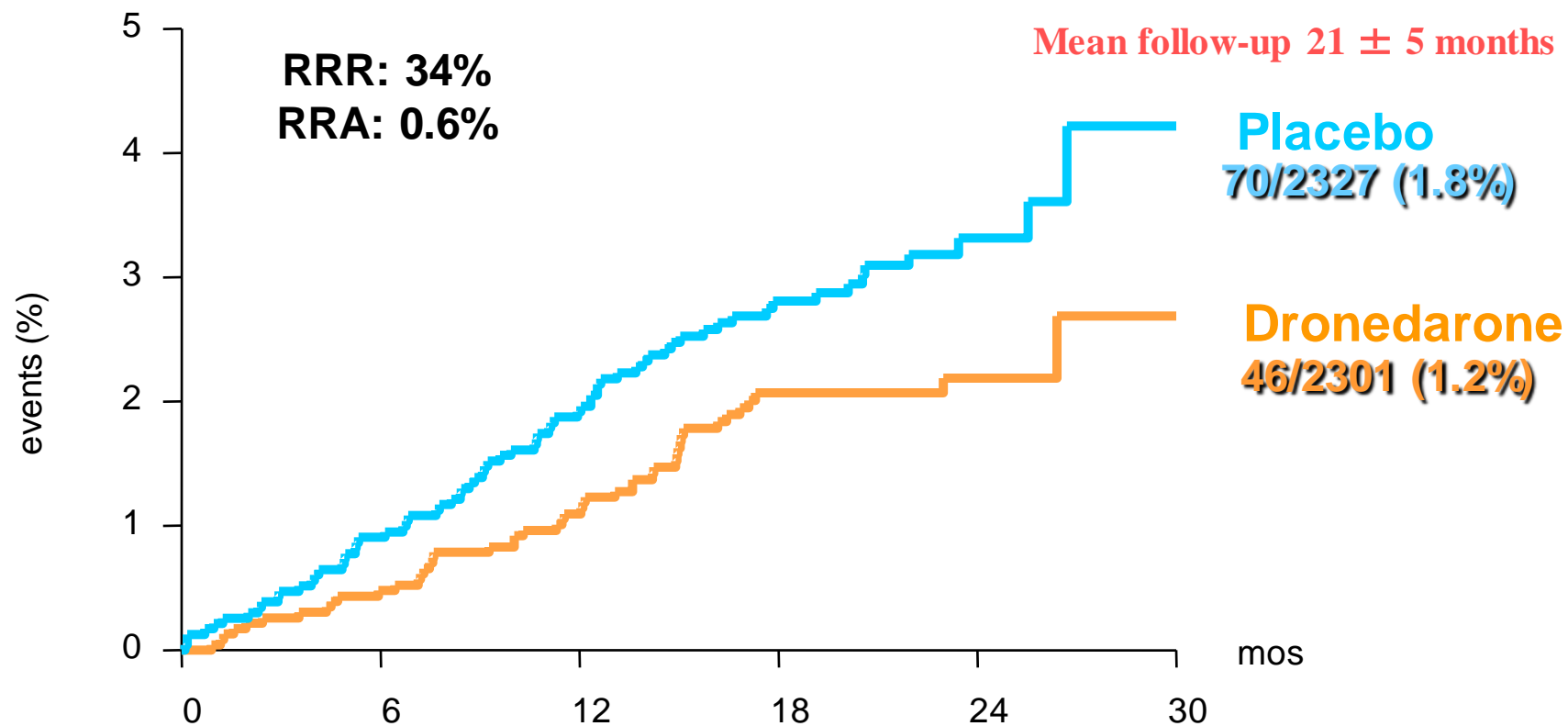


# Primary Endpoint

## Time to first cardiovascular hospitalization or death



# ATHENA Substudy: Analysis of Stroke



Pz a rischio

Placebo

Dronedaronone

2327

2275

2220

1598

618

6

2301

2266

2223

1572

608

4



# ATHENA TRIAL

4500 patients with paroxysmal (75%) / persistent AF (25%)

**Inclusion criteria:**

age  $\geq 75$  years

or  $\geq 70$  years with  $\geq 1$  risk factor:

hypertension; diabetes; prior stroke/TIA;

LA  $\geq 50$  mm; LVEF  $\leq 0.40$

**Dronedaronone 400 mg BID vs Placebo**

**Exclusion criteria:**

***Sinus-node disease, bradycardia (HR < 50 bpm)***

**PR > 0.28 sec, GFR < 10 ml/min; K < 3.5 mmol/l**

# In case of SSS, of about 30% of patients need PM implant

## The preferred stimulation is AAI

### Permanent transvenous Atrial Pacing: an experimental and clinical study

*Smyth N.P.D. et al ann Thorac Surg 1971;11:360-370*

South Med J. 1975 May;68(5):580-3.

### The sick sinus syndrome: treatment by permanent transvenous atrial pacing (a new approach).

Moore CB, Bower PJ.

Bifascicular bundle branch block	37
Blood pressure >250/120 mm Hg	37
Atrial fibrillation >50% of time (2-4 days before implantation)	34
Age <50 yr	17
Atrial fibrillation with RR interval >3 s	12
Cardiac surgery planned	9
Cancer	8

### DANISH STUDY Lancet 1994;344:1523-28

### Long-term follow-up of patients from a randomised trial of atrial versus ventricular pacing for sick-sinus syndrome

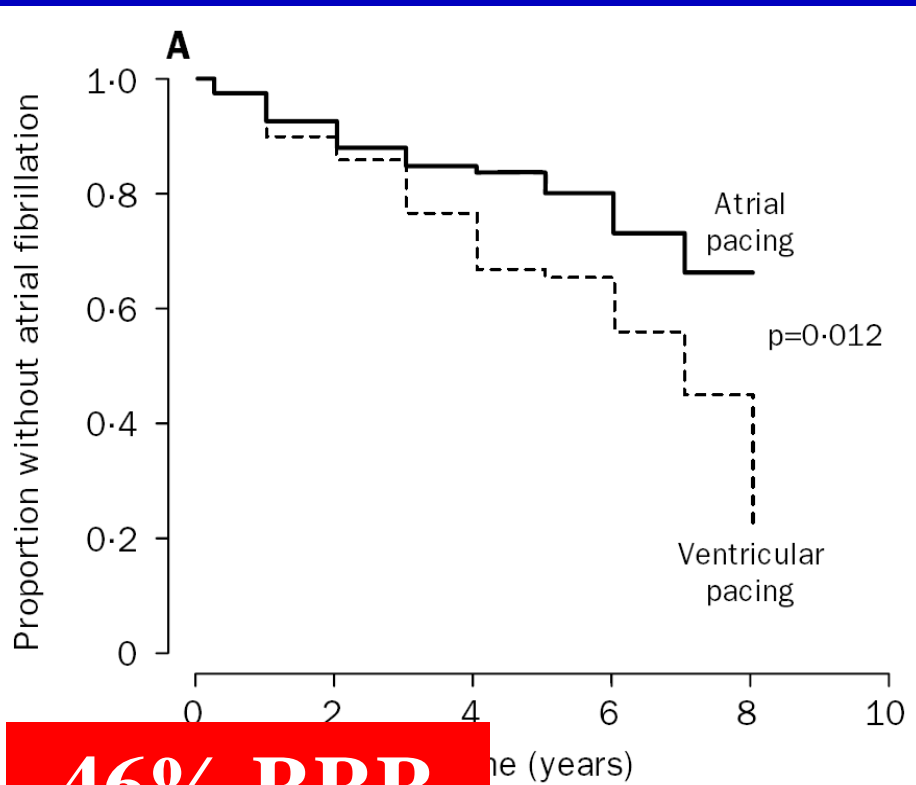
*Henning Rud Andersen, Jens Cosedis Nielsen, Poul Erik Bloch Thomsen, Leif Thuesen, Peter Thomas Mortensen, Thomas Vesterlund, Anders Kirstein Pedersen*

**Lancet 1997;350:1210-16**

# Long-term follow-up of patients from a randomised trial of atrial versus ventricular pacing for sick-sinus syndrome

Henning Rud Andersen, Jens Cosedis Nielsen, Poul Erik Bloch Thomsen, Leif Thuesen, Peter Thomas Mortensen, Thomas Vesterlund, Anders Kirstein Pedersen

## Atrial Fibrillation



**46% RRR**

**225 pts,**  
**age 76 y with SSS**  
**Atrial pacing (104 pts)**  
**VS**  
**Ventricular pacing (115 pts)**



# Clinical Outcomes After Ablation and Pacing Therapy for Atrial Fibrillation

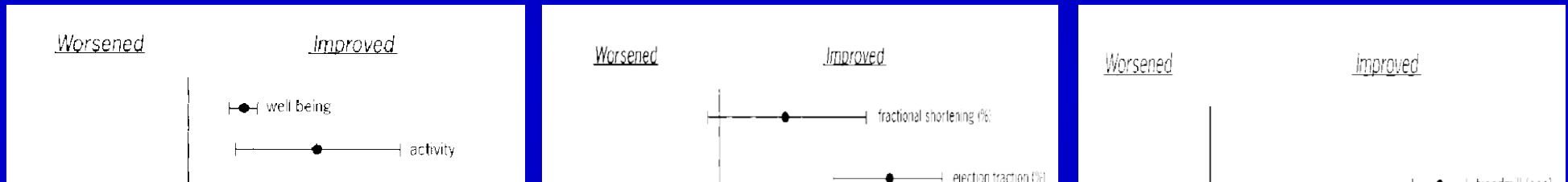
## A Meta-Analysis

*1181 patients with symptomatic and medically refractory AF*

*Qol*

*LV function, Healthcare Use  
NYHA*

*Exercise duration,  
heart rate*



**Ablation and pacing therapy improves a broad range of clinical outcomes for patients with AF.**

**The calculated 1-year mortality rates (6.3%) are low and comparable with medical therapy**

0.4 0.2 0.2 0.4 0.6 0.8 1.0

4 2 2 4 6

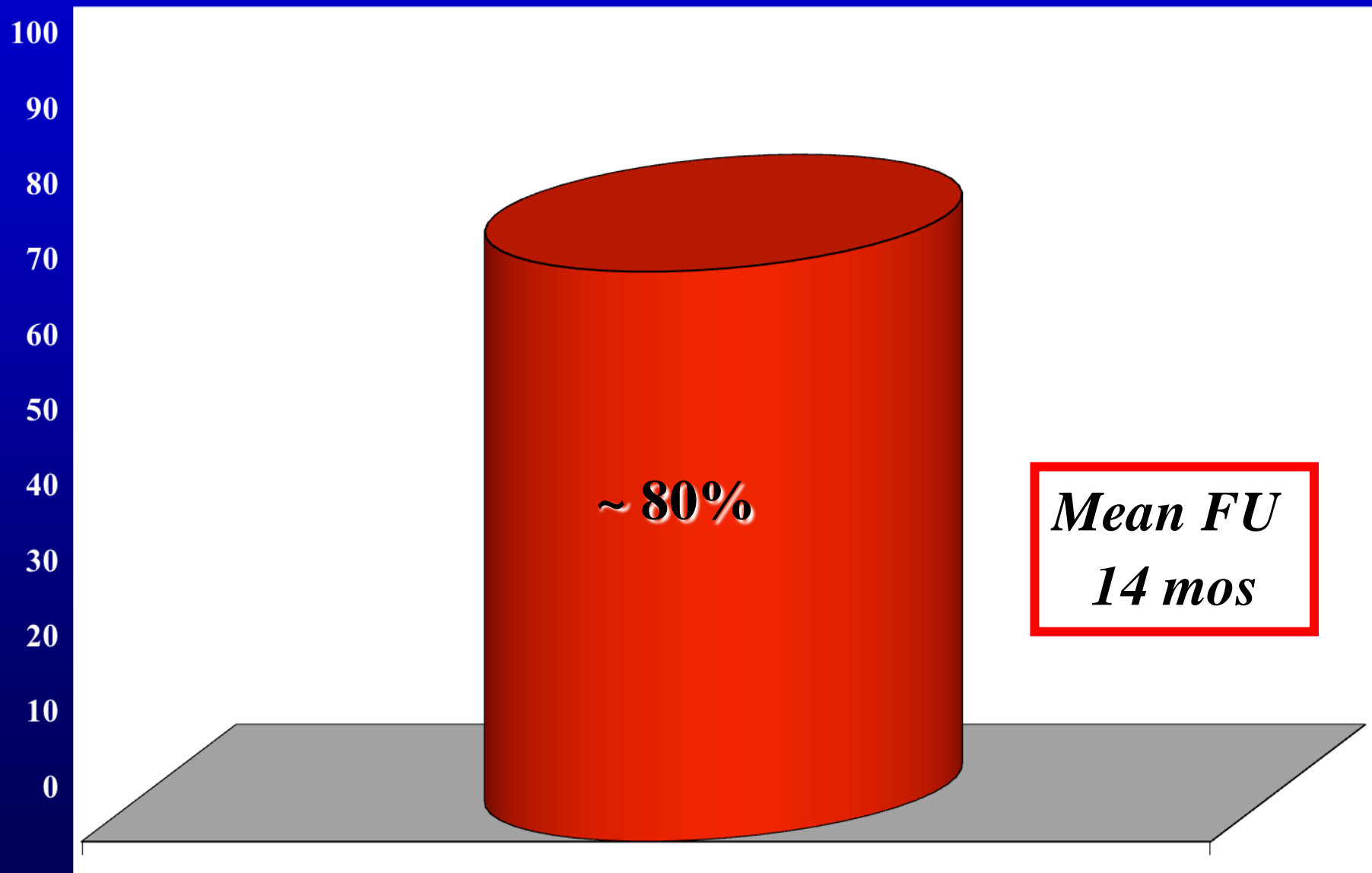
50 25 25 50 75 100 125

# Why don't try Afib ablation in the Elderly?



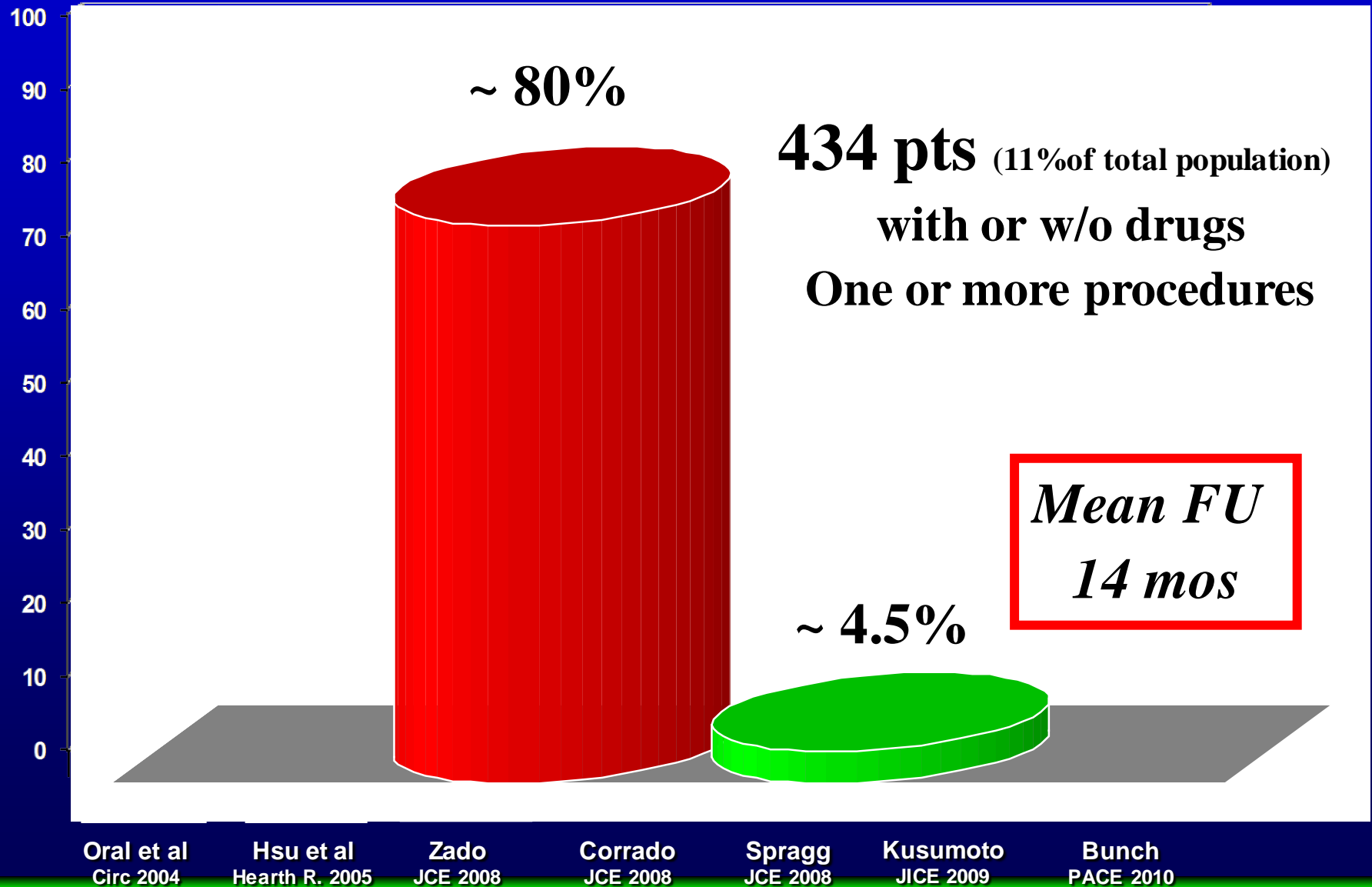
## EFFICACY of AF Ablation in the Elderly:

7 retrospective studies 434 pts > 70 years / 3935 total population (11%)



# SAFETY of AF Ablation in the Elderly:

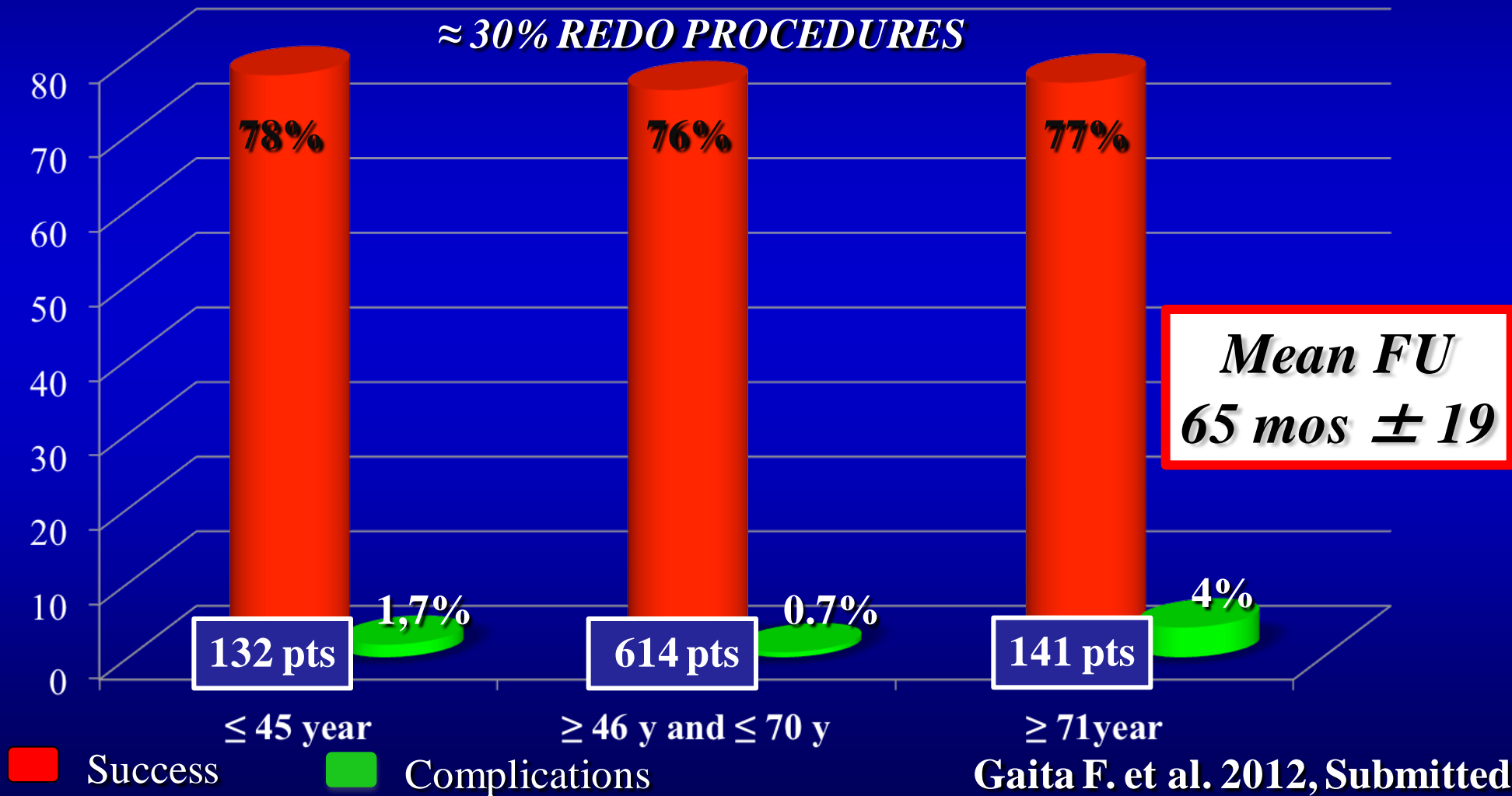
7 retrospective studies 434 pts > 70 years / 3935 total population (11%)



# *Efficacy and Safety of Afib ablation at Long-term follow up stratified for age (paroxysmal and persistent AF)*

*887 pts, 1241 procedures (Jan 2001 and Jan 2009)*

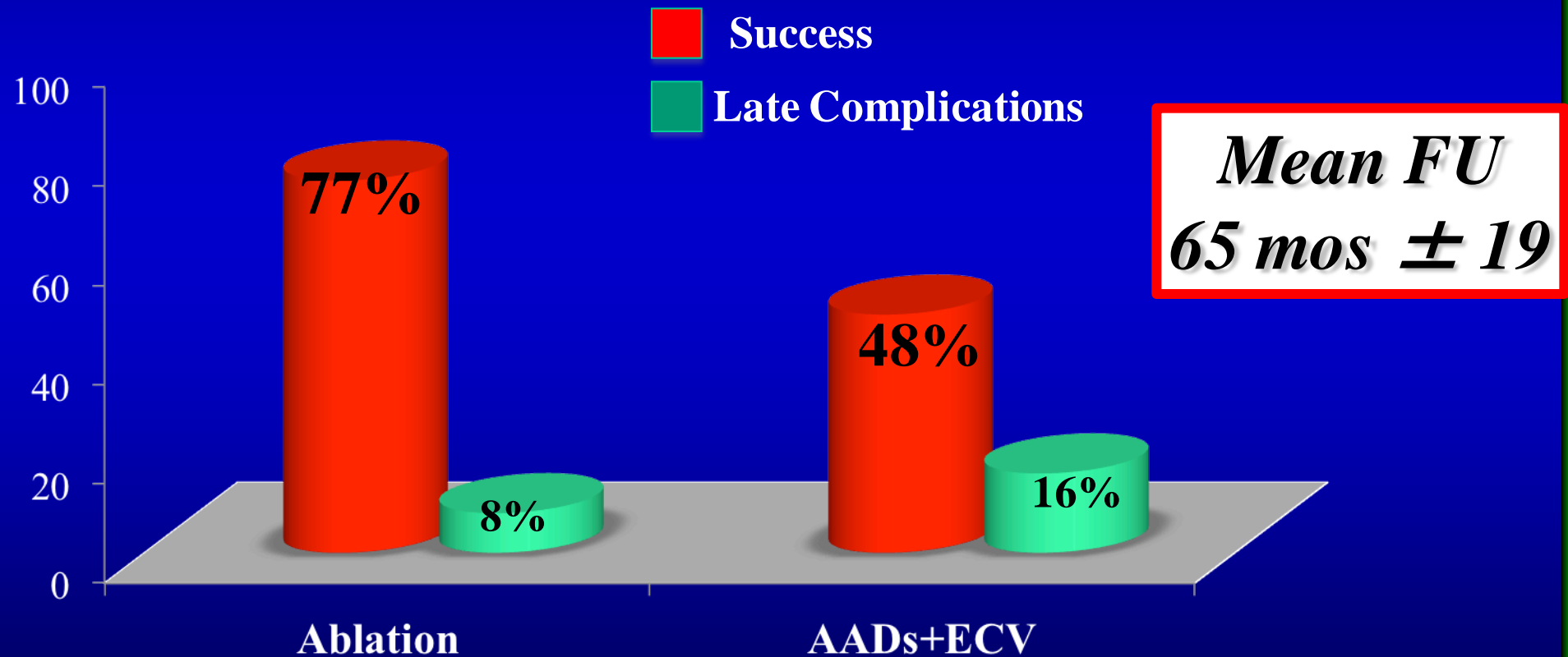
*≈ 30% REDO PROCEDURES*



Long -term efficacy and safety of **ablation** compared to **AADs** in elderly patients with AF

**344 pts age > 70 y**

118 ABLATION (Group A) vs 226 AADs + ECV (Group B)



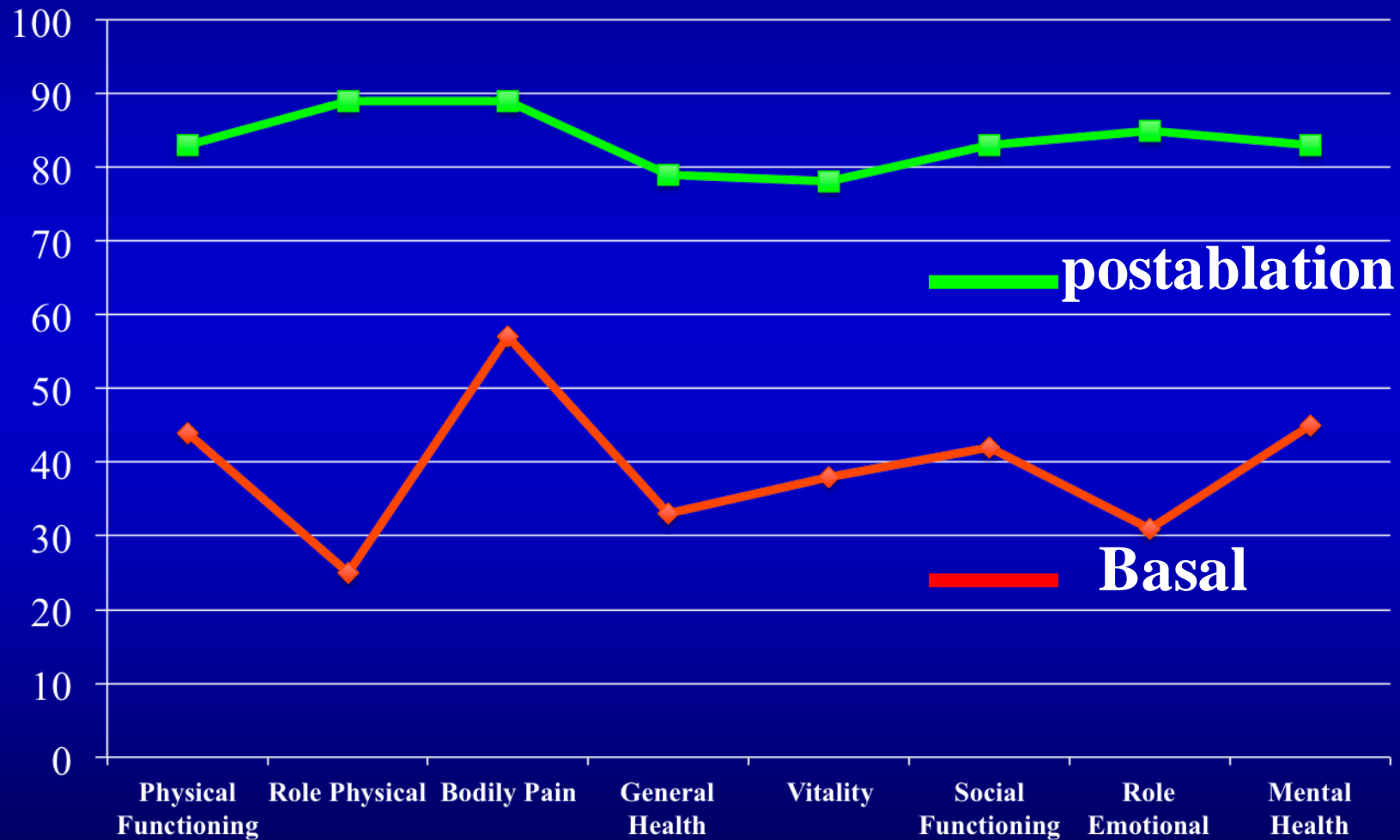
## Safety: Long-term adverse events

<b>Long-term adverse events (&gt; 30 days)</b>	<b>Ablation 118 pts (%)</b>	<b>AADs+ ECV 226 pts (%)</b>	<b>p</b>
<b>Death (CV causes)</b>	2 (1.7)	5 (2)	ns
<b>Stroke/TIA</b>	3 (2.5)	2 (0.8)	ns
<b>Peripheral Embolism</b>	1 (0.8)	1 (0.4)	ns
<b>Myocardial infarction</b>	0	2 (0.8)	ns
<b>Major Bleedings</b>	1 (0.8)	2 (0.8)	ns
<b>Minor Bleedings</b>	1 (0.8)	10 (4)	ns
<b>SSS worsening</b>	2 (1.7)	15 (8)	<b>0.001</b>
<b>Pacemaker implant</b>	0	5	



# *Comparison on QoL (SF-36) at basal and follow up*

## *Ablation*





-Female } CHA<sub>2</sub>DS<sub>2</sub>VASC 3  
-78 y }

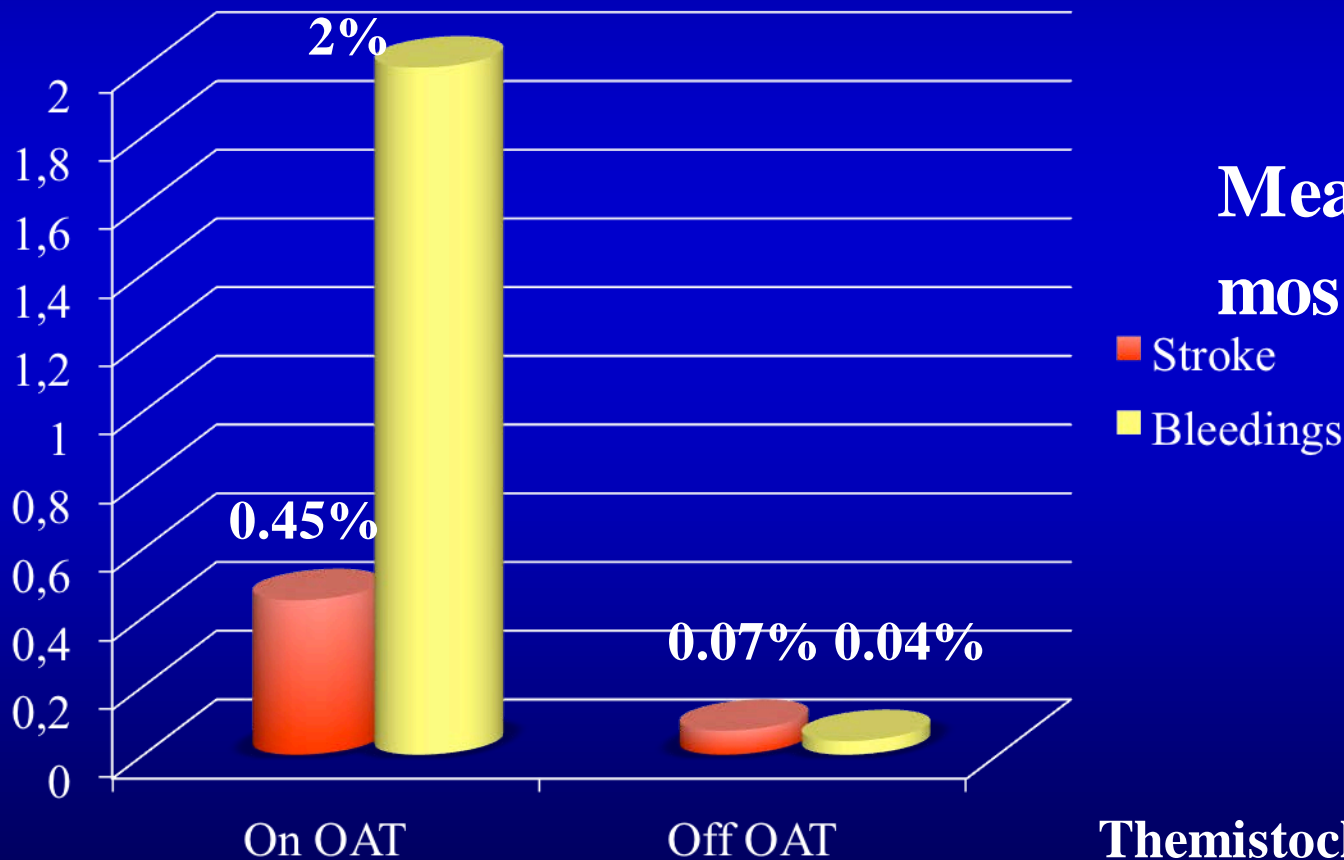
+ Atrial ~~×~~ Fibrillation  
=

Oral Anticoagulation

**...If we restore and maintain SR  
after Afib ablation, what about oral  
anticoagulation?**

# The Risk of Thromboembolism and Need for Oral Anticoagulation After Successful Atrial Fibrillation Ablation

3355 pts, OFF-OAT 2692 pts (80%), ON-OAT 663 pts (20%)



Mean FU  $28 \pm 13$

mos

Stroke

Bleedings

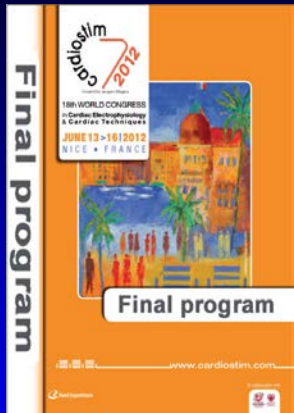
Themistoclakis S. et al. JACC 2010

# The principal answers were...

85 yr, marathon runner

Symptomatic Paroxysmal AF and Sinus bradycardia,

**CHA<sub>2</sub>DS<sub>2</sub>VASC 3, HASBLEED 3**



**AV Node Ablation+  
PM+LA appendage closure**



**Afib Ablation+Loop Recorder  
±LA appendage Closure  
(in case of recurrences)**