

**SERVIZIO SANITARIO REGIONALE  
EMILIA-ROMAGNA**  
Azienda Ospedaliero - Universitaria di Parma



# **La chiusura percutanea dell'auricola sinistra nella prevenzione dell'embolia cardiogenica**

**Umberto Scoditti**

**Programma Stroke Care**

**Dipartimento di Neuroscienze**

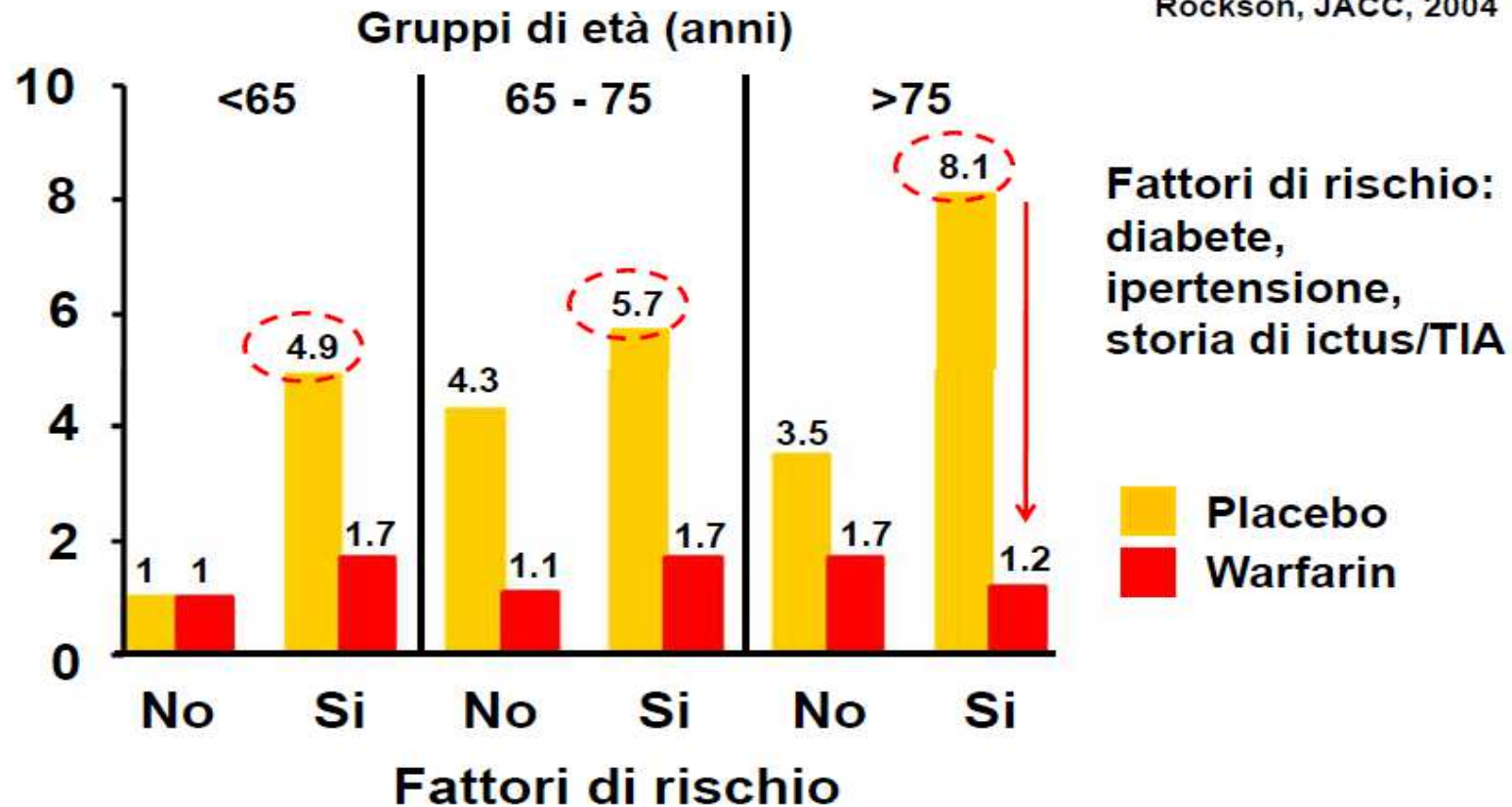
**Azienda Ospedaliero Universitaria di Parma**

**Parma 21 gennaio 2014**

# Rischio annuale di ictus in pazienti con fibrillazione atriale, per gruppi di età

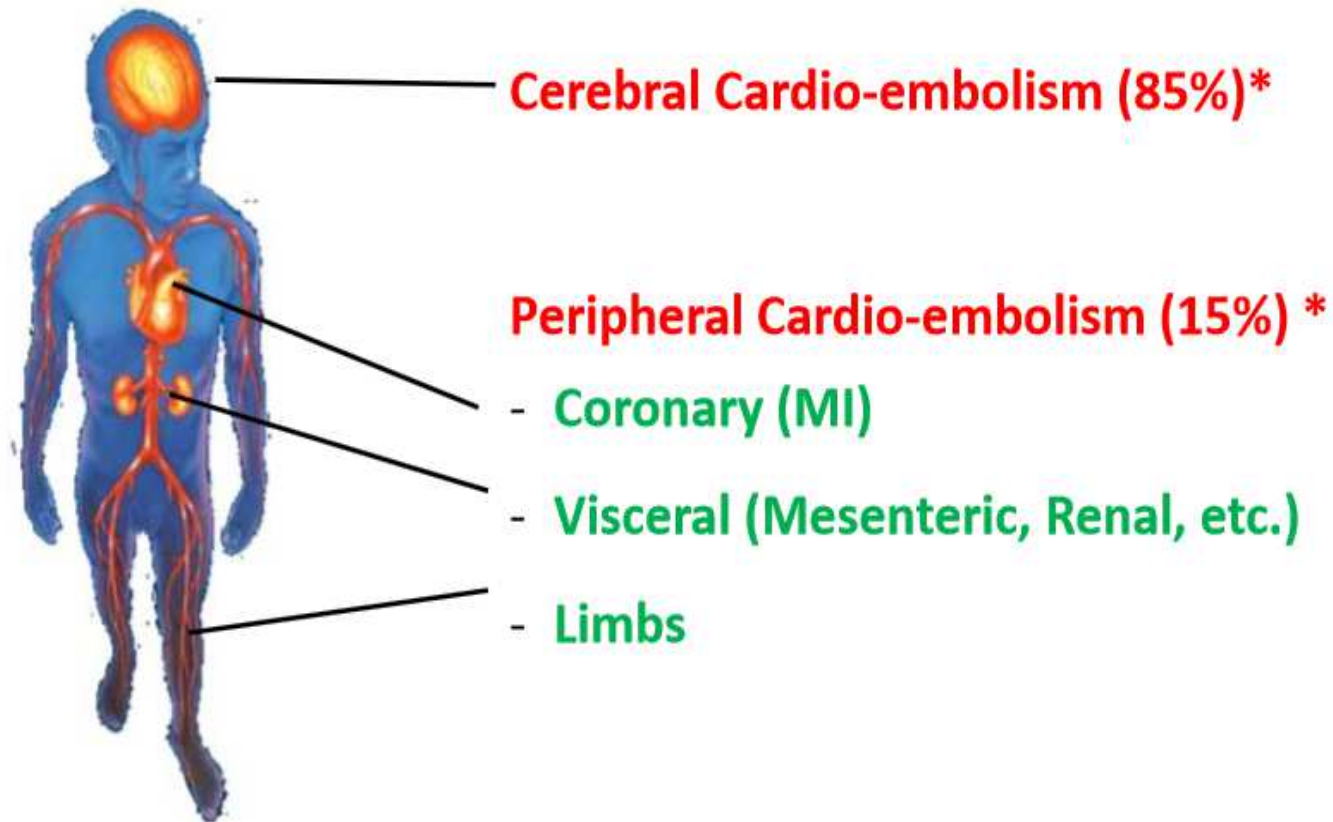
*The Atrial Fibrillation Investigators*

AFI, Arch Int Med, 1994  
Rockson, JACC, 2004



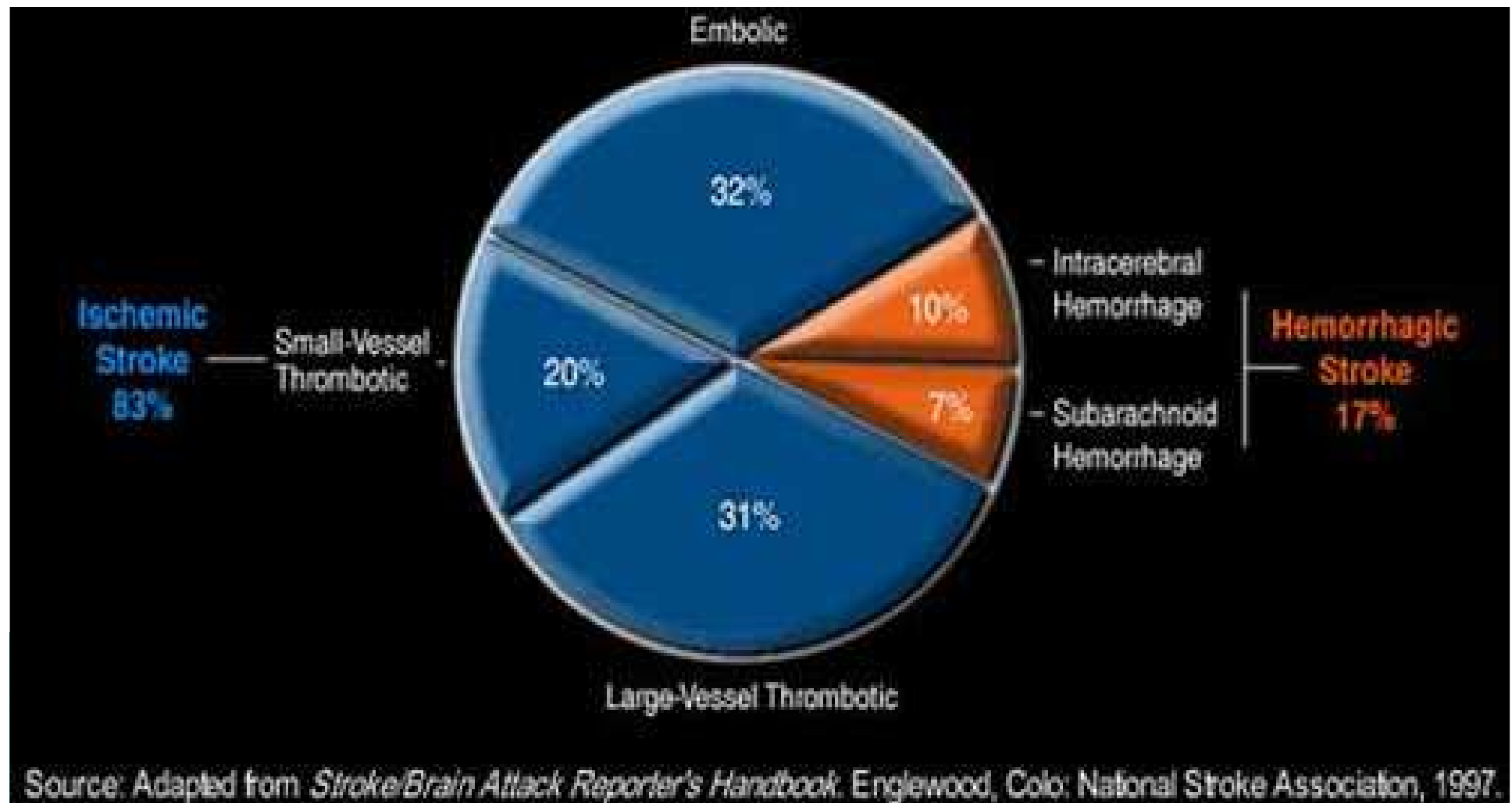
*AF identifies millions of people with a five-fold increased risk of stroke*

# Atrial Fibrillation and Cardioembolic syndromes



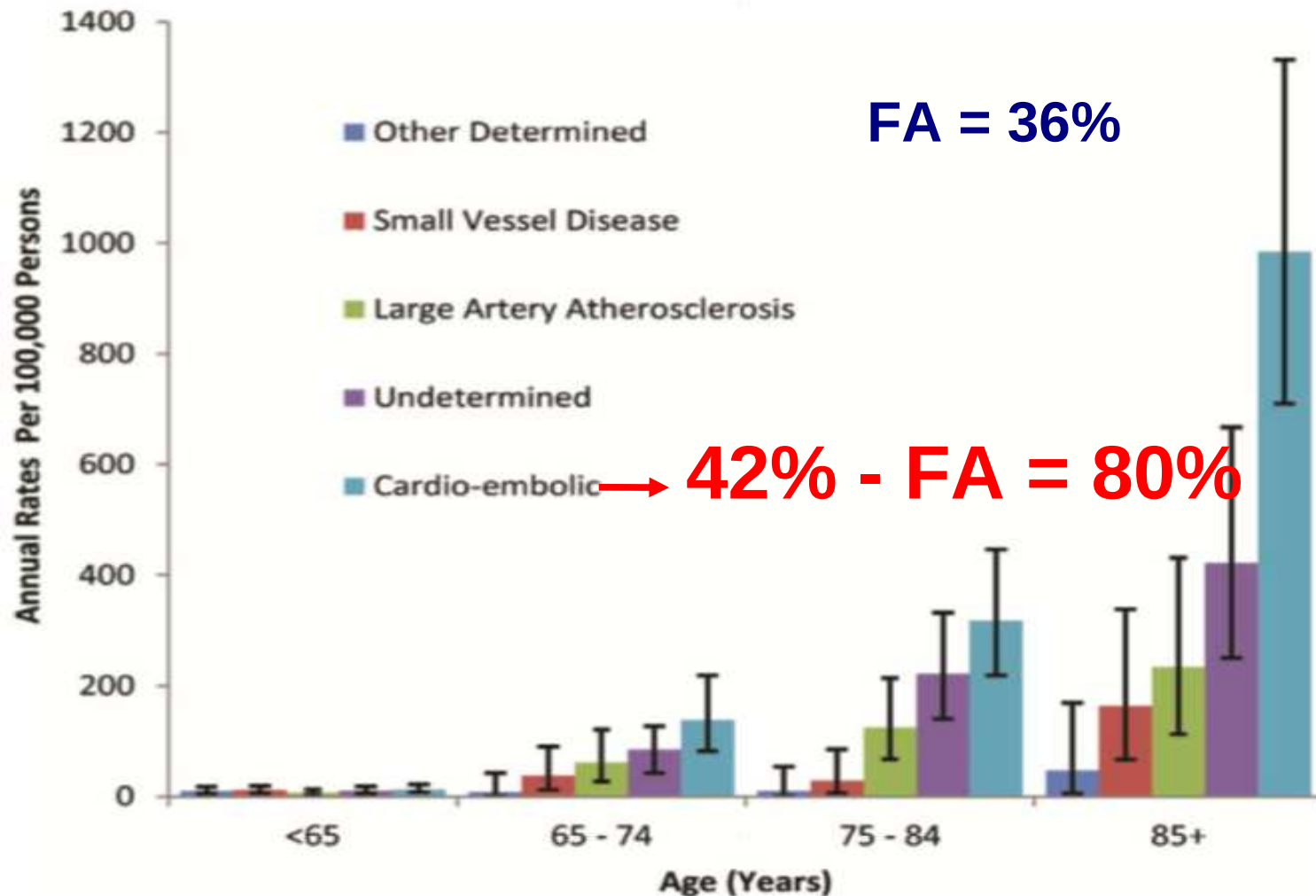
\* Cabin Am J Cardiol 1990

# Frequenza dei sottotipi di stroke



# Adelaide Stroke Incidence Study Declining Stroke Rates but Many Preventable Cardioembolic Strokes

JOURNAL OF THE AMERICAN HEART ASSOCIATION

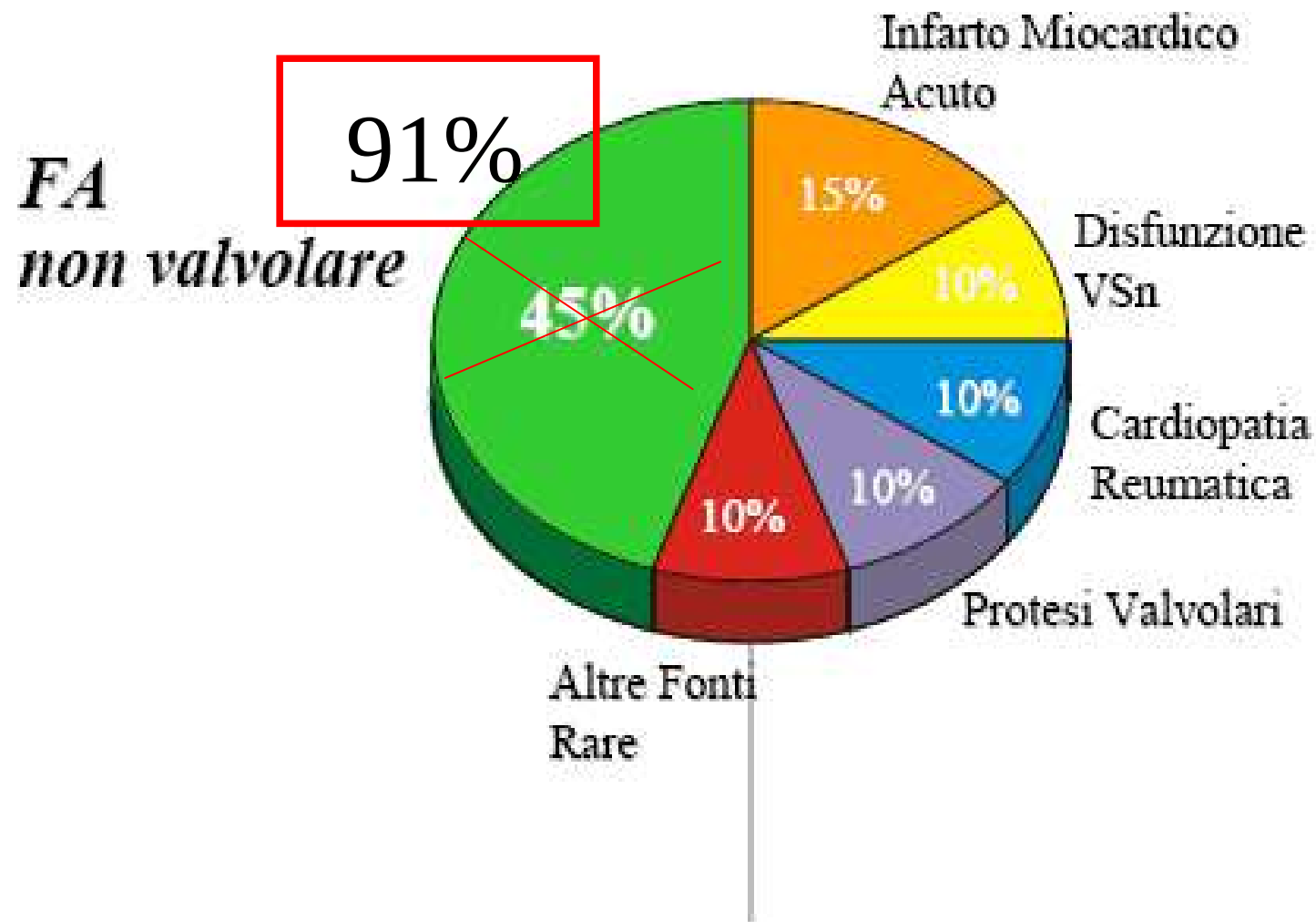
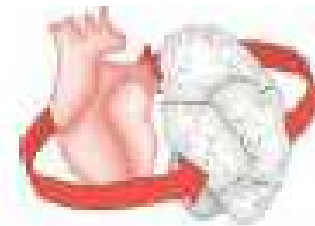


**Figure 3.** Age-specific incidence rates for all ischemic stroke subtypes in Adelaide (2009–2010).

Leyden JM, 2013

# Ictus cardio-embolico

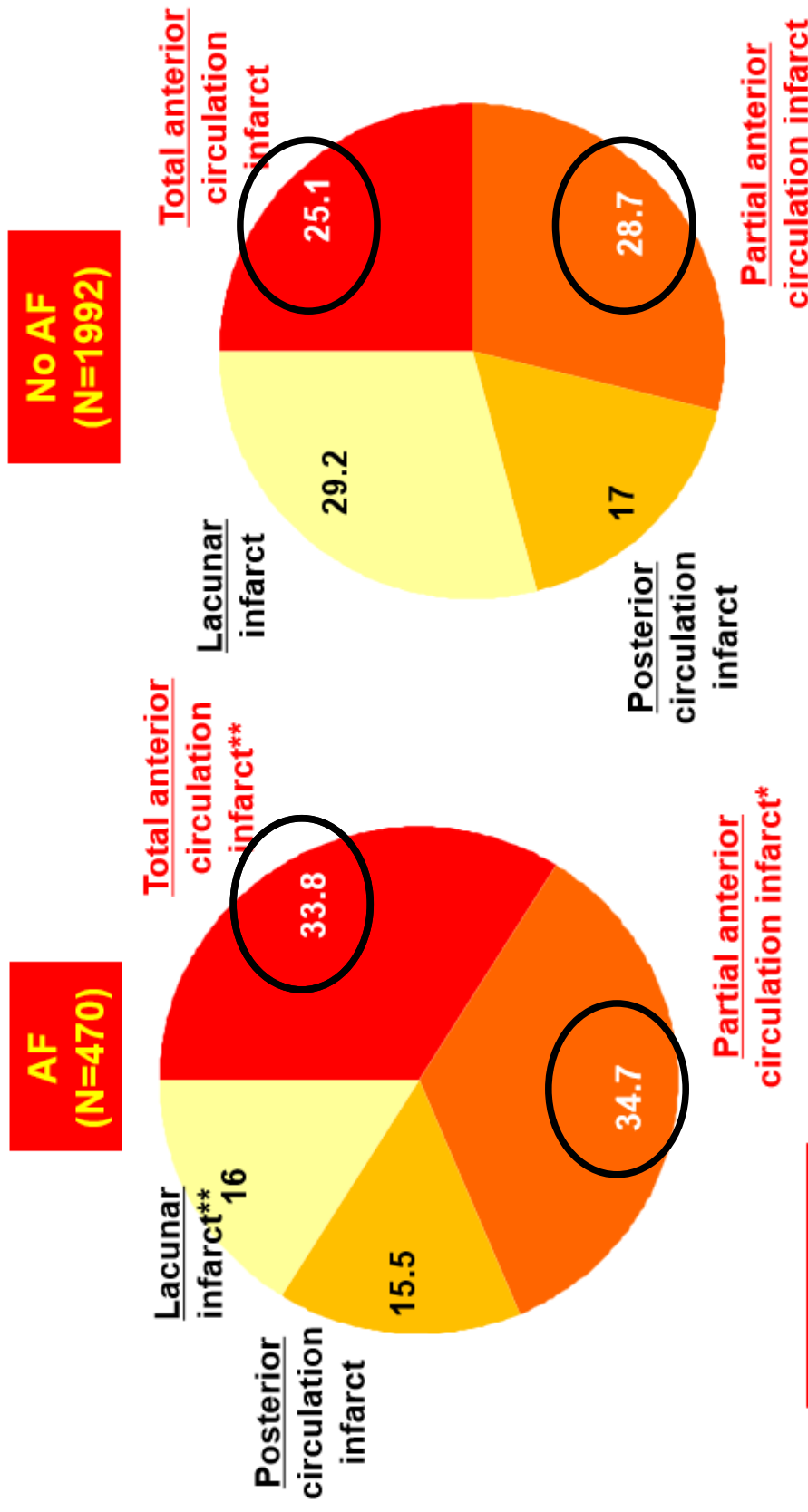
## Ruolo delle diverse cardiopatie





# Characteristics, Outcome, and Care of Stroke Associated With Atrial Fibrillation in Europe

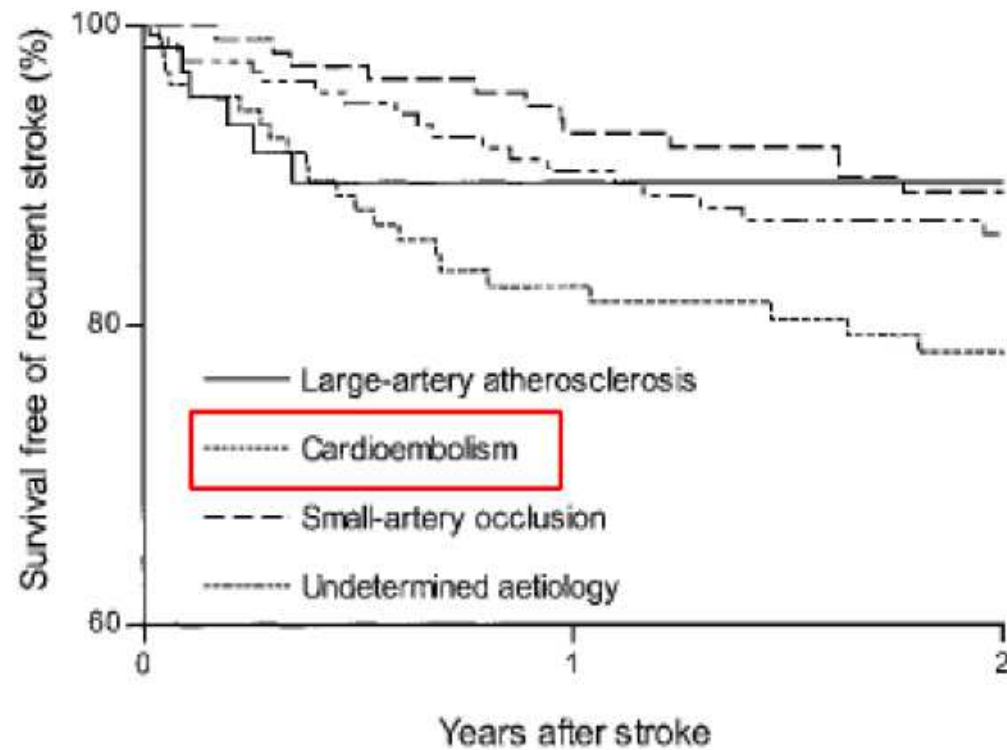
## Data From a Multicenter Multinational Hospital-Based Registry (The European Community Stroke Project)



Stroke

\*:  $p < 0.05$  vs the same category of No AF pts  
 \*\*:  $p < 0.001$  vs the same category of No AF pts

## Rischio di recidive a due anni, per tipo di ictus (n= 531)



Epidemiology of ischemic stroke subtypes according to TOAST criteria: incidence, recurrence, and long-term survival in ischemic stroke subtypes: a population-based study.  
PL Kolominsky-Rabas et al. Stroke. 2001;32:2735-2740

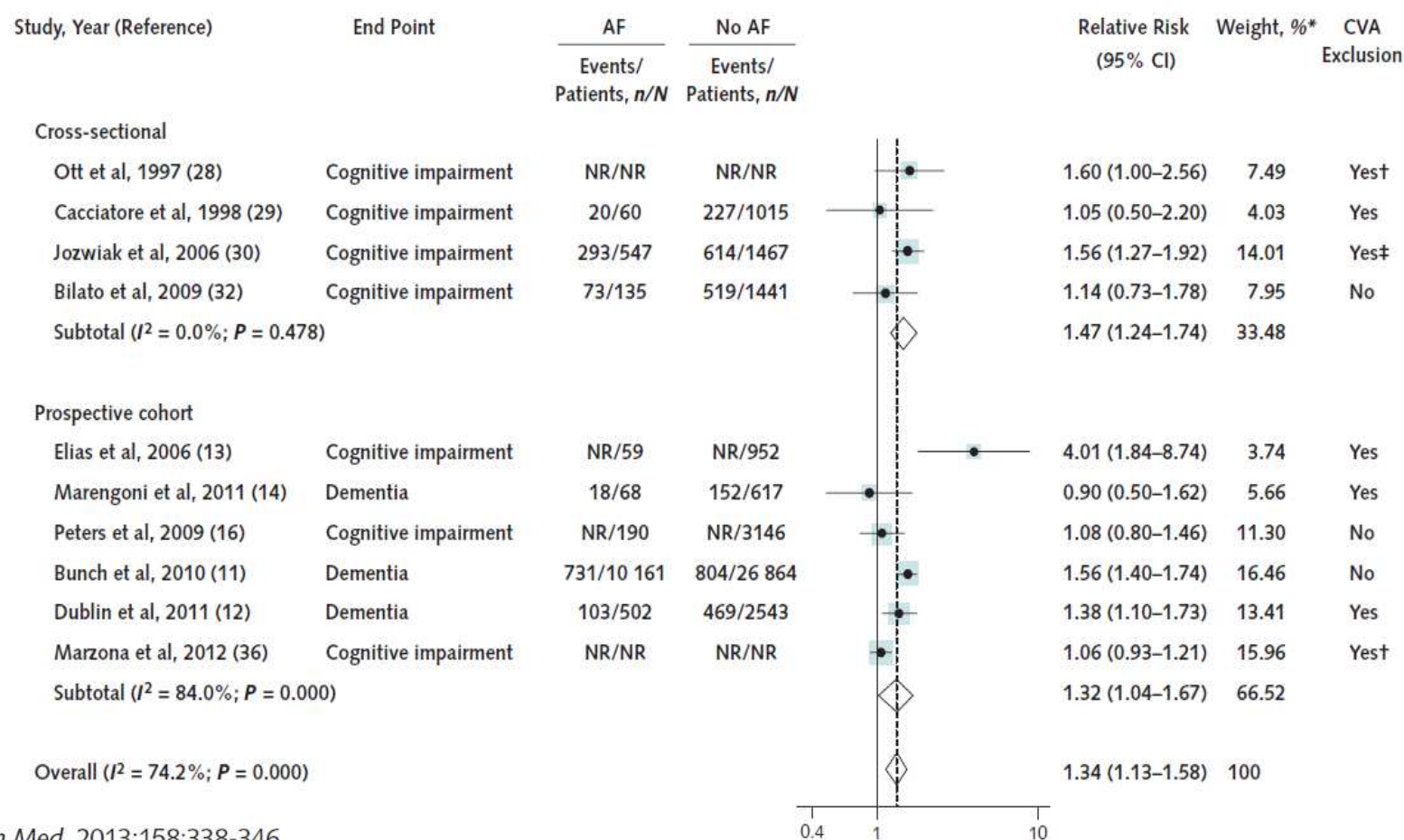


# Cognitive Impairment Associated With Atrial Fibrillation

## A Meta-analysis

Shadi Kalantarian, MD, MPH; Theodore A. Stern, MD; Moussa Mansour, MD; and Jeremy N. Ruskin, MD

Figure 3. The association between AF and cognitive impairment independent of stroke history.



# Atrial Fibrillation is Associated With Reduced Brain Volume and Cognitive Function Independent of Cerebral Infarcts

## Stroke

JOURNAL OF THE AMERICAN HEART ASSOCIATION

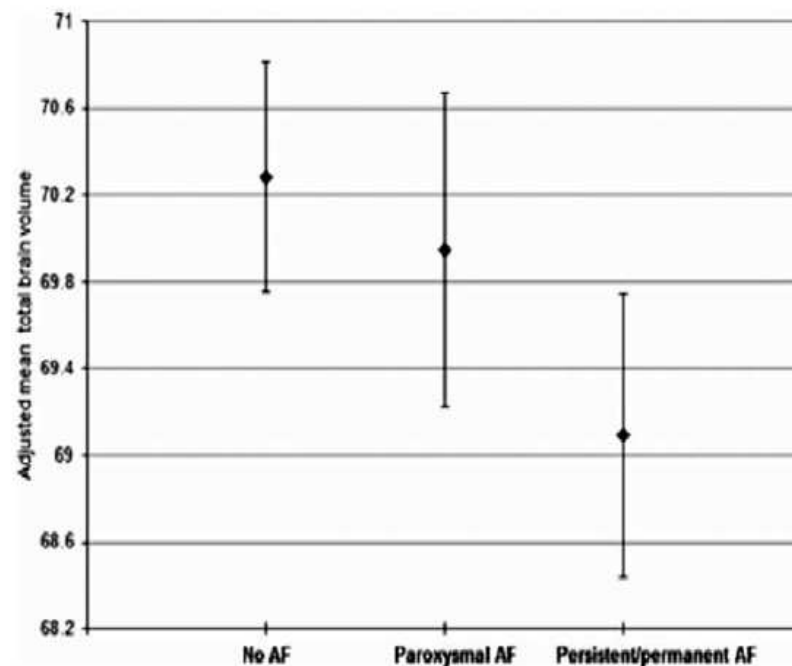


Figure 1. Mean brain volume expressed as percentage of total

Table 4. Association Between Duration of Atrial Fibrillation by Tertile and MRI-Measured Brain Volumes: The Age, Gene/Environment Susceptibility-Reykjavik Study

| Duration of AF     | Total Brain Volume | Gray Matter Volume | White Matter Volume |
|--------------------|--------------------|--------------------|---------------------|
| No AF (reference)  | 70.2               | 43.7               | 25.2                |
| 1                  | 69.5*              | 43.1*              | 25.0                |
| 2                  | 69.4*              | 42.9†              | 24.9*               |
| 3                  | 69.3†              | 43.0†              | 25.0                |
| P for linear trend | 0.012              | 0.011              | 0.16                |

AF indicates atrial fibrillation.

Table 5. Association Between Atrial Fibrillation and Cognitive Domains (Z Scores): The Age, Gene/Environment Susceptibility-Reykjavik Study

|        | Model 1       | Model 2       |
|--------|---------------|---------------|
| Memory | −0.10 (0.05)* | −0.10 (0.05)* |
| SP     | −0.06 (0.04)  | −0.03 (0.04)  |
| EF     | −0.07 (0.04)* | −0.05 (0.04)  |

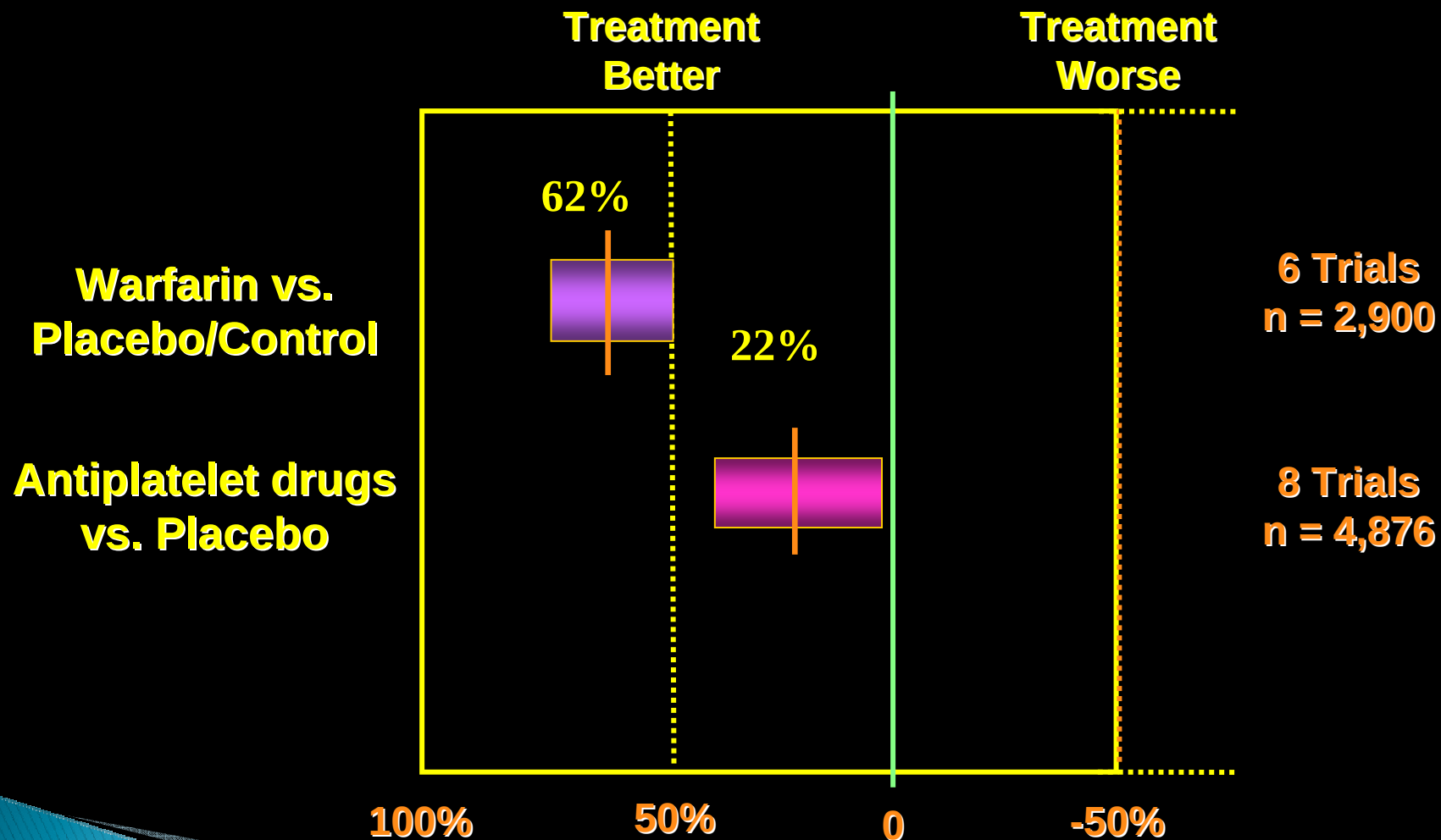
Model 1: adjusted for age, sex, and education level.

Model 2: adjusted for age, sex, education level, depressive symptoms, hypertension, myocardial infarction, diabetes mellitus, heart failure, smoking, body mass index, alcohol consumption, hypercholesterolemia, and cerebral infarcts on MRI.

(Stroke. 2013;44:1020-1025.)

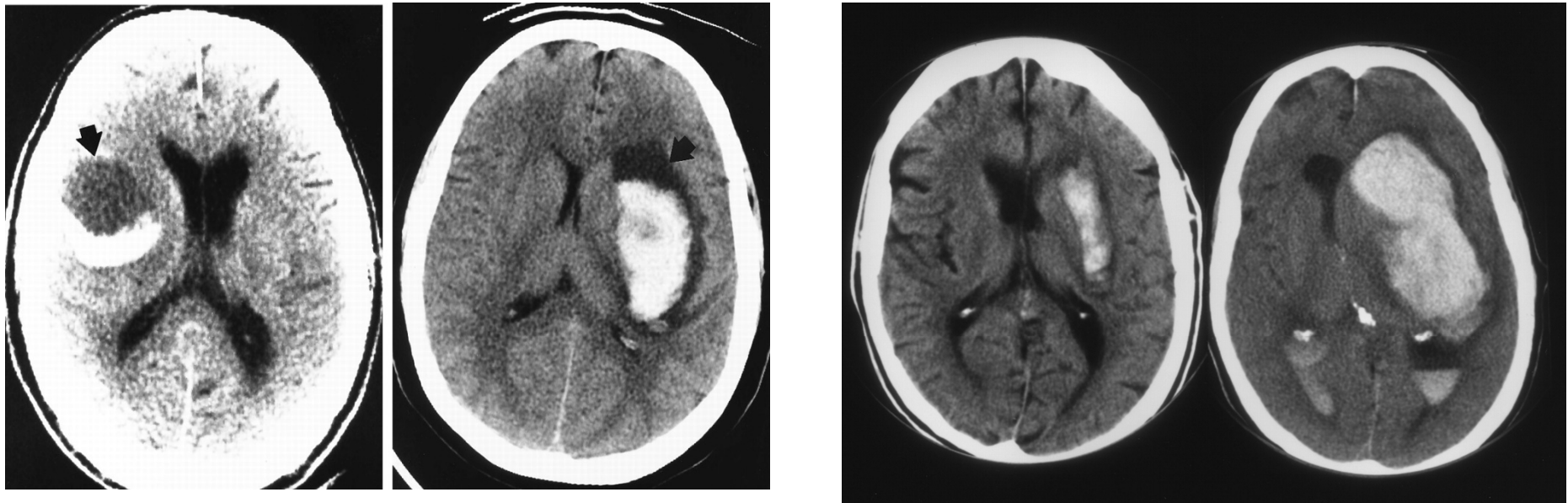
# Antithrombotic Therapy for Atrial Fibrillation

## *Stroke Risk Reduction*



Hart R, et al. *Ann Intern Med* 2007;146:857.

# RADIOLOGICAL FEATURES OF OAC HEMORRHAGE



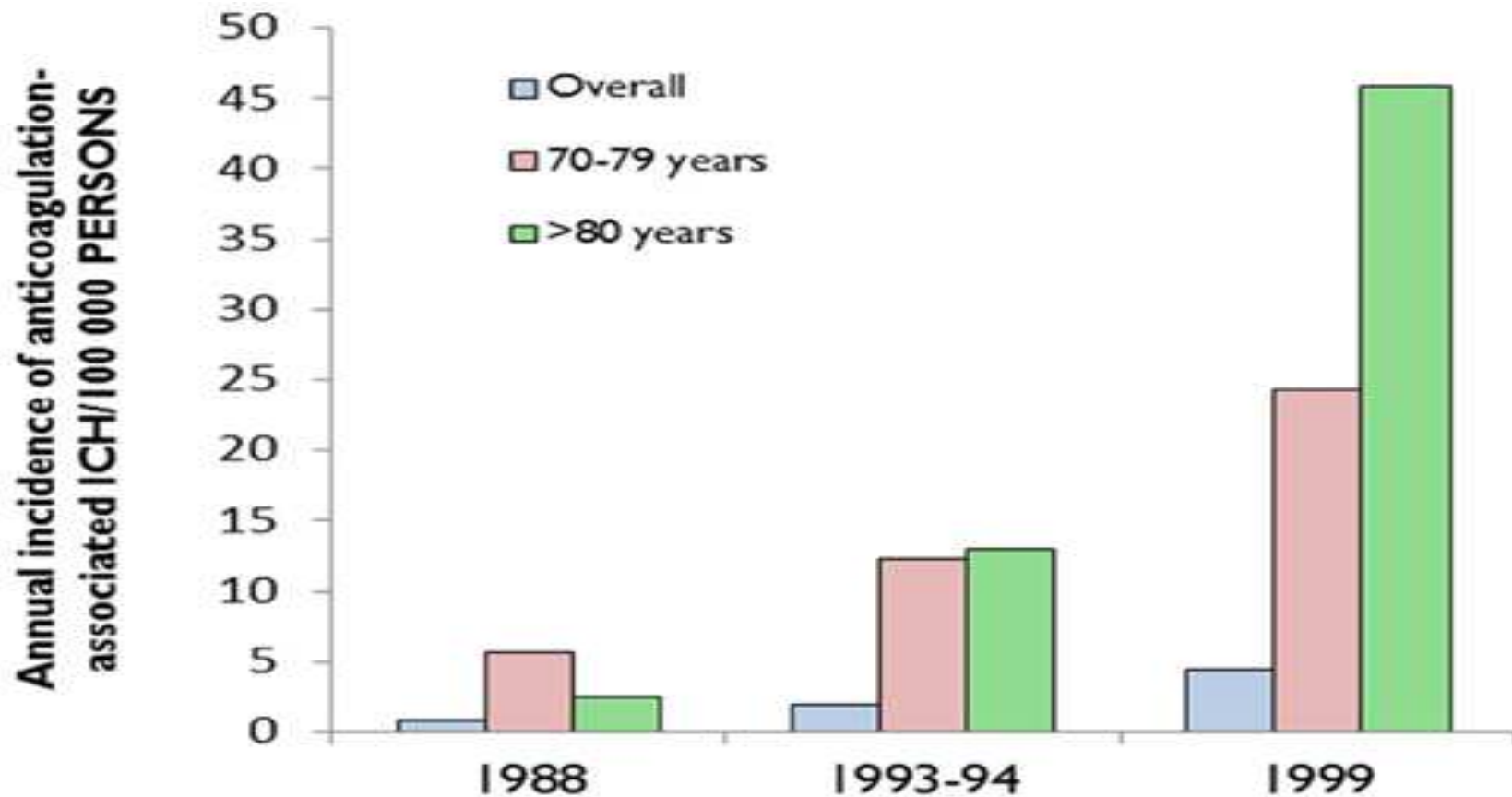
- On CT-scan Blood-Fluid Level
- Higher Risk of Hematoma Expansion

## **DIMENSIONI DEL PROBLEMA**

- 1-1.7% della popolazione europea è in TAO
- 5-15% di tutte le emorragie cerebrali sono correlate alla TAO
- Incidenza di emorragia cerebrale nei pazienti in TAO è 0.3-0.46%
- Nella popolazione l'incidenza di emorragia cerebrale TAO-correlata è 2-9 per 100.000/anno
- Incidenza di emorragia cerebrale e' da 7 a 11 volte maggiore nei pazienti trattati rispetto ai non trattati
- Mortalità 50%

R Hart 1995 Stroke 1995; M. Flaherty et al Neurology 2007;  
T Steiner et al Stroke 2006; AMR Schols et al Stroke 2014

# PROGRESSIVO INCREMENTO DELL'INCIDENZA DELLE EMORRAGIE ASSOCIATE A TAO, SOPRATTUTTO NEGLI ANZIANI



[Flaherty et al., 2007](#)



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**Table 1** Predisposing factors of cerebral hemorrhage in anticoagulated patients [3, 5]

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Proven risk factors

- Advancing age (especially >75 years)
- Hypertension (especially systolic blood pressure >160 mm Hg)
- History of cerebrovascular disease
- Intensity of anticoagulation (mainly if INR > 4.0)

Possible risk factors

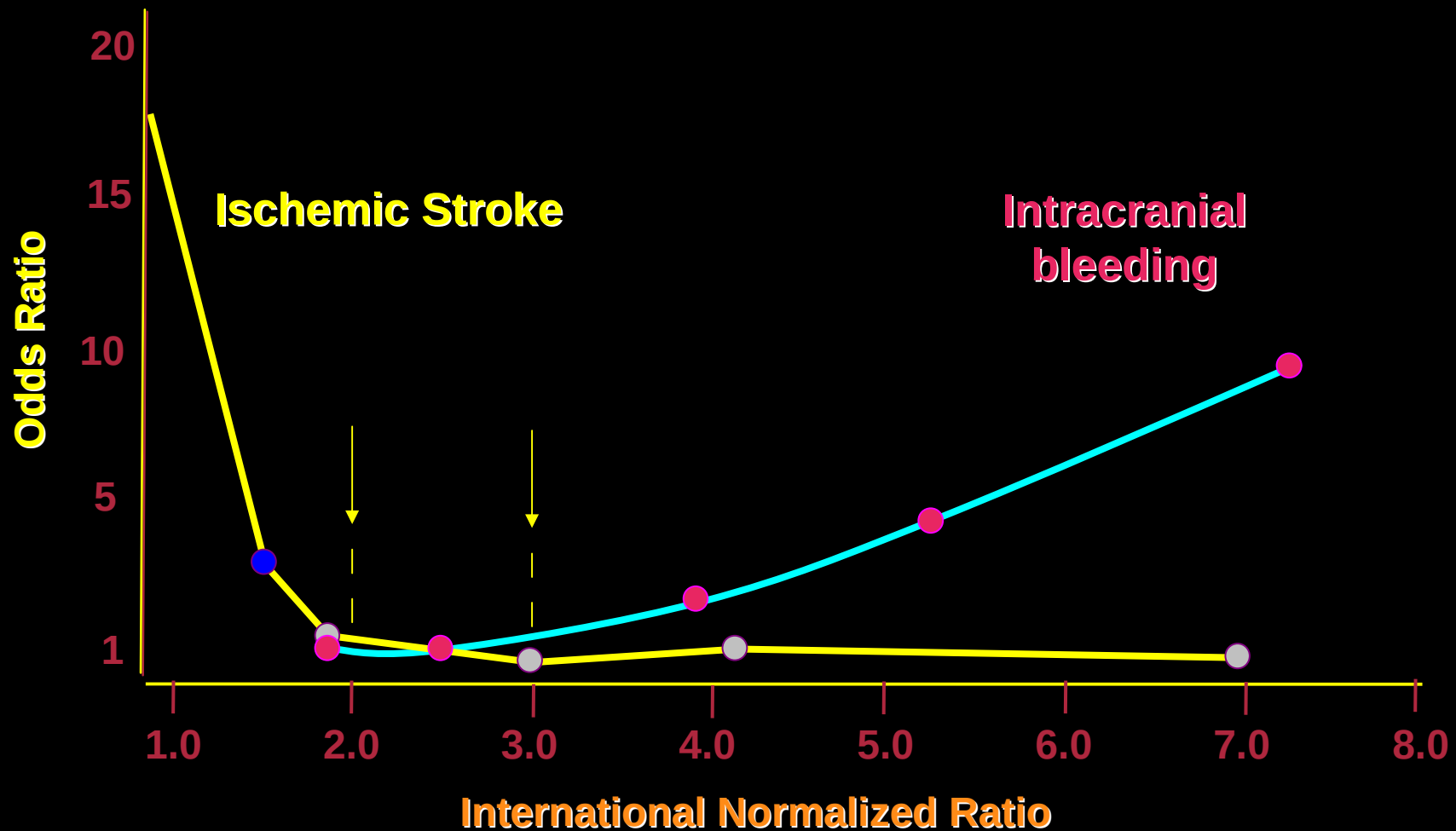
- Increased variation of INR
- Concomitant use of aspirin
- Cerebral amyloid angiopathy
- Tobacco smoking
- Heavy alcohol consumption
- Diabetes
- Serious heart disease
- Liver disease
- Malignancy

Imaging and genetic markers

- Leukoaraiosis detected by brain CT/MRI
  - Microbleeds by T2\*-weighted MRI
  - Apo  $\epsilon$ 2 or  $\epsilon$ 4 genotype
-

# INR at the Time of Stroke or Bleeding

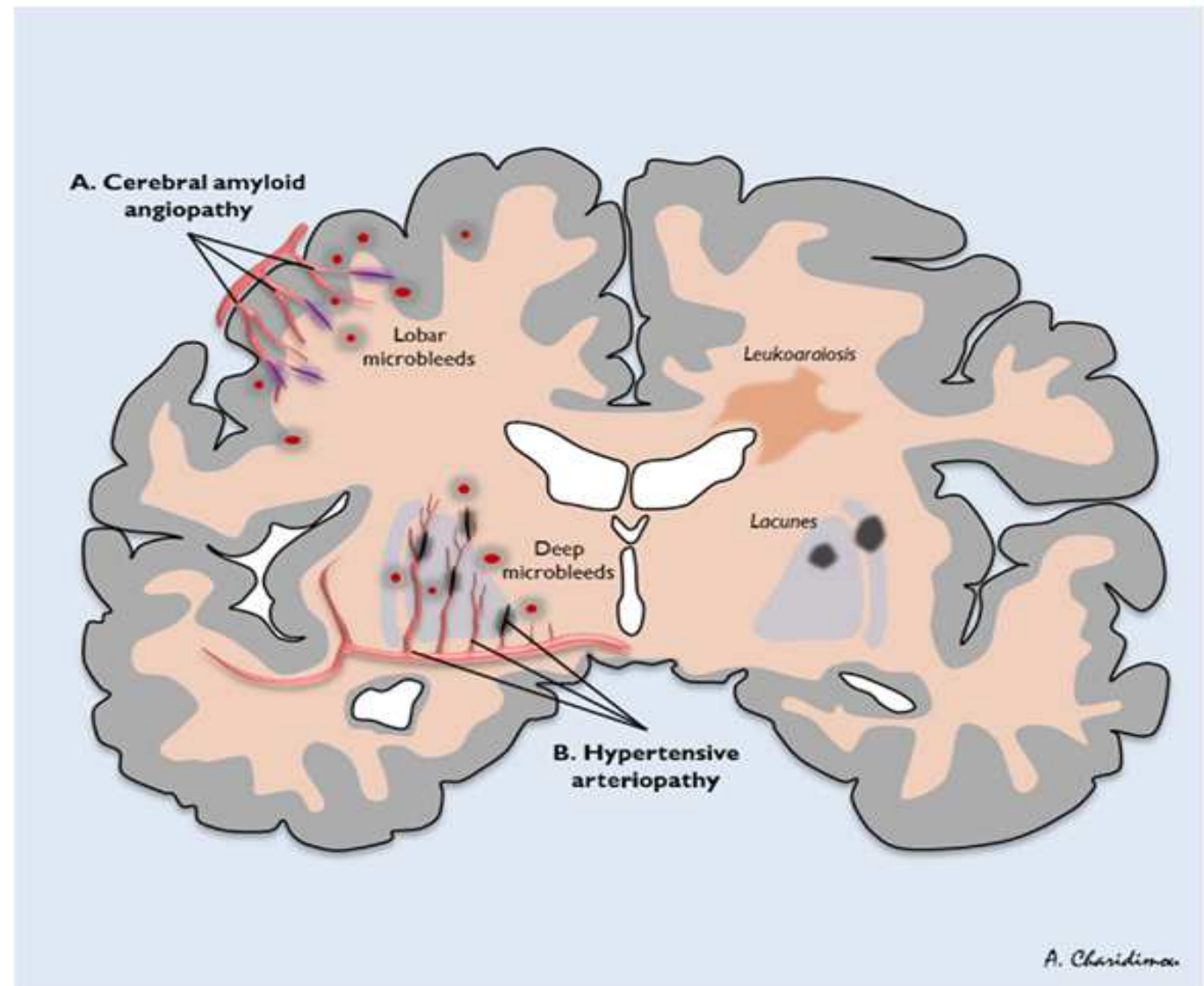
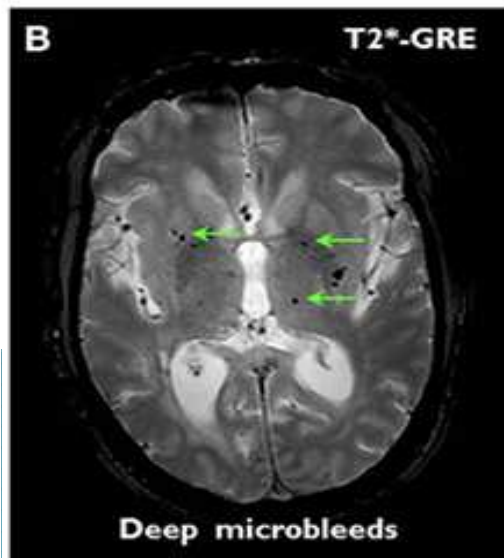
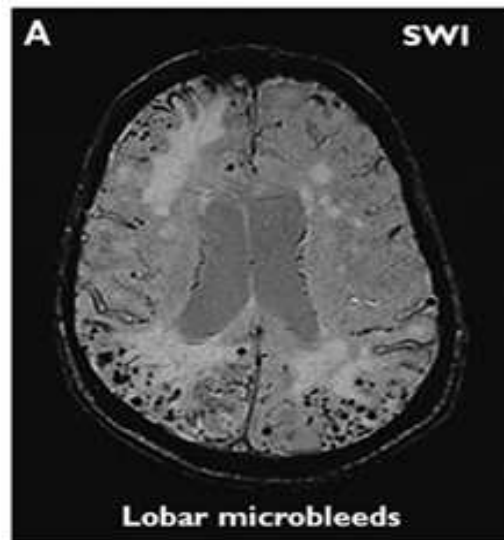
## *Efficacy and Safety of Warfarin*



Fang MC, et al. *Ann Intern Med* 2004; 141:745.  
Hylek EM, et al. *N Engl J Med* 1996; 335:540.

# MARKERS NEURORADIOLOGICI E RISCHIO EMORRAGICO

## "MICROBLEED" E LEUCOARAIOSI



# Use of the CHA<sub>2</sub>DS<sub>2</sub>-VASc and HAS-BLED Scores to Aid Decision Making for Thromboprophylaxis in Nonvalvular Atrial Fibrillation

**Table 2. Assessment of Stroke (CHA<sub>2</sub>DS<sub>2</sub>-VASc)<sup>14</sup> and Bleeding Risk (HAS-BLED)<sup>15</sup> in Atrial Fibrillation Patients**

| CHA <sub>2</sub> DS <sub>2</sub> -VASc             | Score | HAS-BLED                                          | Score  |
|----------------------------------------------------|-------|---------------------------------------------------|--------|
| Congestive heart failure                           | 1     | Hypertension (systolic blood pressure >160 mm Hg) | 1      |
| Hypertension                                       | 1     | Abnormal renal and liver function* (1 point each) | 1 or 2 |
| Age ≥75 y                                          | 2     | Stroke                                            | 1      |
| Diabetes mellitus                                  | 1     | Bleeding tendency/predisposition*                 | 1      |
| Stroke/TIA/TE                                      | 2     | Labile INRs (if on warfarin)*                     | 1      |
| Vascular disease (prior MI, PAD, or aortic plaque) | 1     | Elderly (eg, age >65 y)                           | 1      |
| Aged 65 to 74 y                                    | 1     | Drugs or alcohol (1 point each)*                  | 1 or 2 |
| Sex category (ie, female sex)                      | 1     |                                                   |        |
| Maximum score                                      | 9     | Maximum score                                     | 9      |

TABLE 2. Expert Recommendations for the Management of Warfarin-Associated ICH: Responses to 3 Specific Questions\*

| Question                                                                  | Expert 1                           | Expert 2                                             | Expert 3 <sup>†</sup>      | Expert 4 <sup>†</sup>             | Expert 5 <sup>†</sup>         | Expert 6                      | Expert 7           |
|---------------------------------------------------------------------------|------------------------------------|------------------------------------------------------|----------------------------|-----------------------------------|-------------------------------|-------------------------------|--------------------|
| 1.How can anticoagulation best be reversed (INR=2.5)?                     | rFVIIa and vitamin K               | rFVIIa if deteriorating; FFP and vitamin K otherwise | rFVIIa, FFP, and vitamin K | FFP or PCCs and vitamin K         | PCCs and vitamin K            | PCCs and vitamin K            | PCCs and vitamin K |
| 2.When should anticoagulation with prosthetic cardiac valve be restarted? | >7 d in most patients <sup>‡</sup> | >7 d if computed tomogram stable                     | 5-10 d                     | Low-dose heparin as early as 48 h | 10-14 d <sup>§</sup>          | 1-3 d                         | 7 d                |
| 3.Should warfarin therapy be restarted for atrial fibrillation?           | With reluctance                    | If prior ischemic stroke                             | Probably never             | If ICH is deep (ie, nonlobar)     | If ICH is deep (ie, nonlobar) | If ICH is deep (ie, nonlobar) | In secondary       |

## REVIEW

# Pharmacologic and Nonpharmacologic Therapies for Stroke Prevention in Nonvalvular Atrial Fibrillation

JIANGLI HAN, M.D., PH.D.,\*,† JIE CHENG, M.D., PH.D.,\* and NILESH MATHURIA, M.D.\*

From the \*Texas Heart Institute/St.Luke's Episcopal Hospital, Houston, Texas; and †Peking University Third Hospital, Beijing, China

(PACE 2012; 35:887–896)

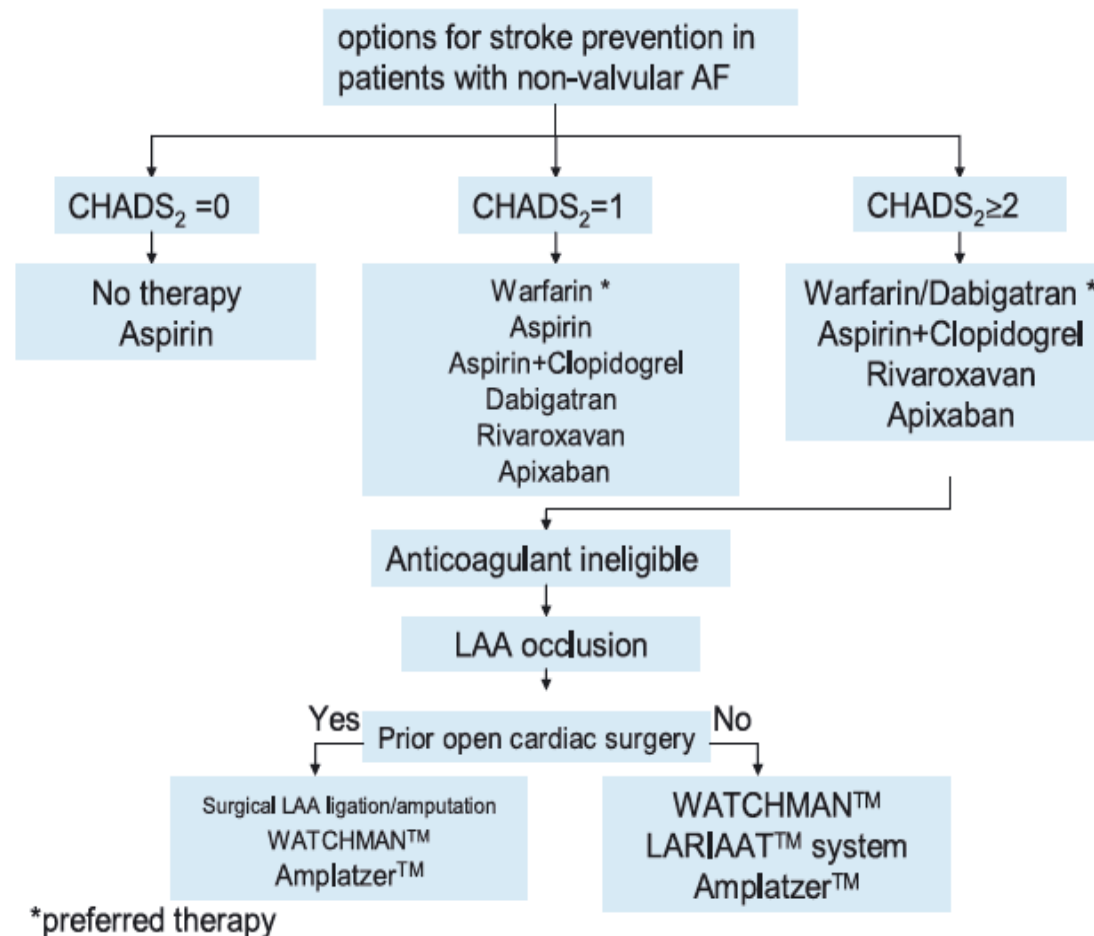
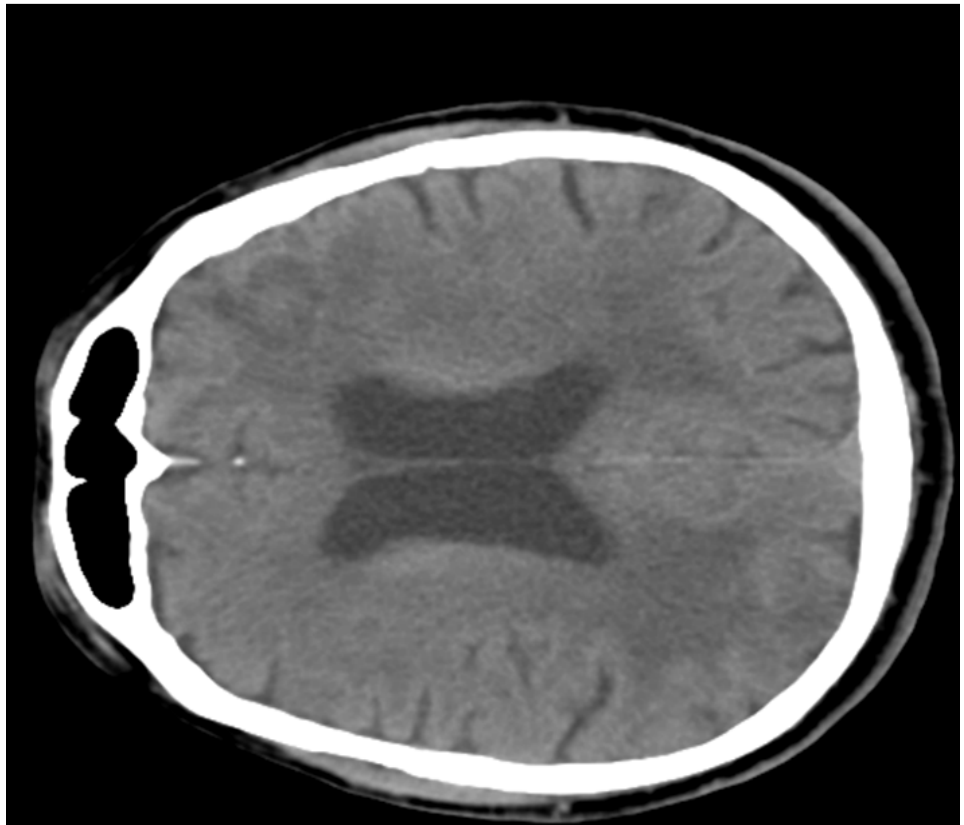
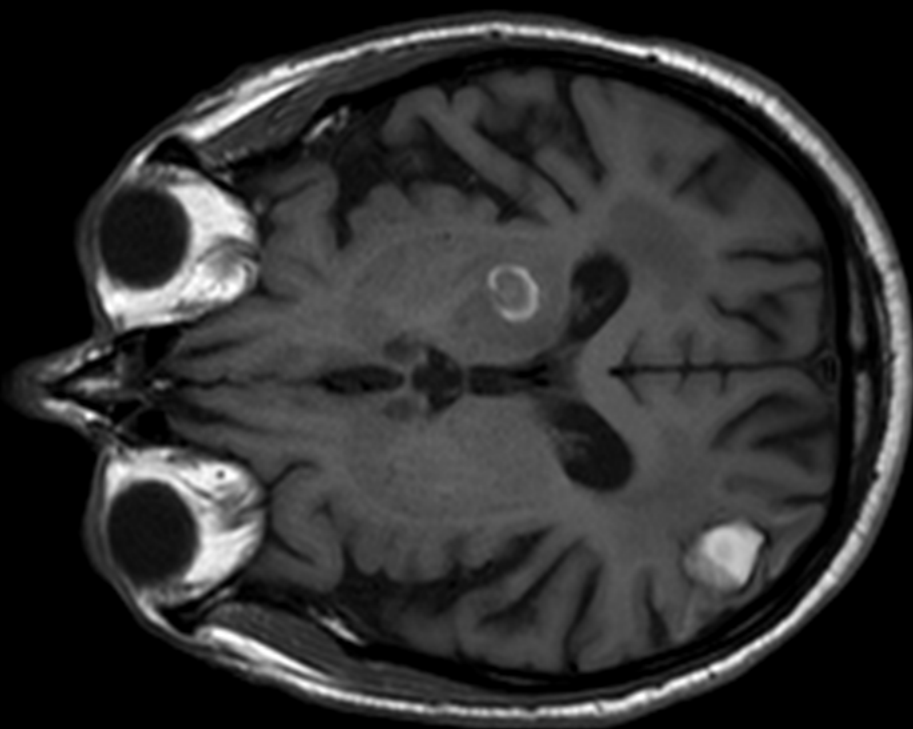
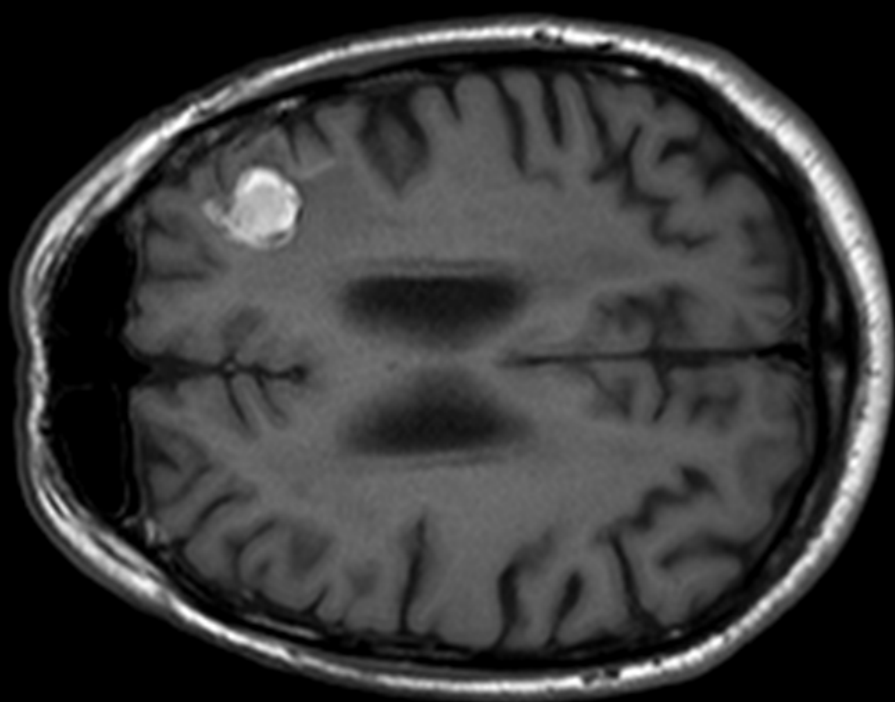
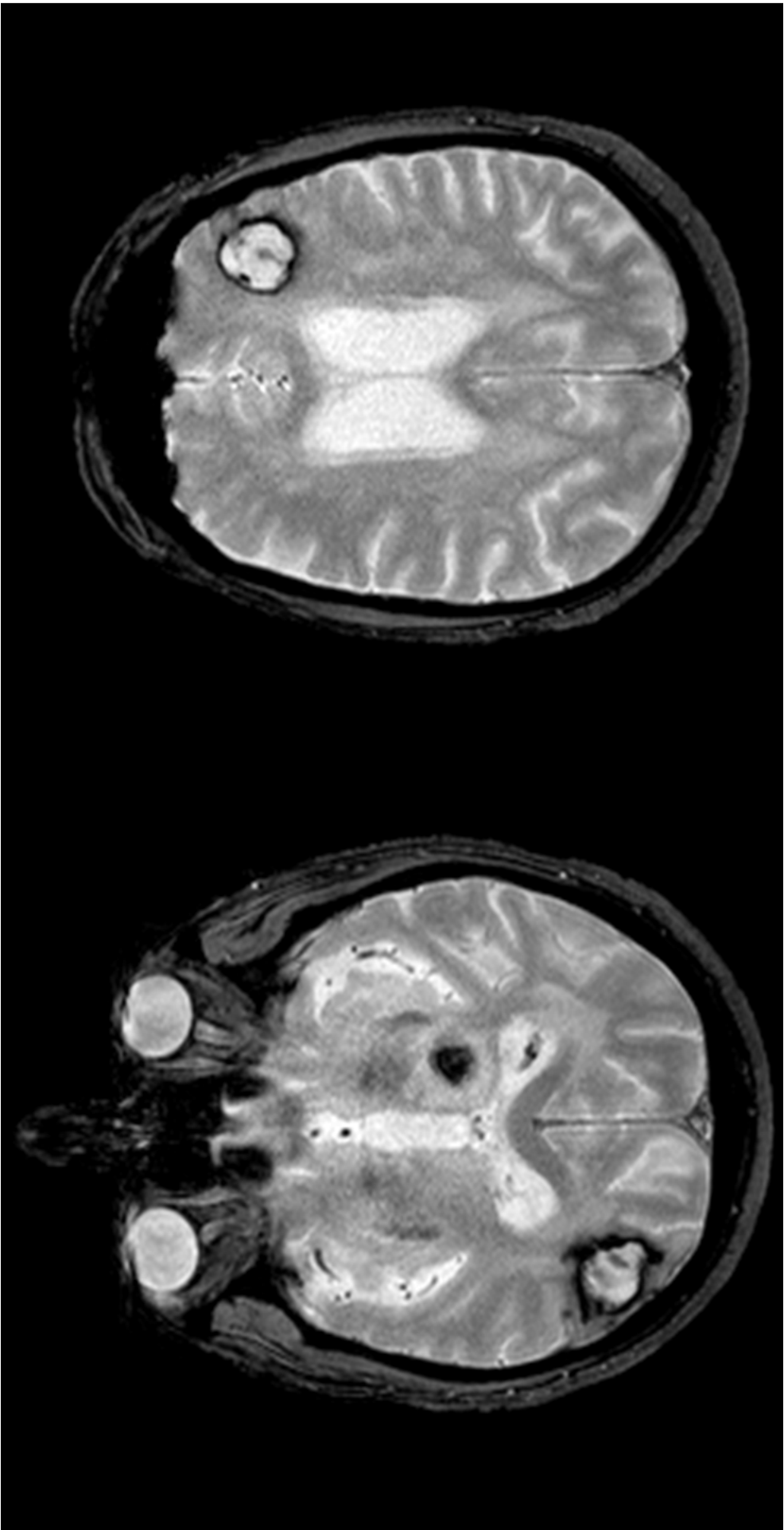


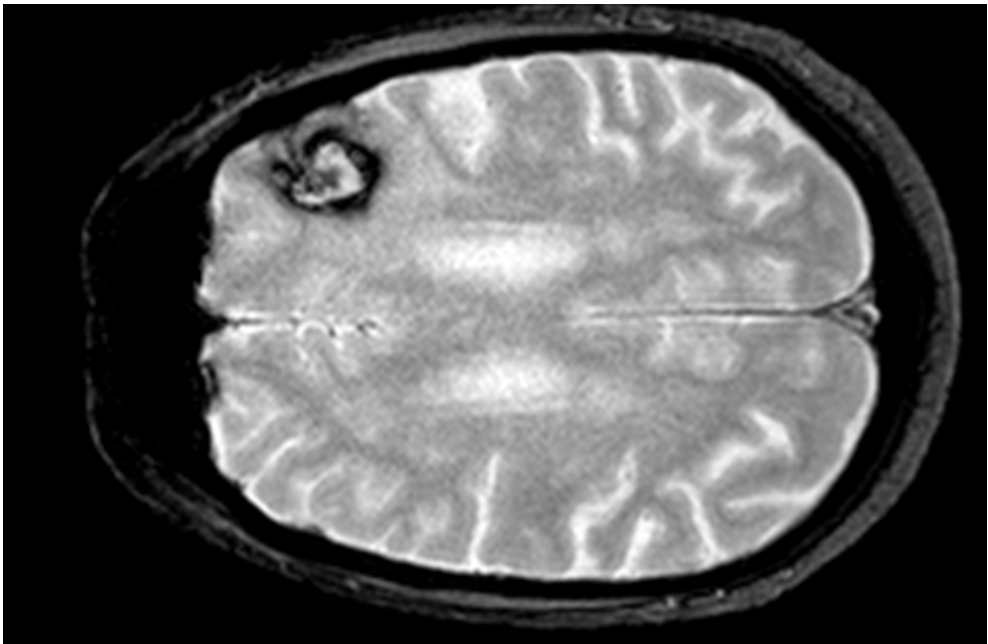
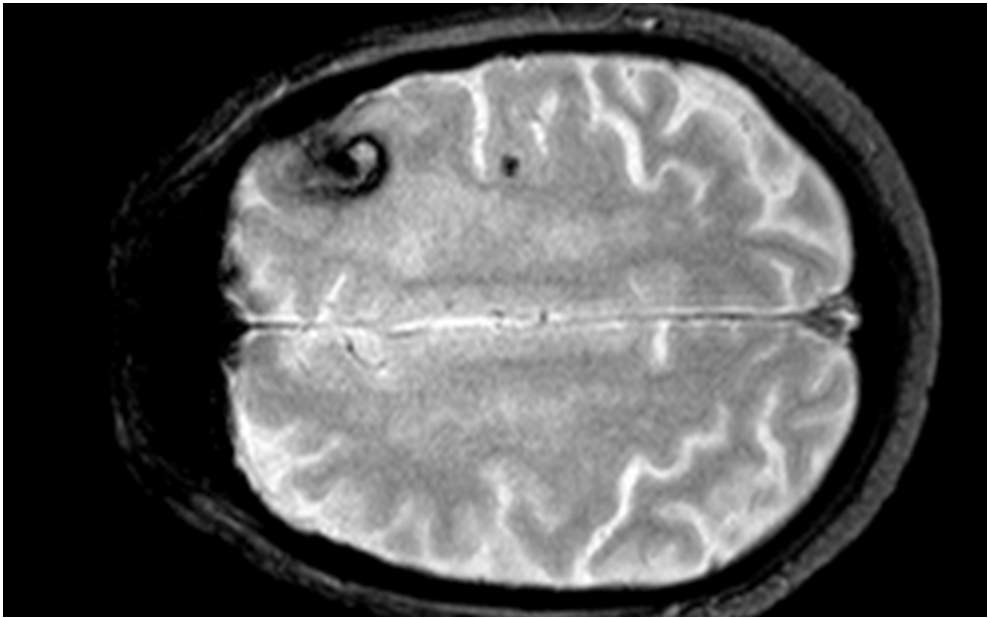
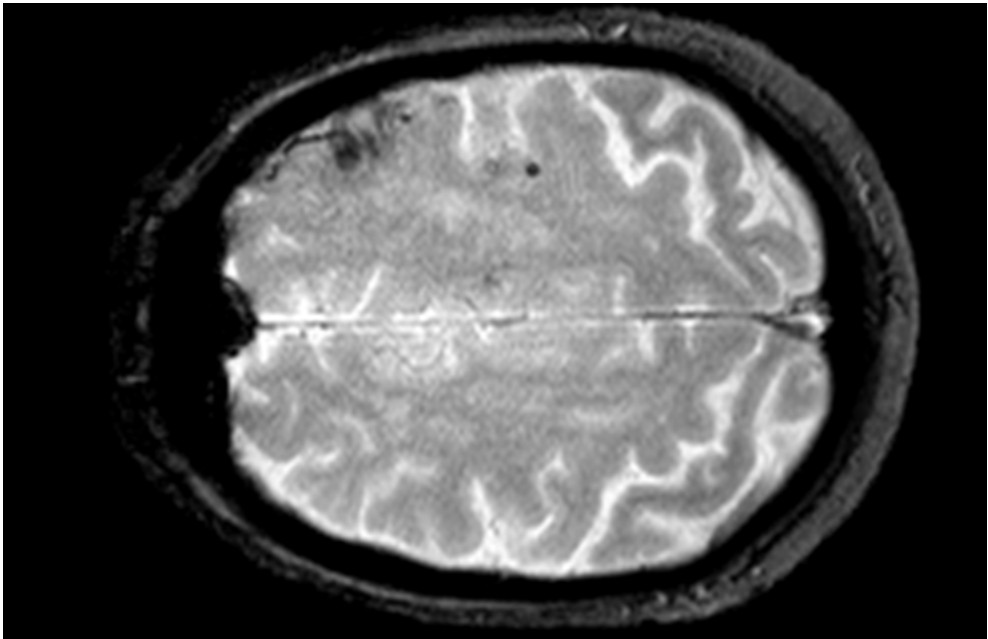
Figure 3. Potential algorithm regarding options for stroke prevention in nonvalvular AF.











**GRAZIE PER L'ATTENZIONE**