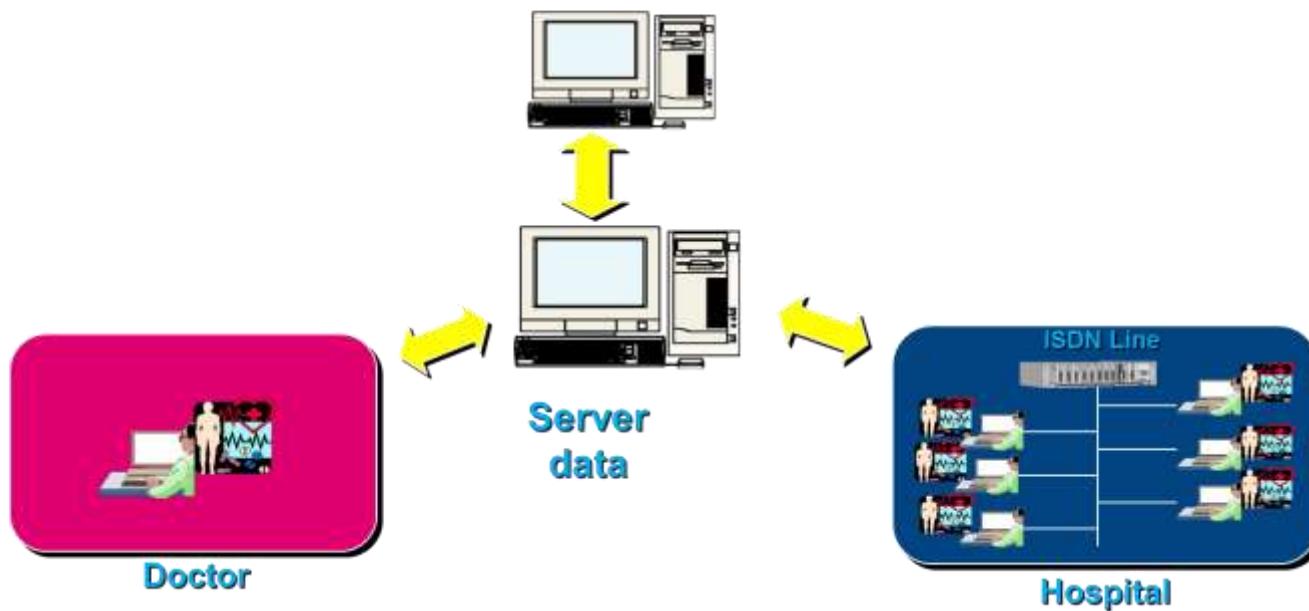


# Il Network dell'Ipertensione

**WWW.CAMPANIASALUTE.COM**



# Campania Salute Network

www.campaniasalute.com

Versione Italiana English version

## Campania Salute Project

WEB-based connection between Specialties Centers and General Practitioners for Clinical Data-Base consultations of High Cardiovascular Risk patient.

Telework, Clinical and Diagnostic evaluations Reservations & Clinical Referrals Consultations

General Practitioner's Program

CODICE PERSONALE  
PERSONAL CODE

EMERGENCY  
ACCESS

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Scientific Consultant  
Prof. Giovanni de Simone  
Professor of Medicine

AIFA Project

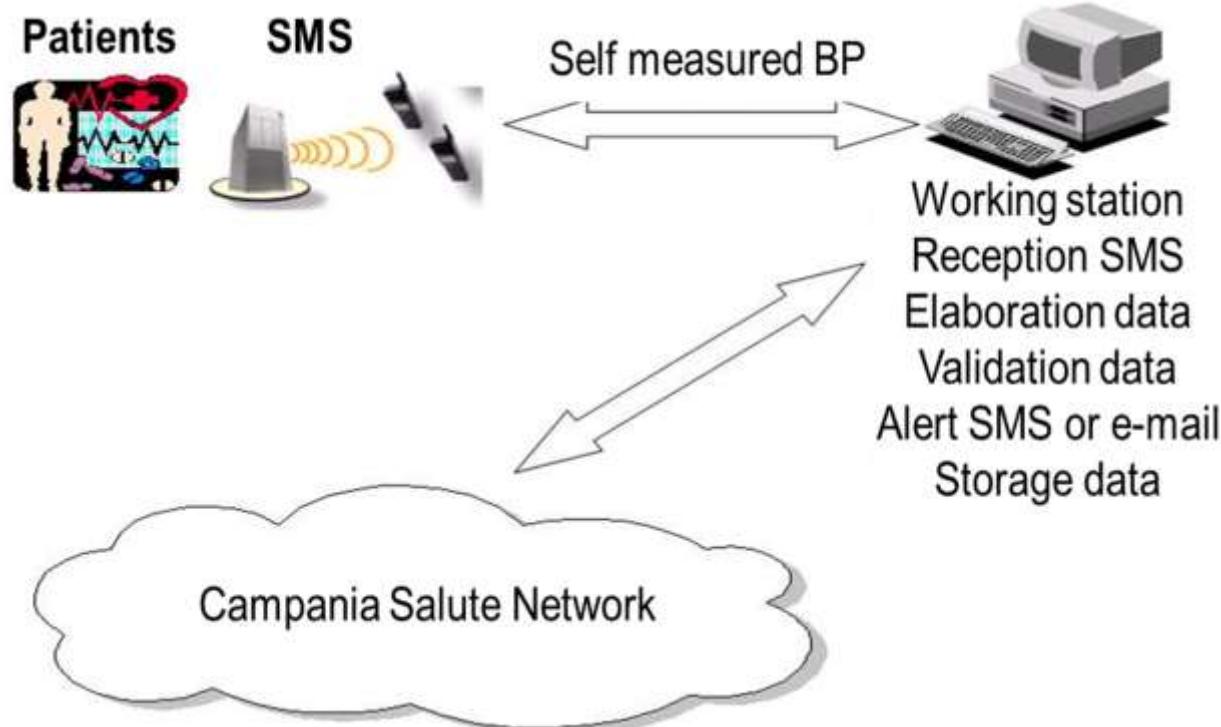
Tasca Project

Telematic ECG Referrals

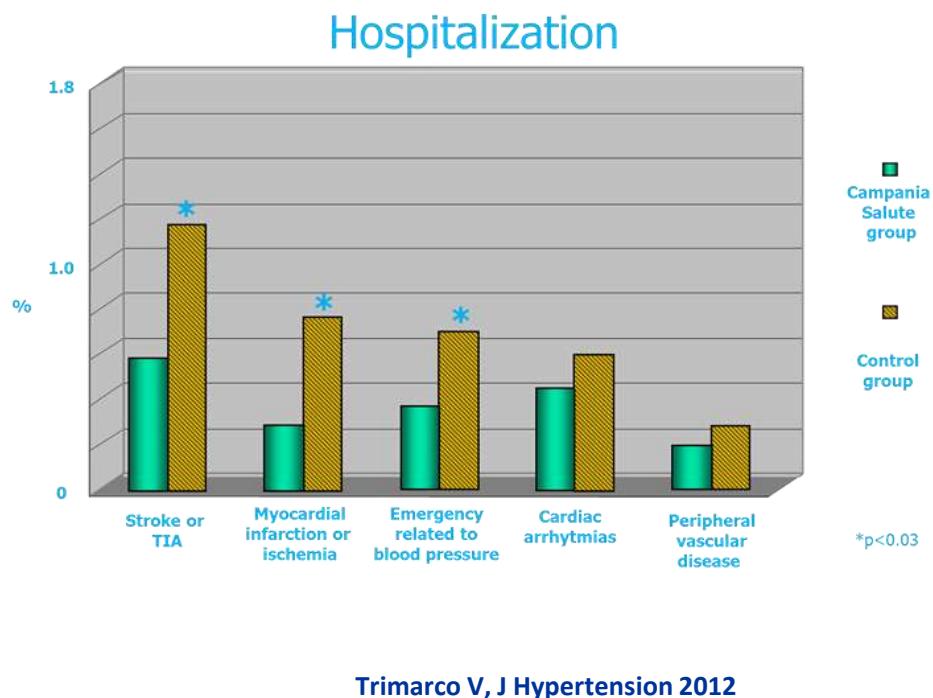
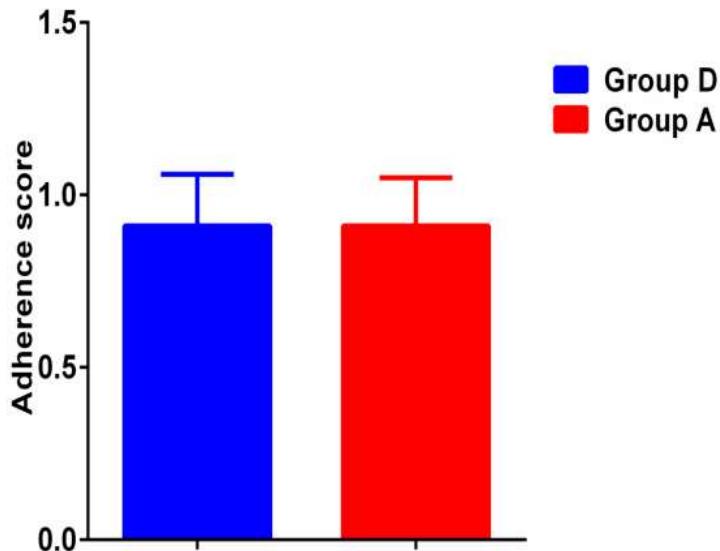
Share Project

The screenshot shows the homepage of the Campania Salute Network website. At the top, there is a header with the URL 'www.campaniasalute.com/index\_english.html' and a search bar. Below the header, there are language links for 'Versione Italiana' and 'English version', and a logo of the University of Naples 'Federico II'. The main content area features a title 'Campania Salute Project' and a subtitle 'WEB-based connection between Specialties Centers and General Practitioners for Clinical Data-Base consultations of High Cardiovascular Risk patient.' Below this, there is a section titled 'Telework, Clinical and Diagnostic evaluations Reservations & Clinical Referrals Consultations' and 'General Practitioner's Program'. To the left, a vertical menu lists various project components: Introduction, Participants, Goals, Description, Structure, Pilot Experience, Results, Services, Slides, Steering Committee, and Proposal Studies. On the right, there is a sidebar titled 'EMERGENCY ACCESS' containing a form for entering a 'CODICE PERSONALE' (Personal Code). The sidebar also includes a note about inserting the personal code onto the back of a card and a QR code at the bottom.

# CSN

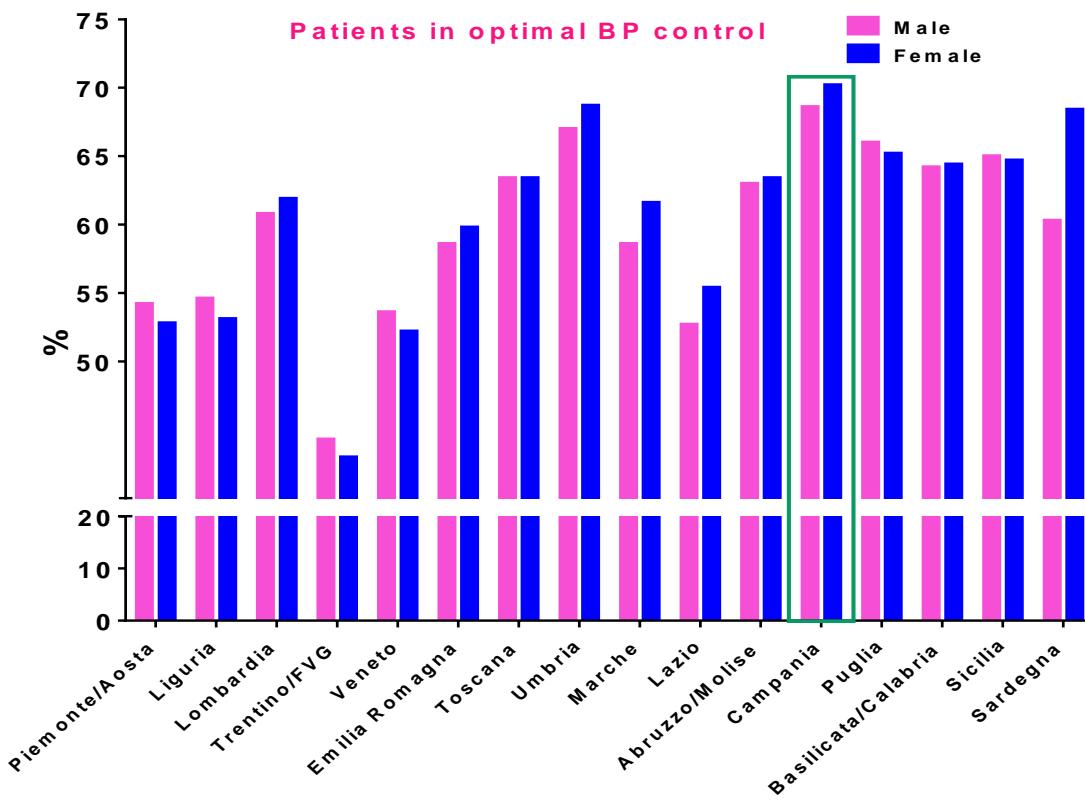


# Persistenza ed Aderenza alla Terapia Antiipertensiva



Trimarco V, J Hypertension 2012

# Controllo della Pressione Arteriosa in Italia



WINCARE - Popolazione Selezionata: GLOBALE

Operazioni generali | Paziente | Servizi | Funzioni di Gestione | ?

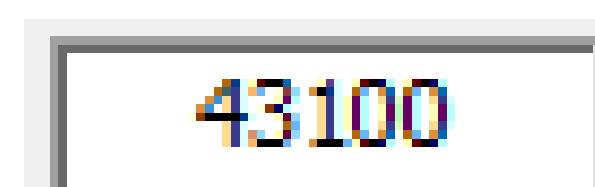
Eseguire Nuovo Vista Eventi Smart Card Duplicazione Utility About Progetto AITA Analitica Centrale

Ricerca per Cognome Condizione marito

Cerca Avanzata Aggiorna

A/S	Centro	Codice	Cognome	Nome	Data di nascita	Ingresso in	Score
A	1	7809	MARINO	Giovanna	02/09/1949	Iperpressione	
A	1	107383	MARINO	Giovanni	28/07/1943	Esterno	
A	1	102709	MARINO	Giulia	17/11/1937	Cardiologia	
S	1	4170	MARINO	Giuliana	16/10/1954	Iperpressione	
A	1	10571	MARINO	Giuliana	12/01/1951	Normale	
A	100	837	MARINO	GIUSEPPE	18/01/1954	Cardiopatia Ischemica	
A	1	105206	MARINO	Giuseppe	01/04/1963	Esterno	
A	1	106315	MARINO	Ida	22/07/1959	Esterno	
A	100	1057	MARINO	Isabella	09/03/1928	Cardiopatia Ischemica	
A	1	7373	MARINO	Italia	07/05/1934	Iperpressione	
A	1	3842	MARINO	Liliana	15/10/1942	Iperpressione	
A	1	39546	MARINO	Lorenzo	16/12/1952	Iperpressione	
A	100	634	MARINO	Lorenzo	05/04/1937	Cardiopatia Ischemica	
A	1	104875	MARINO	Luisa	02/12/1964	Iperpressione	
A	1	54249	MARINO	Luigi	13/09/1955	UTSC	
A	1	51370	MARINO	Luigi	01/04/1950	UTSC	
A	1	42473	MARINO	Maria	18/02/1939	Iperpressione	
A	1	10564	MARINO	Maria	05/06/1931	UTSC	
A	1	34650	MARINO	Maria	10/10/1950	Iperpressione	
S	1	31971	MARINO	Maria Luisa	19/07/1948	Iperpressione	
A	1	38000	MARINO	Maria Rosaria	09/12/1954	Iperpressione	
A	1	37188	MARINO	Maria Rosaria	13/01/1946	DH	
A	1	8197	MARINO	Maria Rosaria	14/09/1939	Iperpressione	
A	1	47983	MARINO	Maria Rosaria	10/05/1950	Iperpressione	
A	1	50432	MARINO	Maria Rosaria	24/09/1942	Iperpressione	
A	1	58720	MARINO	Massimiliano	26/01/1966	Iperpressione	
A	1	106329	MARINO	Massimo	13/08/1963	UTSC	
A	1	50069	MARINO	Mattia	23/05/2000	Esterno	
A	1	44709	MARINO	Maurizio	03/11/1974	Esterno	
A	1	104124	MARINO	Michèle	30/09/1956	UTSC	
A	1	56255	MARINO	Nada	12/05/1952	Neurologia	
A	1	57510	MARINO	Natalino	07/07/1951	AOU GASTRODENT	
A	1	44265	MARINO	Nicola	13/10/1963	Esterno	
A	1	41369	MARINO	Paolo	10/12/1993	Iperpressione	
A	1	50055	MARINO	Paolo	02/09/1951	Cardiologia	
A	1	31516	MARINO	Paolo	19/01/1949	Iperpressione	
A	1	31520	MARINO	Pasquale	27/02/1972	Esterno	
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A	1	40133	MARINO	Patrizia	21/11/1950	Esterno	
A	1	44920	MARINO	Patrizia	14/02/1970	Iperpressione	
A	1	44804	MARINO	Pietro	06/05/1956	Iperpressione	
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A	1	41248	MARINO	Rachele	08/11/1952	Esterno	
A	1	104051	MARINO	Raffaele	02/08/1976	Esterno	
A	1	53125	MARINO	Renata	24/10/1950	UTSC	
A	1	39540	MARINO	Rita	18/05/1963	Esterno	
A	1	105715	MARINO	Roberta	03/08/1968	Esterno	
S	1	31016	MARINO	Roberto	16/11/1952	Iperpressione	
A	1	35292	MARINO	Rocco	08/04/1969	DH	
A	1	57427	MARINO	Rosa	12/04/1939	Esterno	
A	1	40196	MARINO	Rosalba	02/04/1956	DH	
A	1	33008	MARINO	Rosaria	04/02/1944	Esterno	
A	1	54029	MARINO	Rubria	26/03/1945	Iperpressione	
A	1	39842	MARINO	Salvatore	05/07/1938	Iperpressione	
A	1	51340	MARINO	Sebastiano	14/03/1985	Iperpressione	
A	1	106439	MARINO	Serafina	12/03/1955	Iperpressione	
A	1	51287	MARINO	Sergio	17/02/1975	Iperpressione	
A	1	30175	MARINO	Sergio	15/01/1959	Iperpressione	
A	1	46394	MARINO	Stefania	25/08/1969	Esterno	
A	1	102916	MARINO	Tullio	26/04/1961	Esterno	
A	1	58732	MARINO	Vincenzo	20/01/1958	Iperpressione	

Patenti nella popolazione: 43100



Dr. Raffaele Izzo OO, MM, n°. CEB 16/02/2017 11:51



Ricerca per Ingresso In Condizione



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A	1	3521	ABAGNALE	Giuseppe	22/02/1966	Iper tensione	
A	1	4429	ABALSIANO	Antonio	17/02/1955	Iper tensione	
A	1	8334	ABATE	Antroglia	06/03/1955	Iper tensione	
A	1	2413	ABATE	Anna	07/03/1949	Iper tensione	
A	1	6574	ABATE	Carmina	19/10/1951	Iper tensione	
A	1	33054	ABATE	Carmine	27/01/1949	Iper tensione	
A	1	44808	ABATE	Ciro	29/07/1965	Iper tensione	
A	1	40397	ABATE	Domenico	15/12/1978	Iper tensione	
A	1	57871	ABATE	Gennaro	10/07/1963	Iper tensione	
A	1	43196	ABATE	Giuseppe	22/12/1941	Iper tensione	
A	1	6007	ABATE	Giuseppe	21/06/1954	Iper tensione	
A	1	811	ABATE	Giuseppe	02/04/1961	Iper tensione	
A	1	5630	ABATE	Lorenzo	02/07/1949	Iper tensione	
A	1	1179	ABATE	Luigi	19/03/1975	Iper tensione	
A	1	2678	ABATE	Pasquale	11/02/1954	Iper tensione	
A	1	7907	ABATE	Quirino	04/06/1951	Iper tensione	
A	1	35510	ABATE	Rosal	06/09/1942	Iper tensione	
A	1	911	ABATE	Saverio	25/06/1962	Iper tensione	
A	1	9870	ABATE	Vincenzo	06/02/1959	Iper tensione	
A	1	104644	ABATECOLA	Adriana	31/07/1946	Iper tensione	
A	1	7502	ABATEGIOVANNI	Giovanna	11/10/1935	Iper tensione	
A	1	37212	ABAZIA	Elisabetta	23/03/1968	Iper tensione	
A	1	34273	ABAGNALE	Umberto	08/05/1954	Iper tensione	
A	1	6405	ABAGNAO	Antonio	11/06/1946	Iper tensione	
A	1	50108	ABAGNAO	Liliana	11/08/1945	Iper tensione	
A	1	102278	ABBAGNATO	Carmina	22/05/1952	Iper tensione	
A	1	33988	ABBAMONDI	Dante	09/08/1957	Iper tensione	
A	1	3906	ABBADE	Bruno	26/11/1953	Iper tensione	
A	1	903	ABBADE	Carmina	16/07/1943	Iper tensione	
A	1	6209	ABBADE	Ciro	04/07/1945	Iper tensione	
A	1	42205	ABBADE	Concetta	14/11/1962	Iper tensione	
A	1	53520	ABBADE	Genaro	14/04/1966	Iper tensione	
A	1	6197	ABBADE	Giuseppe	17/09/1950	Iper tensione	
A	1	8137	ABBADE	Lorenzo	19/04/1971	Iper tensione	
A	1	34750	ABBADE	Maria	16/04/1949	Iper tensione	
A	1	6173	ABBADE	Mario	17/09/1956	Iper tensione	
A	1	46372	ABBADE	Renato	01/06/1939	Iper tensione	
A	1	33449	ABBADE	Salvatore	12/04/1955	Iper tensione	
A	1	30942	ABBATI	Antonio	07/09/1944	Iper tensione	
A	1	34091	ABBENANTE	Anne	01/07/1945	Iper tensione	
A	1	49701	ABBENANTE	Umberto	16/11/1951	Iper tensione	
A	1	5460	ABBUSO	Primilla	29/05/1965	Iper tensione	
A	1	42592	ABBONDANTE	Vincenzo	23/01/1947	Iper tensione	
A	1	2874	ABRUZZESE	Francesca	20/07/1957	Iper tensione	
A	1	46663	ABELLE	Amalia	27/08/1949	Iper tensione	
A	1	9136	ABELANDI	Angelina	13/09/1938	Iper tensione	
A	1	33108	ABRAMO	Liberia	01/08/1962	Iper tensione	
A	1	50358	ABRATE	Massimo	14/01/1966	Iper tensione	
A	1	6685	ABRUZZESE	Carmino	08/09/1952	Iper tensione	
A	1	8041	ABRUZZESE	Giovanna	10/04/1951	Iper tensione	
A	1	7885	ABRUZZESE	Ituriala	01/06/1954	Iper tensione	
A	1	44740	ACAMPORA	Sergio	09/01/1966	Iper tensione	
A	1	48223	ACAMPORA	Wanda	19/05/1932	Iper tensione	
A	1	39911	ACAMPORA	Angelina	14/05/1949	Iper tensione	
A	1	1196	ACAMPORA	Anna	09/02/1948	Iper tensione	
A	1	39910	ACAMPORA	Anna Maria	06/05/1947	Iper tensione	
A	1	127	ACAMPORA	Emanuele	28/06/1973	Iper tensione	
A	1	3635	ACAMPORA	Raimondo	11/02/1948	Iper tensione	
A	1	7266	ACAMPORA	Annunzieta	25/06/1946	Iper tensione	
A	1	41218	ACAMPORA	Bernardo	21/01/1943	Iper tensione	
A	1	9939	ACAMPORA	Carmina	04/01/1956	Iper tensione	

Papienti nella popolazione:

15990

15:59:00

Query n.1 di 1

Dr. Raffaele Izzo OO: MM: yy: CC: 16/02/2017 11:58



Ricerca per Ingresso In Condizione Cerca Avanzata Aggiorna

A/S	Centro	Codice	Cognome	Nome	Data di nascita	Impronta In	Score
A	I	58509	ABAGNALE	Roberto	16/09/1963	UTIC	
A	I	31209	ABATE	Andrea	16/10/1958	UTIC	
A	I	58993	ABATE	Bruno	13/10/1967	UTIC	
A	I	36403	ABATE	Carmela	27/01/1925	UTIC	
A	I	44511	ABATE	Nicola	09/03/1953	UTIC	
A	I	31923	ABATE	Vincenzo	23/09/1961	UTIC	
A	I	107663	ABBAMONDI	Alessandra	30/05/1994	UTIC	
A	I	42334	ABBASTE	Brigida	11/04/1940	UTIC	
A	I	42818	ABBATTI	Antonio	14/05/1945	UTIC	
A	I	105964	ABBATINO	Gennaro	12/06/1963	UTIC	
A	I	32478	ABBRECCIA	Lugi	27/03/1949	UTIC	
A	I	38991	ABBRUNZO	Alfonso	05/01/1931	UTIC	
A	I	35170	ABRUZZESE	Renato	31/03/1937	UTIC	
A	I	58579	ABILITATO	Ciro	05/11/1948	UTIC	
A	I	107614	ABISSI	Anna	12/07/1927	UTIC	
A	I	10184	ACAMPORA	Augusto	16/01/1929	UTIC	
A	I	53467	ACAMPORA	Luisa	11/10/1924	UTIC	
A	I	105373	ACAMPORA	Salvatore	06/05/1942	UTIC	
A	I	42766	ACAMPORA	Vincenzo	16/12/1945	UTIC	
A	I	106499	ACANFORA	Gina	15/07/1950	UTIC	
A	I	54475	ACCARDI	Agostino	16/09/1959	UTIC	
A	I	55599	ACCARDO	Anna Rosaria	03/03/1945	UTIC	
A	I	34126	ACCARDO	Salvatore	17/02/1945	UTIC	
A	I	42395	ACCARINO	Vincenzo	11/11/1953	UTIC	
A	I	51093	ACCARINO	Mario	18/08/1930	UTIC	
A	I	49991	ACCETTO	Filomena	24/11/1935	UTIC	
A	I	52102	ACORSO	Giacetano	07/03/1945	UTIC	
A	I	43094	ACERRA	Salvatore	01/12/1927	UTIC	
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A	I	52676	ACINTO	Leonardo	06/03/1937	UTIC	
A	I	37100	ACINTO	Gennaro	20/07/1950	UTIC	
A	I	31571	ACINTO	Pasquale	07/06/1944	UTIC	
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A	I	53903	ADAMI	Enrico	02/06/1963	UTIC	
A	I	42388	ADAMI	Rosario	25/06/1943	UTIC	
A	I	41413	ADAMO	Pasquale	17/01/1946	UTIC	
A	I	10046	ADARDI	Giacomo	11/02/1929	UTIC	
A	I	54837	ADEO	Amorino	03/04/1949	UTIC	
A	I	31450	ADEO	Angela	14/06/1939	UTIC	
A	I	39898	ADEO	Italia	29/05/1934	UTIC	
A	I	33571	ADEO	Santolo	29/06/1972	UTIC	
A	I	44708	ADDEYCO	Antonio	30/11/1946	UTIC	
A	I	10231	ADEL FAUL	Mohammed	14/11/1956	UTIC	
A	I	105339	ADONI	Gennaro	18/09/1957	UTIC	
A	I	48325	ADORNO	Renato	18/04/1949	UTIC	
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A	I	35168	AFFINITO	Bartolomeo	06/06/1940	UTIC	
A	I	108343	AFFINITO	Bento	12/04/1962	UTIC	
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A	I	90618	APRATELLANZA	Rosa	10/08/1935	UTIC	
A	I	43315	AGATA	Silvana	27/11/1949	UTIC	
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A	I	59028	AGNETTI	Paolo	08/01/1990	UTIC	
A	I	31860	AGORINE	Emilio	25/10/1963	UTIC	
A	I	43034	AGOVINO	Giovanni	07/03/1936	UTIC	
A	I	32336	AGRESTI	Aldo Francesco	27/05/1936	UTIC	
A	I	10073	AGRESTI	Vincenzo	09/12/1927	UTIC	
A	I	50758	AGRILLO	Luig	22/04/1954	UTIC	
A	I	52425	AHMAD	Khaled	03/02/1964	UTIC	
A	I	105966	AIELLO	Antonio	08/01/1964	UTIC	

Patienti nella popolazione: 7000

Risultati Query





## Risk of AF in General Population

- Hypertension is the most important risk factor for new AF
  - Population-attributable risk
    - 14-22%

Benjamin EJ, JAMA 1994; Psaty BM, Circulation 1997



# Atrial Fibrillation (AF) Risk Factors



Established Risk Factors	Estimated Increased Risk	Comments
Age	2	Per decade
Male Sex	1.5	
Hypertension	1.2-1.5	BP>140/90 mmHg
Valvular heart disease	1.8-3.4	
LV systolic dysfunction	4.5-5.9	
Obesity	1.4-2.4	
Alcohol consumption	1.3-1.5	Heavy alcohol use (>36 g/d)

Andrade J, Circ Res 2014



## Risk of AF in Hypertensives

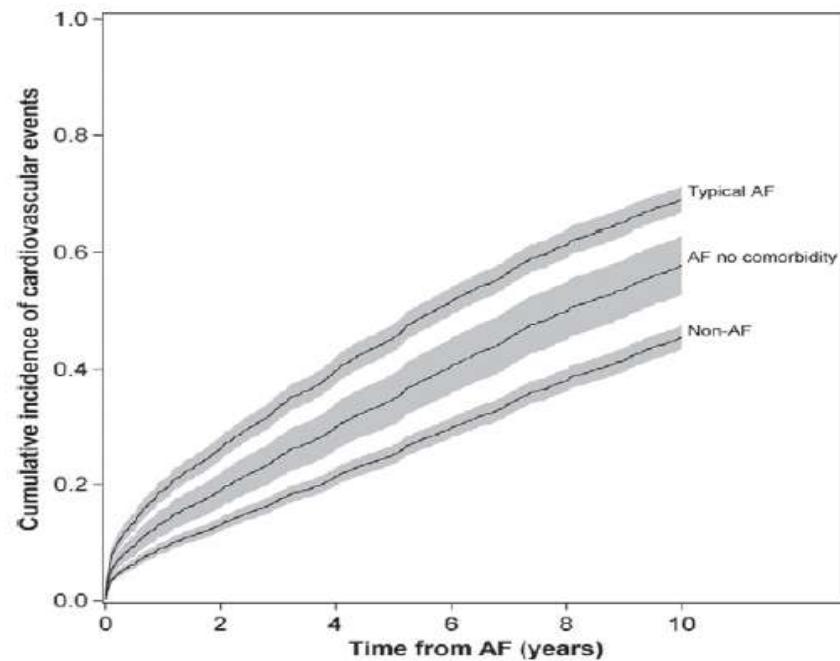
- Role of Systolic Blood Pressure (BP) at baseline:
  - Higher in patients who develop AF
    - Registries
    - Untreated patients
    - Treated patients

Verdecchia P, Hypertension 2003

Losi MA, Int J Cardiol 2015



# AF with and without Comorbidities



Kim EJ, Am Heart J 2016

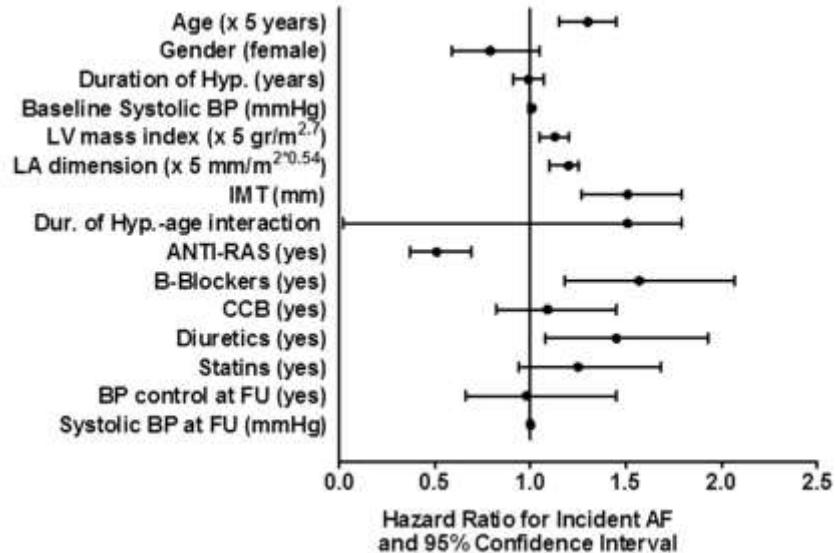


## Risk of AF in Hypertensives Systolic BP at baseline

TABLE 2. Independent Predictors of Atrial Fibrillation\*

Variable	Comparison	Relative Risk (95% Confidence Intervals)	P
Acute atrial fibrillation			
Age	10 years	1.75 (1.36–2.26)	0.001
LV mass	1 SD (14 g/height <sup>2.7</sup> )	1.20 (1.07–1.34)	0.001
Chronic atrial fibrillation			
Age	10 years	2.86 (1.72–4.76)	0.001
LV mass	1 SD (14 g/height <sup>2.7</sup> )	1.70 (1.19–2.43)	0.003
Left atrial diameter	1 SD (0.57 cm)	2.08 (1.31–3.31)	0.002

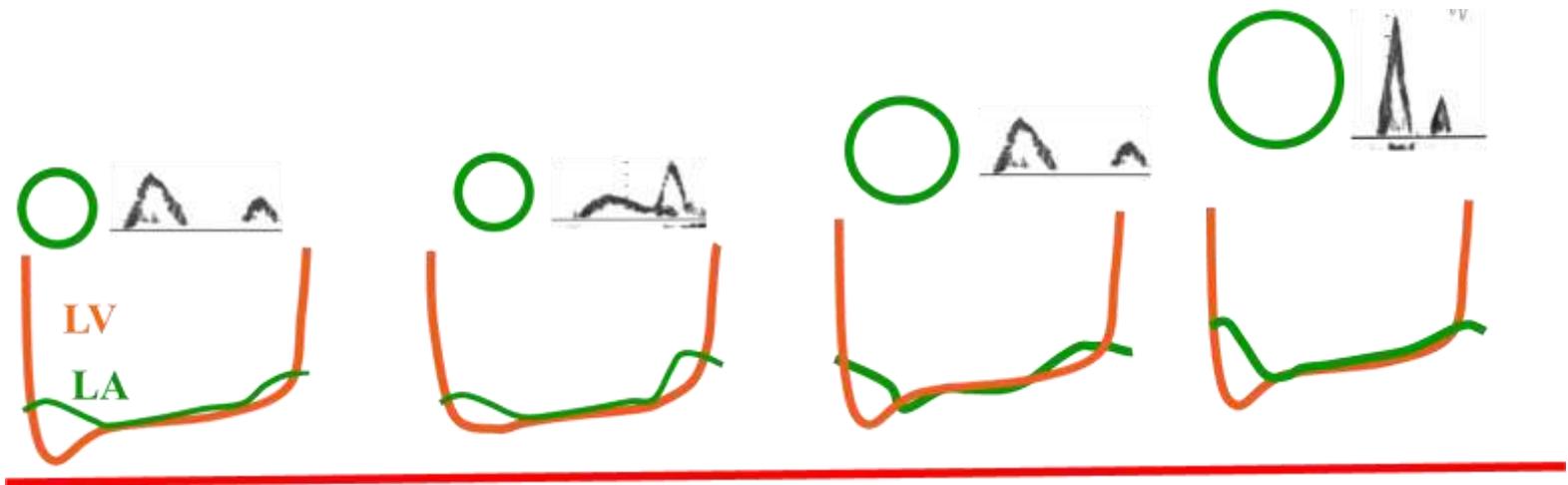
Verdecchia P, Hypertension 2003



Losi MA, Int J Cardiol 2015

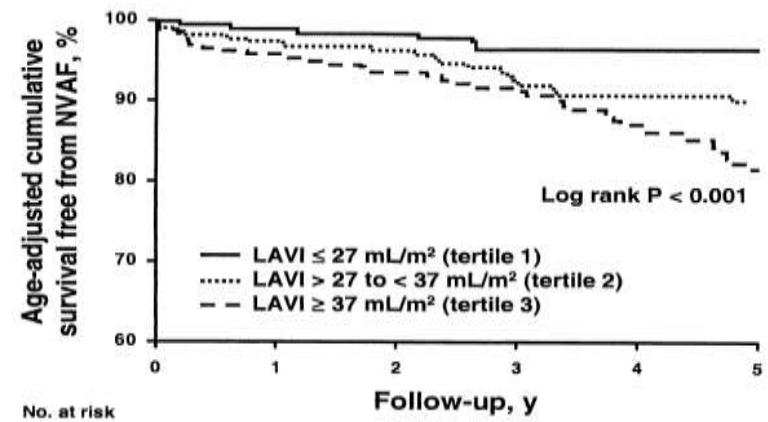
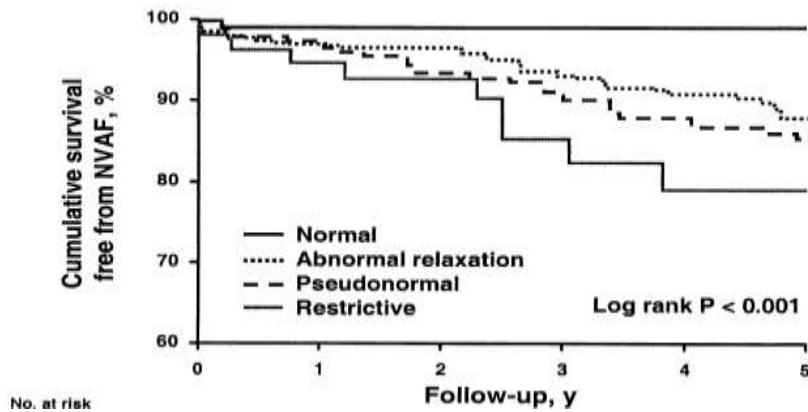


# Left Atrial Size and Diastolic Dysfunction





# Doppler «diastolic dysfunction» and AF

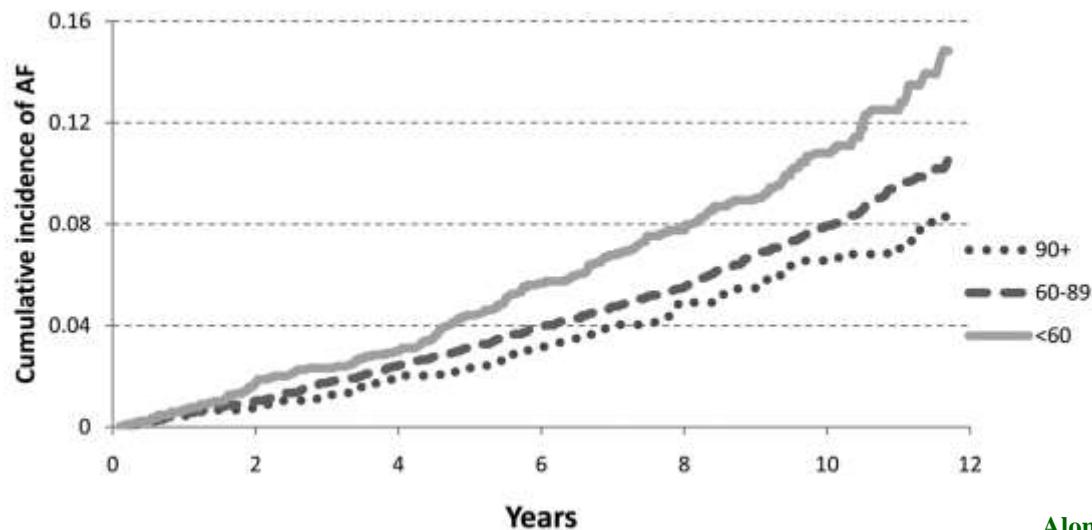


Tsang TSM, J Am Coll Cardiol 2002



## AF and Renal Function in General Population

- AF increases with severity of renal dysfunction



Alonso A, Circulation 2011

# Renal Function and Incident AF in Hypertensives



TABLE 1. Baseline Characteristics of Patients With and Without Future Occurrence of Atrial Fibrillation

Variable	Future Occurrence of Atrial Fibrillation		
	Total Population (N=2482)	No (n=2421)	Yes (n=61)
Creatinine, mmol/L	87.1 (21.8)	87.1 (22)	90.3 (25)

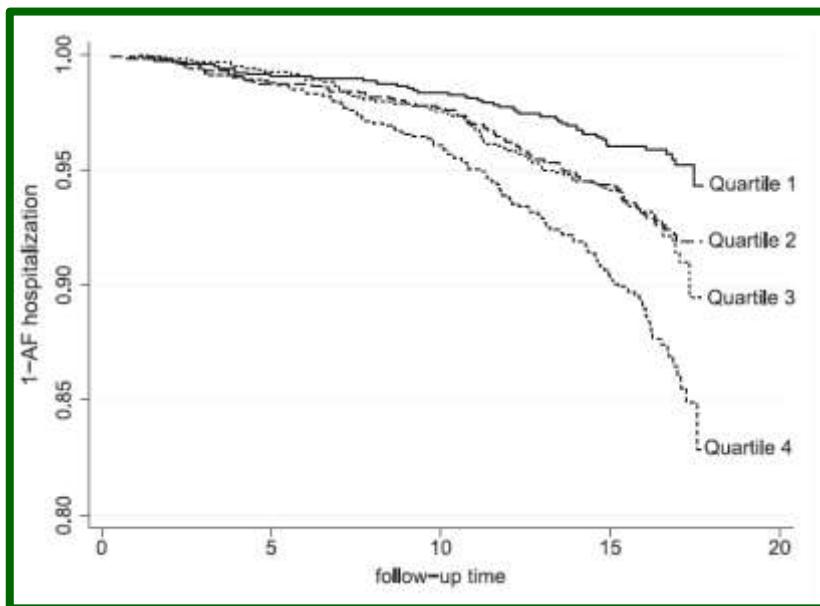
Verdecchia P, Hypertension 2003

Table 1  
Baseline characteristics of the studied population.

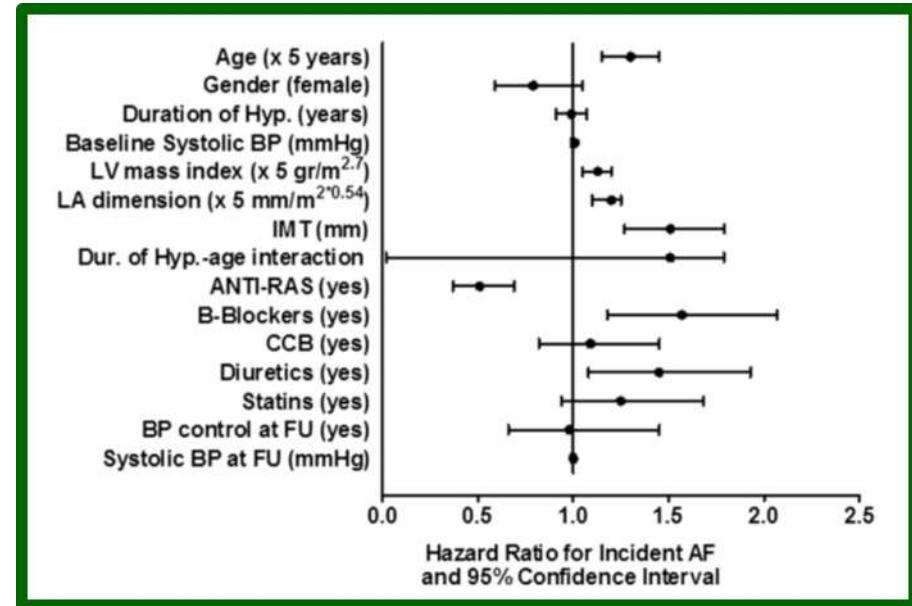
Parameter	No AF group 6945	AF group 117	p
GFR by simplified MDRD (ml/min/1.73 m <sup>2</sup> )	79 ± 18	77 ± 21	NS

Losi MA, Int J Cardiol 2015

# Carotid Intimal Medial Thickness and Incident AF



Eryd SA, Atherosclerosis 2013



Losi MA, Int J Cardiol 2015

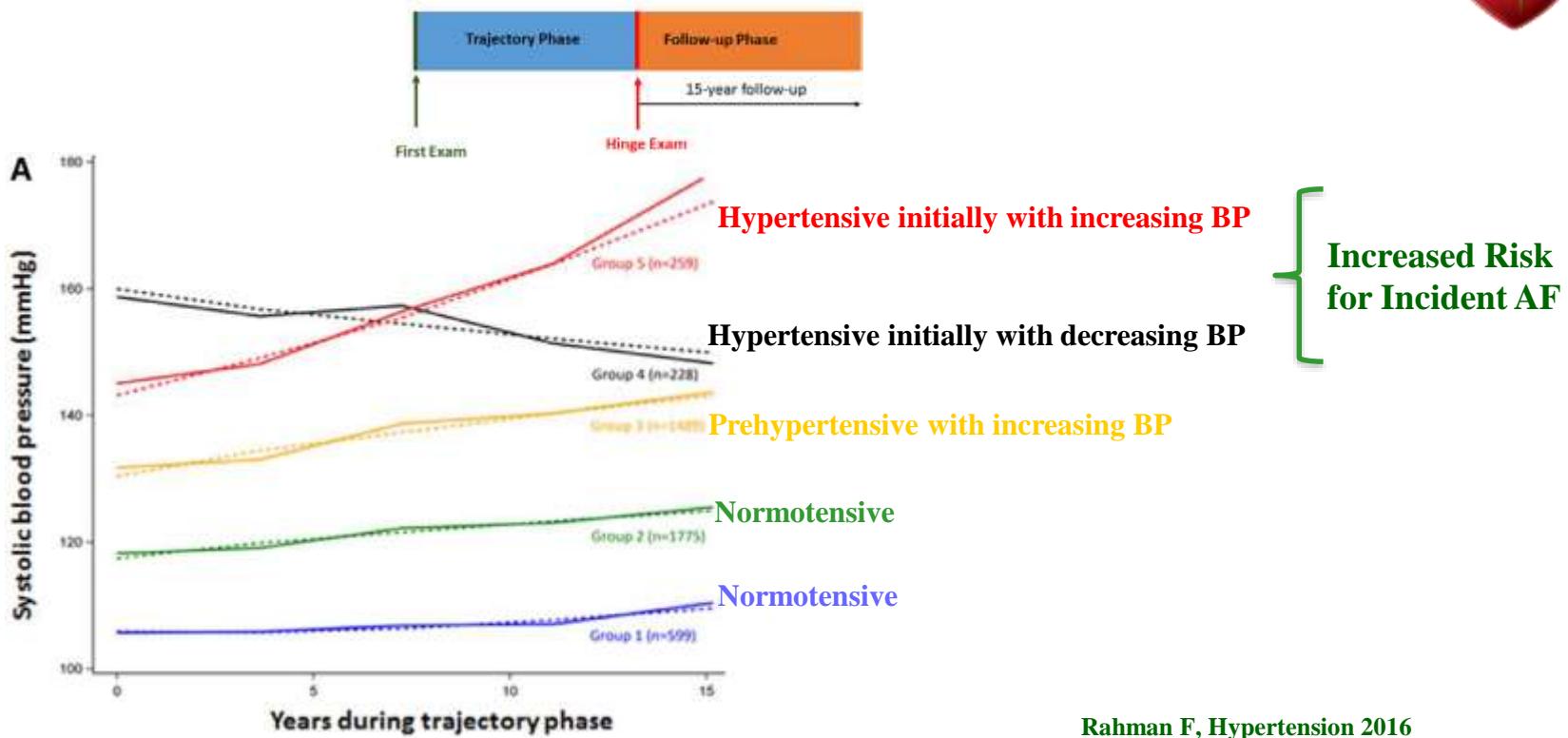


# Characteristics during follow-up and risk of AF

- Changes in
  - Systolic BP
  - LV hypertrophy
  - LA size
  - Renal Function

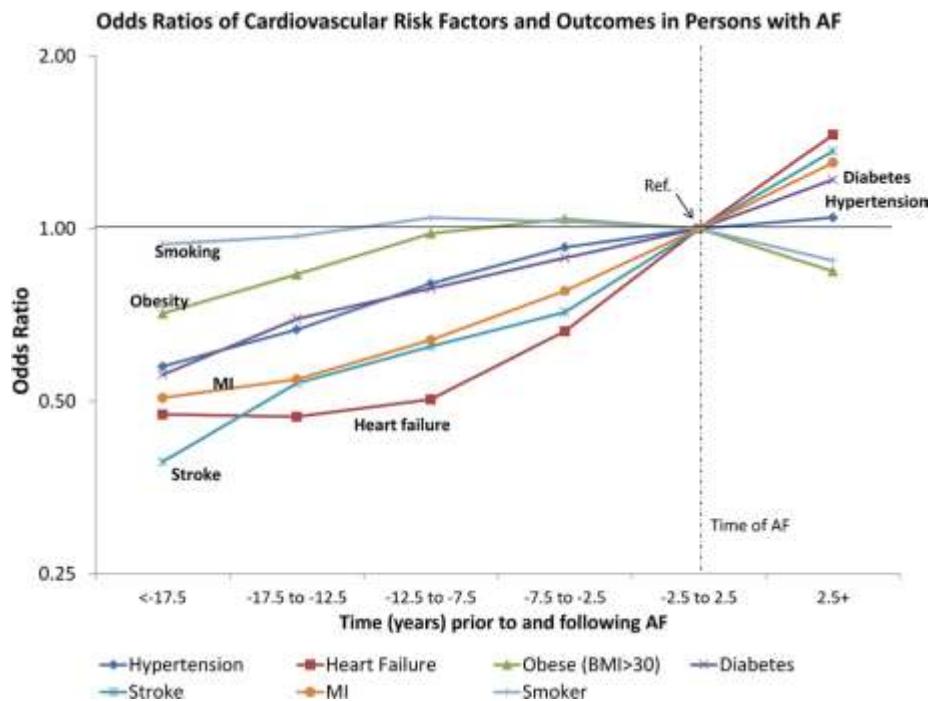


# SBP during Follow-up and AF

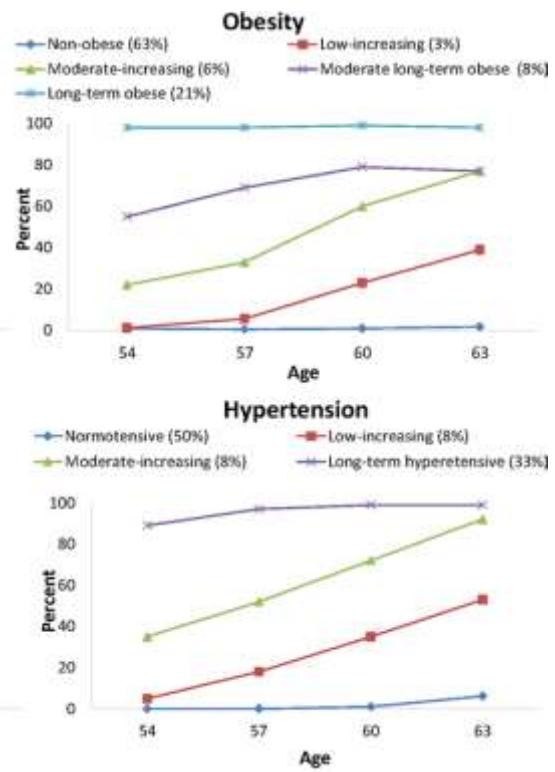




# SBP during Follow-up and AF

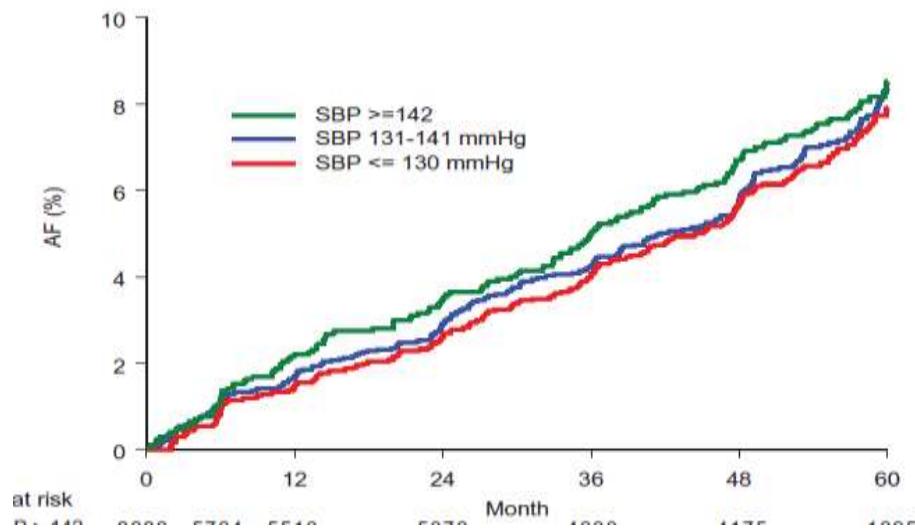


Norby FL, Circulation 2016





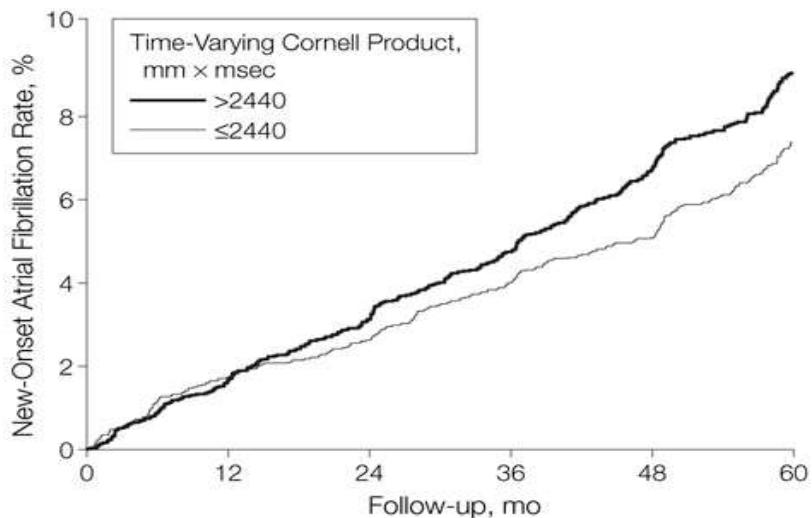
# SBP during Follow-up and AF



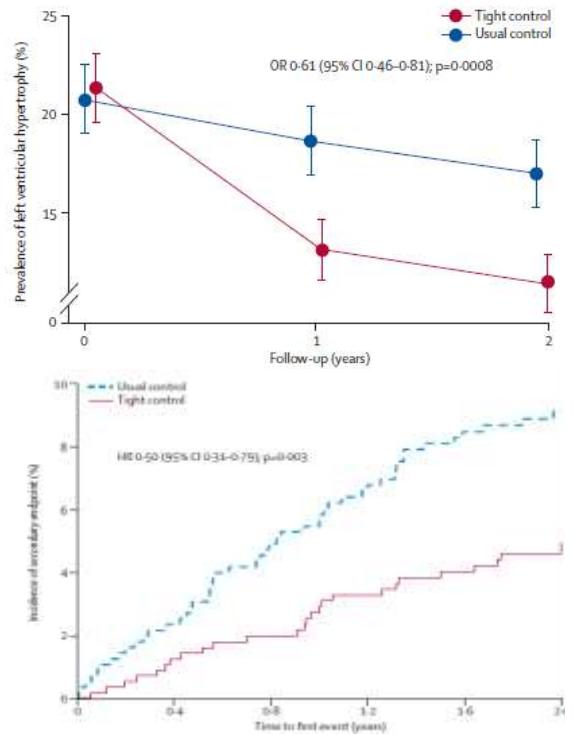
Okin PM, Hypertension 2015



# LVM during Follow-up and AF



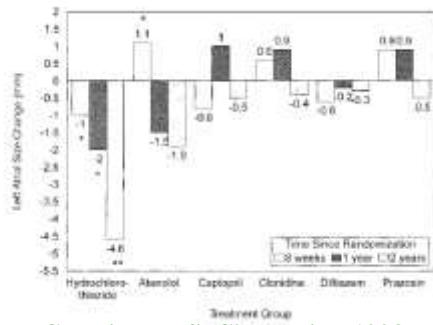
Okin PM, JAMA 2006



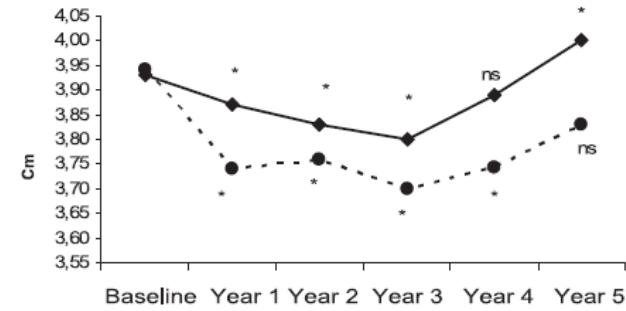
Verdecchia P, Lancet 2009



# LA Size During Follow-up

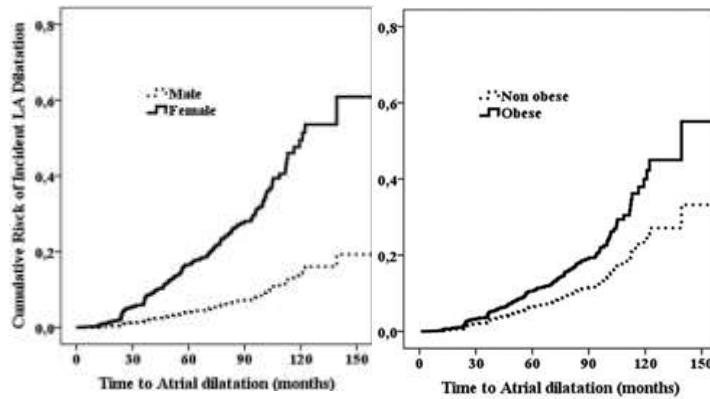


Gottdiener JS, Circulation 1998



Gerdts E, Hypertension 2007

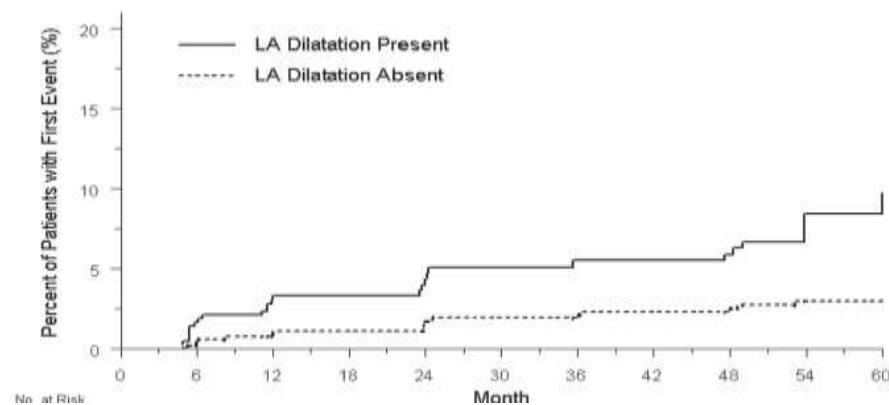
Left atrium dilates in 12% of hypertensive treated patients



Losi MA, Am J Hypert 2016



## LA Size during follow-up and AF



Wachtell K, Blood Pressure 2010



## Worsening Renal Function & AF

- We can hypothesize that
  - Development of worsening renal function contributes to a greater incidence of LV hypertrophy and LA dilatation
  - The development of AF should negatively affect kidney function
    - AF abolishes the contribution of atrial contraction to ventricular filling
      - Reduction of cardiac output



## Conclusions

- Hypertension and AF are strictly linked
  - At baseline predictors of incident AF:
    - Target organ damage
    - LA size
  - During follow-up the rate of AF is reduced by
    - Lower value of Systolic BP
    - LV hypertrophy and LA dilation regression

# Atrial fibrillation and cognitive decline

## A longitudinal cohort study

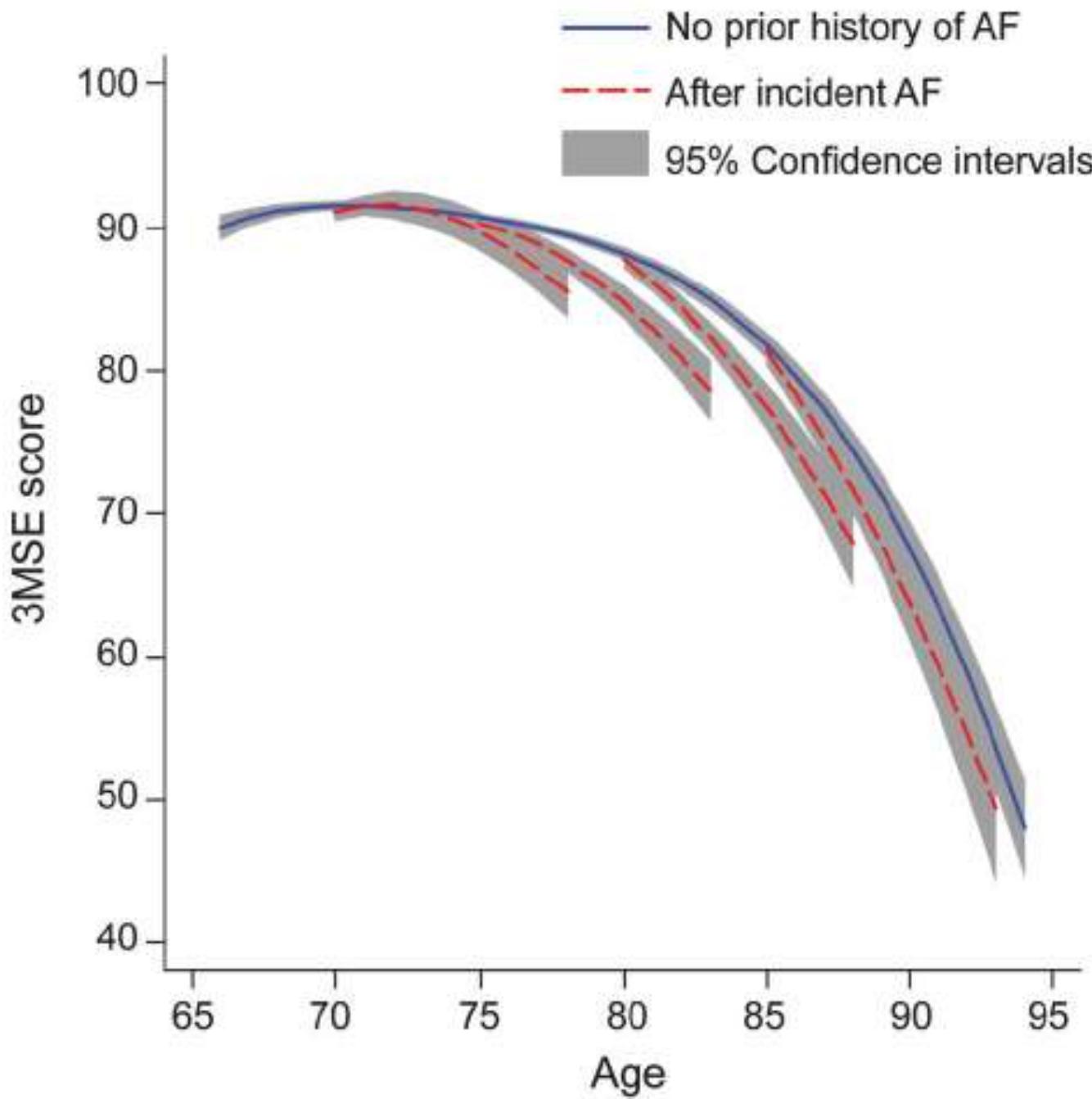
### ABSTRACT

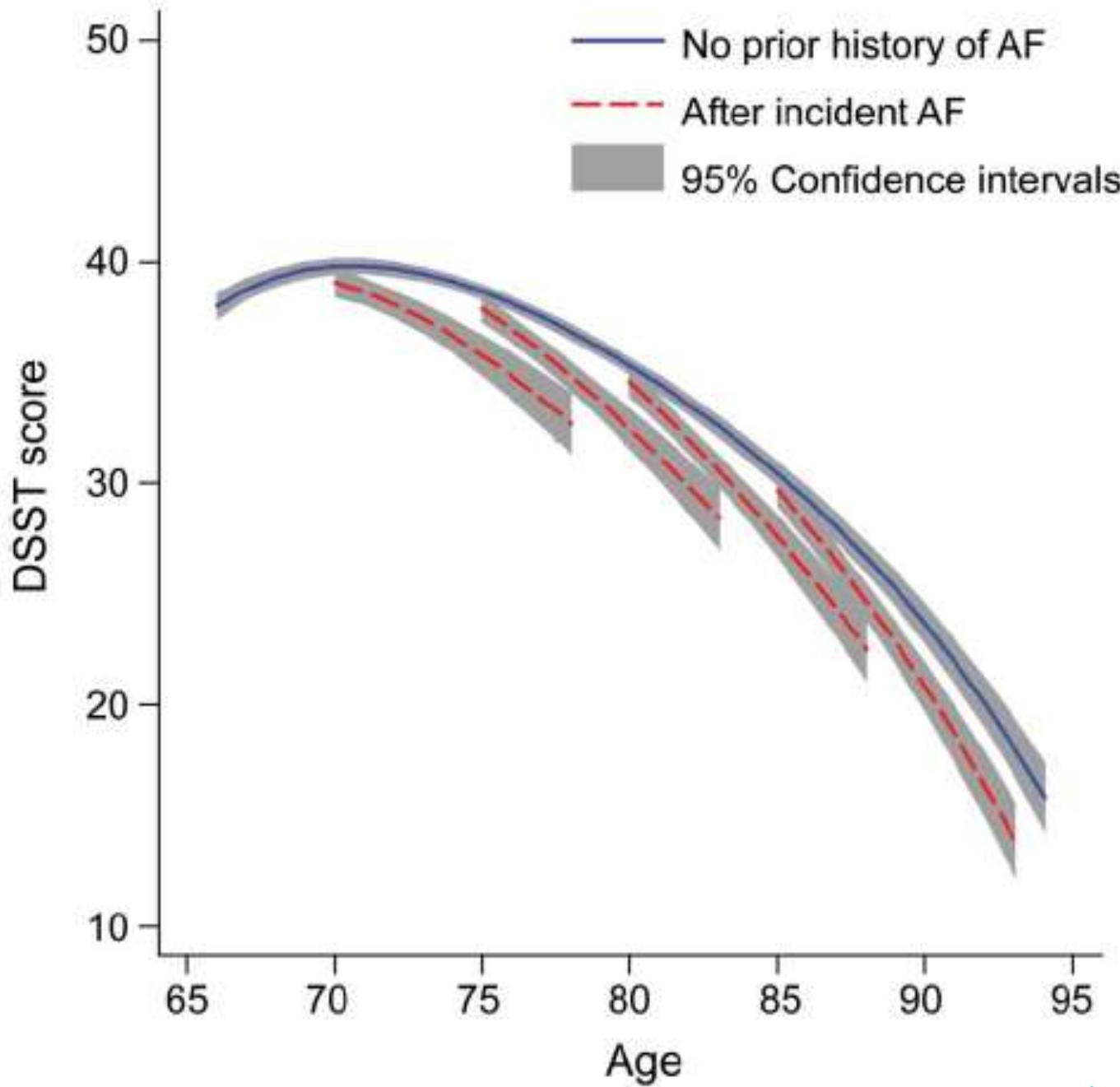
**Objective:** We sought to determine whether in the absence of clinical stroke, people with atrial fibrillation experience faster cognitive decline than people without atrial fibrillation.

**Methods:** We conducted a longitudinal analysis in the Cardiovascular Health Study, a community-based study of 5,888 men and women aged 65 years and older, enrolled in 1989/1990 or 1992/1993. Participants did not have atrial fibrillation or a history of stroke at baseline. Participants were censored when they experienced incident clinical stroke. Incident atrial fibrillation was identified by hospital discharge diagnosis codes and annual study ECGs. The main outcome was rate of decline in mean scores on the 100-point Modified Mini-Mental State Examination (3MSE), administered annually up to 9 times.

**Results:** Analyses included 5,150 participants, of whom 552 (10.7%) developed incident atrial fibrillation during a mean of 7 years of follow-up. Mean 3MSE scores declined faster after incident atrial fibrillation compared with no prior atrial fibrillation. For example, the predicted 5-year decline in mean 3MSE score from age 80 to age 85 was -6.4 points (95% confidence interval [CI]: -7.0, -5.9) for participants without a history of atrial fibrillation, but was -10.3 points (95% CI: -11.8, -8.9) for participants experiencing incident atrial fibrillation at age 80, a 5-year difference of -3.9 points (95% CI: -5.3, -2.5).

**Conclusions:** In the absence of clinical stroke, people with incident atrial fibrillation are likely to reach thresholds of cognitive impairment or dementia at earlier ages than people with no history of atrial fibrillation. *Neurology®* 2013;81:119-125





# 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

**Background:** Atrial fibrillation (AF) has been linked with an increased risk for cognitive impairment and dementia.

**Purpose:** To complete a meta-analysis of studies examining the association between AF and cognitive impairment.

**Data Sources:** Search of MEDLINE, PsycINFO, Cochrane Library, CINAHL, and EMBASE databases and hand search of article references.

**Study Selection:** Prospective and nonprospective studies reporting adjusted risk estimates for the association between AF and cognitive impairment.

**Data Extraction:** Two abstracters independently extracted data on study characteristics, risk estimates, methods of AF and outcome ascertainment, and methodological quality.

**Data Synthesis:** Twenty-one studies were included in the meta-analysis. Atrial fibrillation was significantly associated with a higher risk for cognitive impairment in patients with first-ever or recurrent stroke (relative risk [RR], 2.70 [95% CI, 1.82 to 4.00]) and in a broader population including patients with or without a history of stroke (RR, 1.40 [CI, 1.19 to 1.64]). The association in the latter group remained significant independent proof of clinical stroke his-

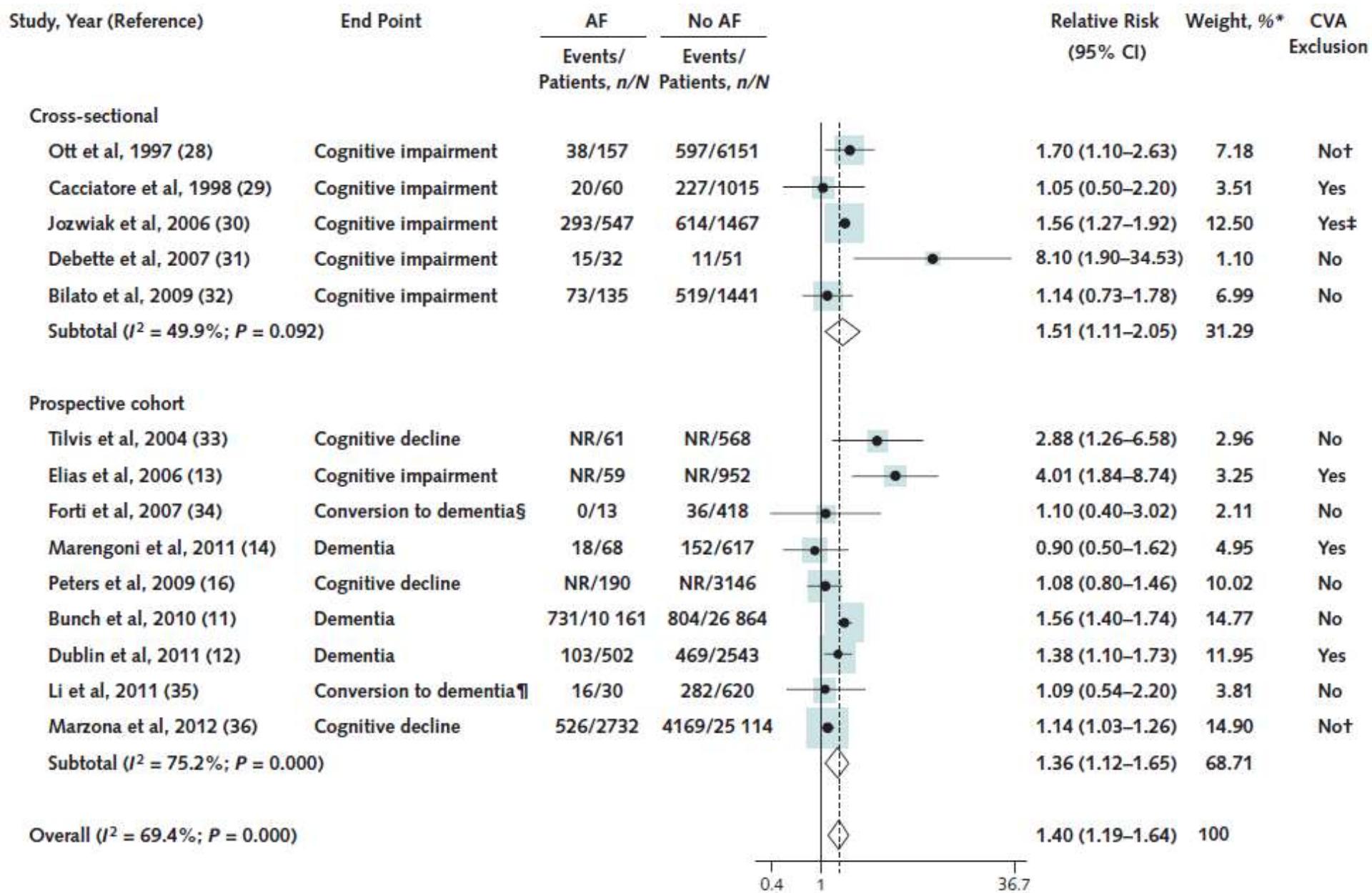
tory (RR, 1.34 [CI, 1.13 to 1.58]). However, there was significant heterogeneity among studies of the broader population ( $I^2 = 69.4\%$ ). Limiting the analysis to prospective studies yielded similar results (RR, 1.36 [CI, 1.12 to 1.65]). Restricting the analysis to studies of dementia eliminated the significant heterogeneity ( $P = 0.137$ ) but did not alter the pooled estimate substantially (RR, 1.38 [CI, 1.22 to 1.56]).

**Limitations:** There is an inherent bias because of confounding variables in observational studies. There was significant heterogeneity among included studies.

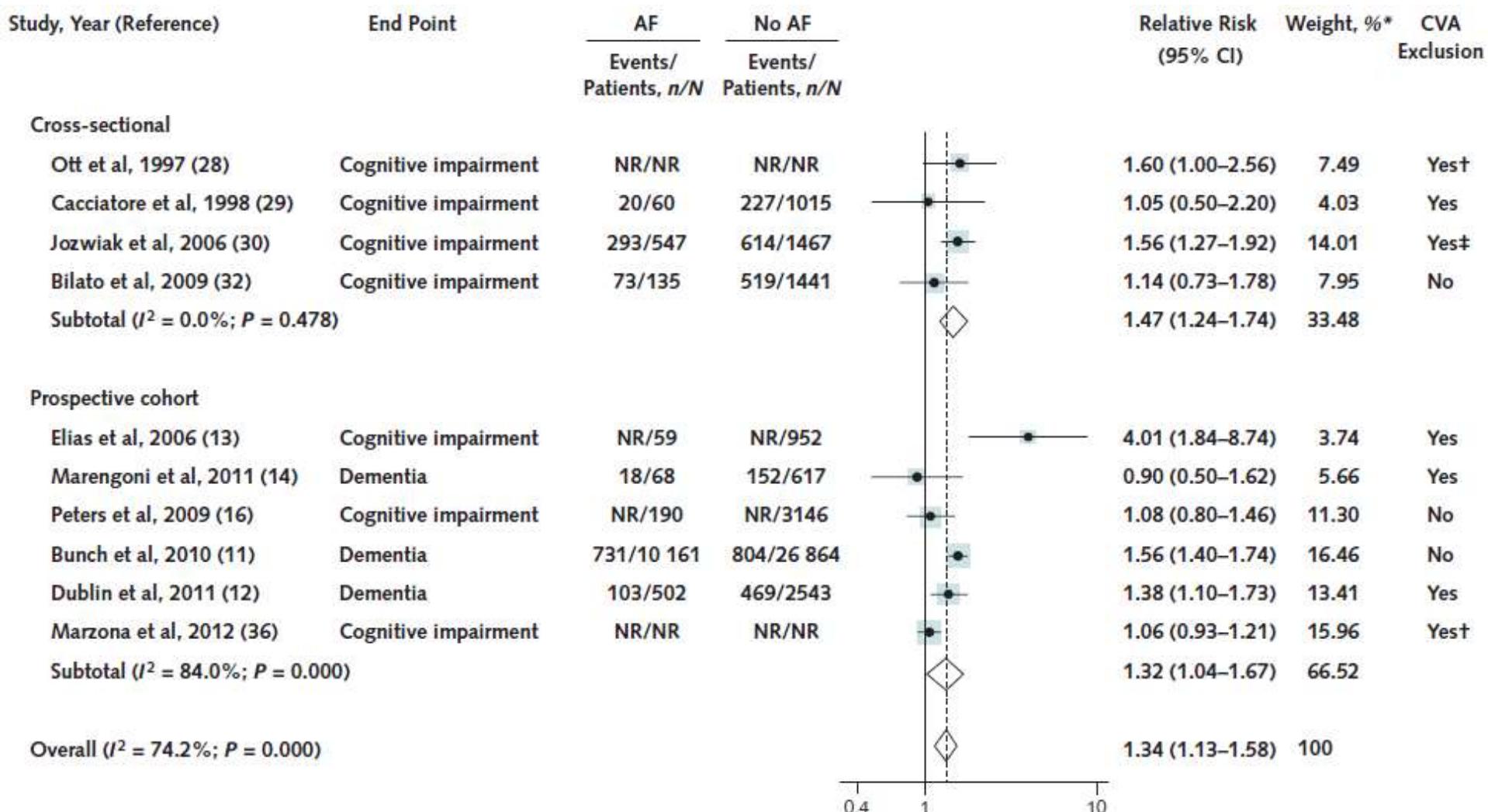
**Conclusion:** Evidence suggests that AF is associated with a higher risk for cognitive impairment and dementia, with or without a history of clinical stroke. Further studies are required to elucidate the association between AF and subtypes of dementia as well as the cause of cognitive impairment.

**Primary Funding Source:** Deane Institute for Integrative Research in Atrial Fibrillation and Stroke at the Massachusetts General Hospital.

# The association between AF and cognitive impairment in patients with or without history of stroke



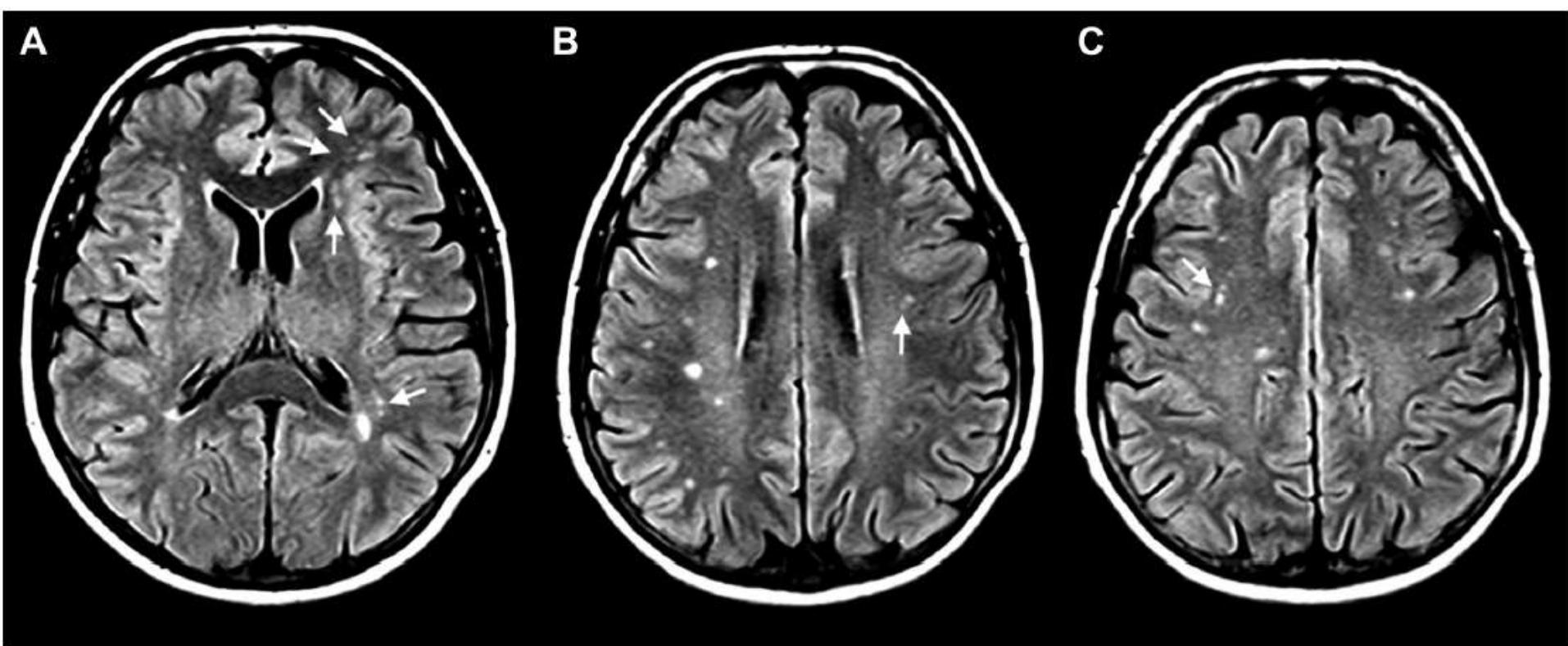
# The association between AF and cognitive impairment independent of stroke history

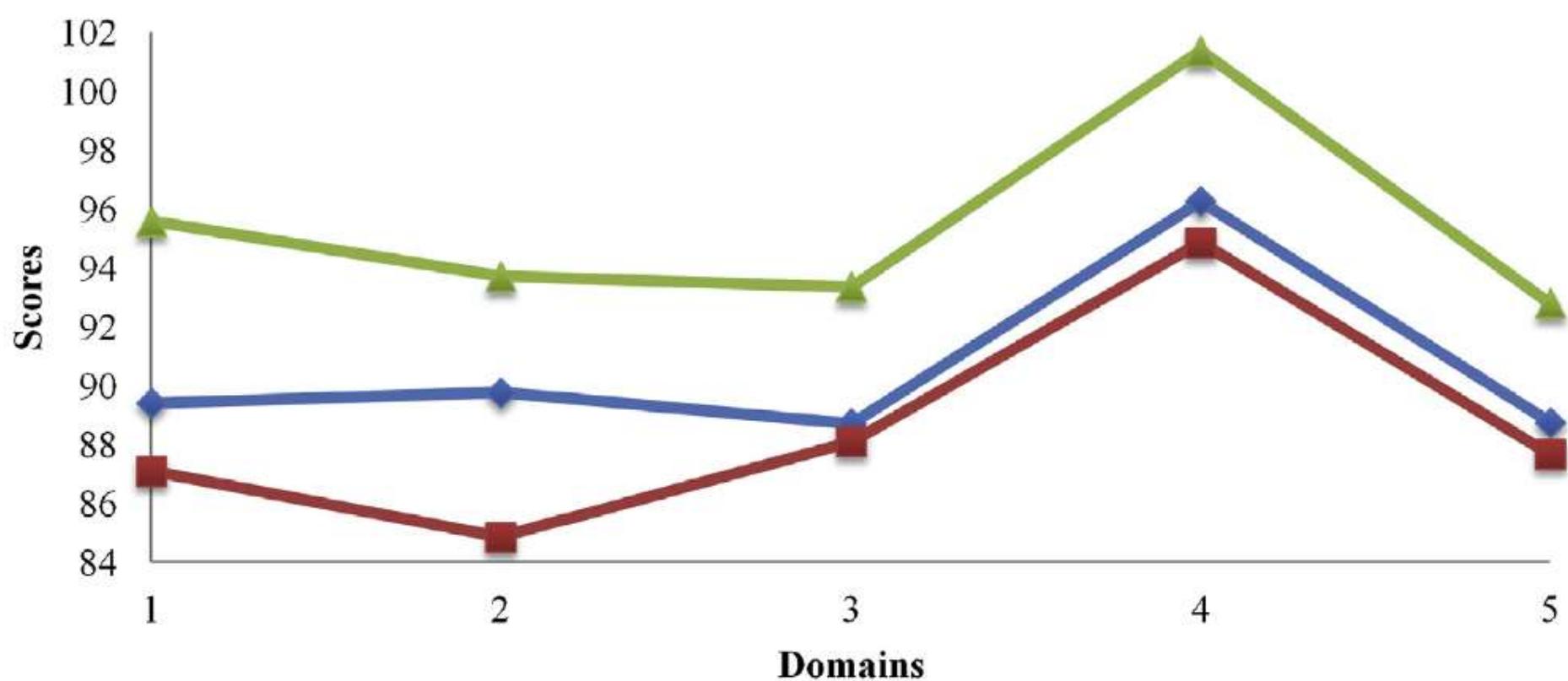


# Prevalence of Silent Cerebral Ischemia in Paroxysmal and Persistent Atrial Fibrillation and Correlation With Cognitive Function

Fiorenzo Gaita, MD,\* Laura Corsinovi, MD, PhD,\* Matteo Anselmino, MD, PhD,\*  
Cristina Raimondo, MD,\* Martina Pianelli, MD,\* Elisabetta Toso, MD,\* Laura Bergamasco, PROF,†  
Carlo Boffano, MD,‡ Maria Consuelo Valentini, MD,§ Federico Cesarani, MD,||  
Marco Scaglione, MD¶

Turin, Milan, and Asti, Italy





	Controls (N = 90)	PRX AF (N = 90)	PER AF (N = 90)	p PRX / controls	p PER / controls	p PRX / PER
Domains	92.4 ± 15.4	86.2 ± 13.8	82.9 ± 11.5	< 0.01	< 0.01	0.08
1-Immediate Memory	95.6 ± 17.5	89.9 ± 14.7	87.1 ± 16.9	0.02	< 0.01	0.24
2-Visuo-spatial abilities	93.8 ± 16.7	89.9 ± 18.2	84.8 ± 14.8	0.14	< 0.01	0.04
3-Language	92.9 ± 11.4	88.8 ± 9.1	88.1 ± 8.7	< 0.01	< 0.01	0.59
4-Attention	101.4 ± 21.2	96.6 ± 16.6	94.9 ± 15.6	0.09	0.02	0.47
5-Delayed memory	93.5 ± 11.7	88.7 ± 14.7	87.7 ± 14	0.02	< 0.01	0.64

# Increased risk of cognitive and functional decline in patients with atrial fibrillation: results of the ONTARGET and TRANSCEND studies

Irene Marzona MSc PharmD, Martin O'Donnell MB PhD, Koon Teo MB PhD, Peggy Gao MSc, Craig Anderson MD PhD, Jackie Bosch BScOT MSc, Salim Yusuf MD DPhil

## ABSTRACT

**Background:** The role of atrial fibrillation in cognitive impairment and dementia, independent of stroke, is uncertain. We sought to determine the association of atrial fibrillation with cognitive and physical impairment in a large group of patients at high cardiovascular risk.

**Methods:** We conducted a post-hoc analysis of two randomized controlled trials involving 31 546 patients, the aims of which were to evaluate the efficacy of treatment with ramipril plus telmisartan (ONTARGET) or telmisartan alone (TRANSCEND) in reducing cardiovascular disease. We evaluated the cognitive function of participants at baseline and after two and five years using the Mini-Mental State Examination (MMSE). In addition, we recorded incident dementia, loss of independence in activities of daily living and admission to long-term care facilities. We used a Cox regression model adjusting for main confounders to determine the association between atrial fibrillation and our primary outcomes: a decrease of three or more points in MMSE score, incident dementia, loss of

independence in performing activities of daily living and admission to long-term care.

**Results:** We enrolled 31 506 participants for whom complete information on atrial fibrillation was available, 70.4% of whom were men. The mean age of participants was 66.5 years, and the mean baseline MMSE score was 27.7 (standard deviation 2.9) points. At baseline, 1016 participants (3.3%) had atrial fibrillation, with the condition developing in an additional 2052 participants (6.5%) during a median follow-up of 56 months. Atrial fibrillation was associated with an increased risk of cognitive decline (hazard ratio [HR] 1.14, 95% confidence interval [CI] 1.03–1.26), new dementia (HR 1.30, 95% CI 1.14–1.49), loss of independence in performing activities of daily living (HR 1.35, 95% CI 1.19–1.54) and admission to long-term care facilities (HR 1.53, 95% CI 1.31–1.79). Results were consistent among participants with and without stroke or receiving antihypertensive drugs.

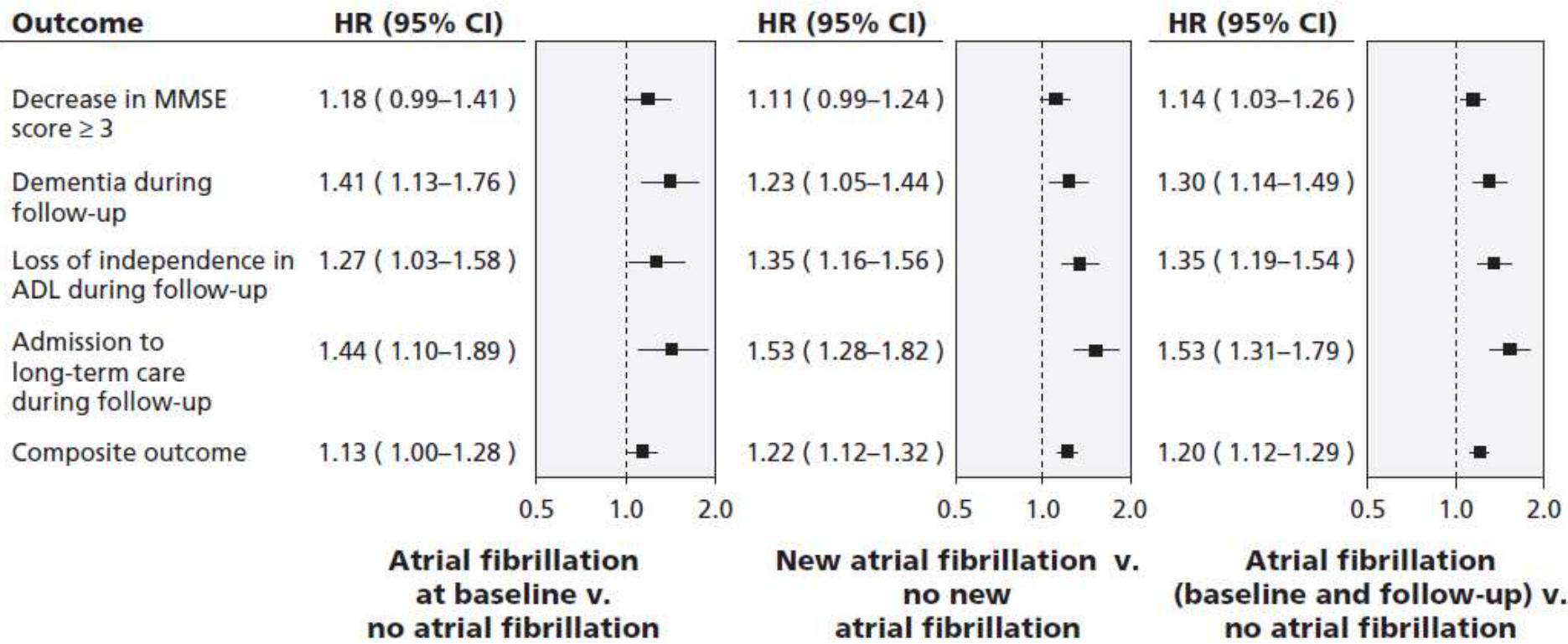
**Interpretation:** Cognitive and functional decline are important consequences of atrial fibrillation, even in the absence of overt stroke.

**Competing interests:** Martin O'Donnell has received grants, consulting fees and payment for lectures from Boehringer-Ingelheim. Koon Teo has received research grants from the Canadian Institutes of Health Research and the Heart and Stroke Foundation of Ontario. Craig Anderson has received project and fellowship grants from the National Health and Medical Research Council of Australia. No other competing interests were declared.

This article has been peer reviewed.

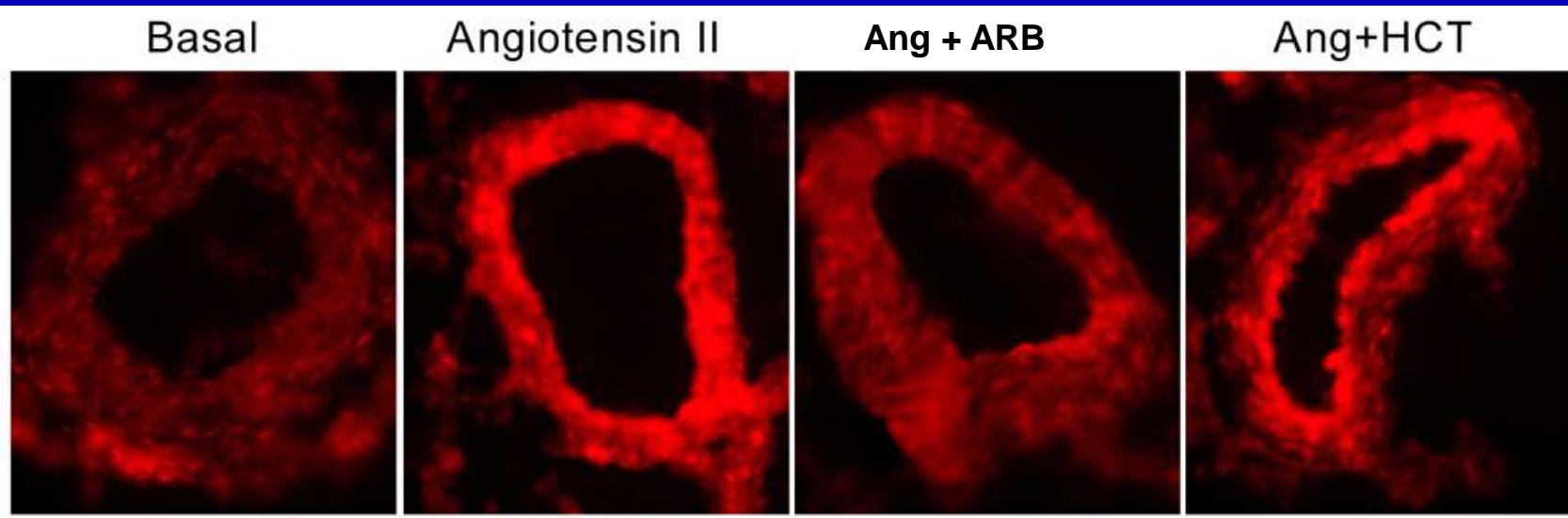
**Correspondence to:**  
Dr. Irene Marzona,  
[irene.marzona@phri.ca](mailto:irene.marzona@phri.ca)

**CMAJ 2012. DOI:10.1503/cmaj.111173**

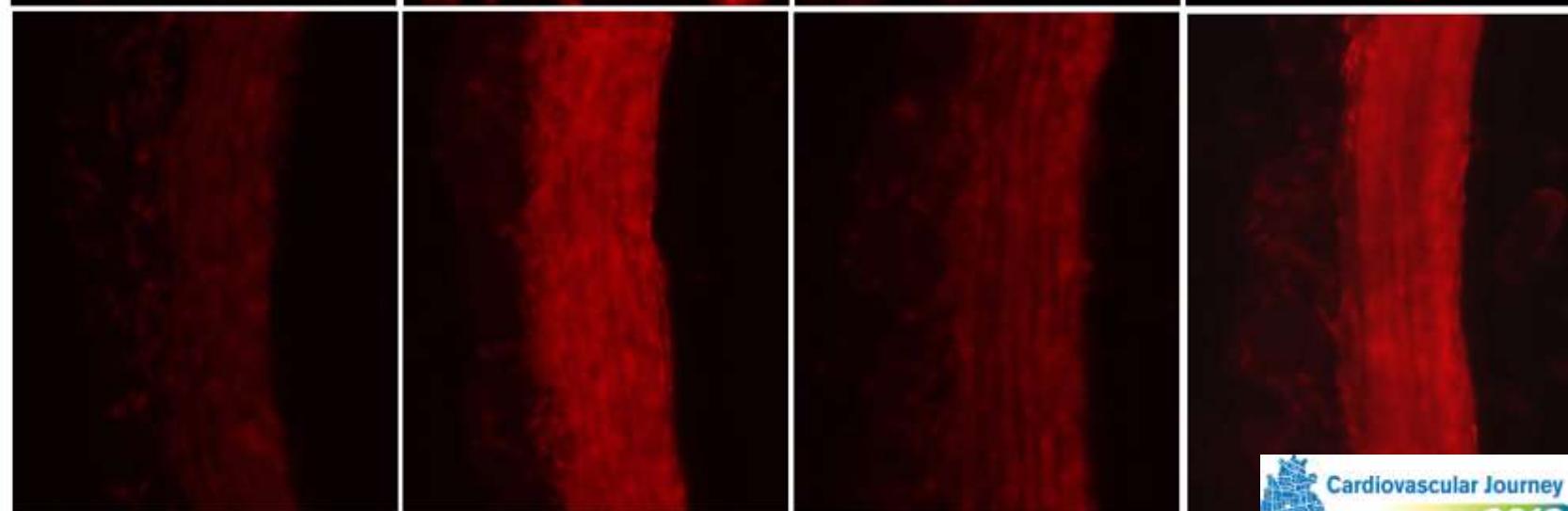


# AT-1 antagonism protects against angiotensin- induced vascular oxidative stress

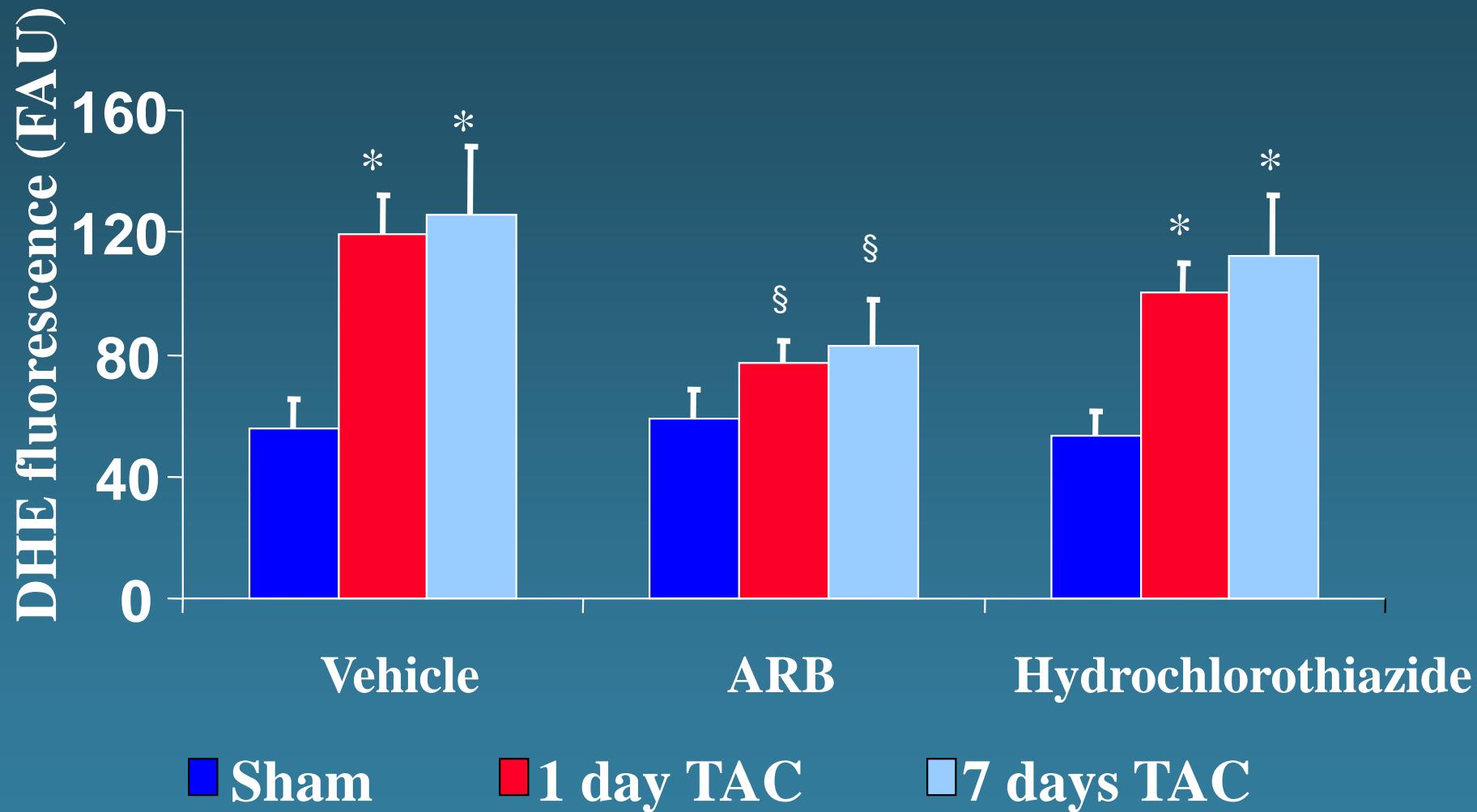
Basilar artery



Aorta



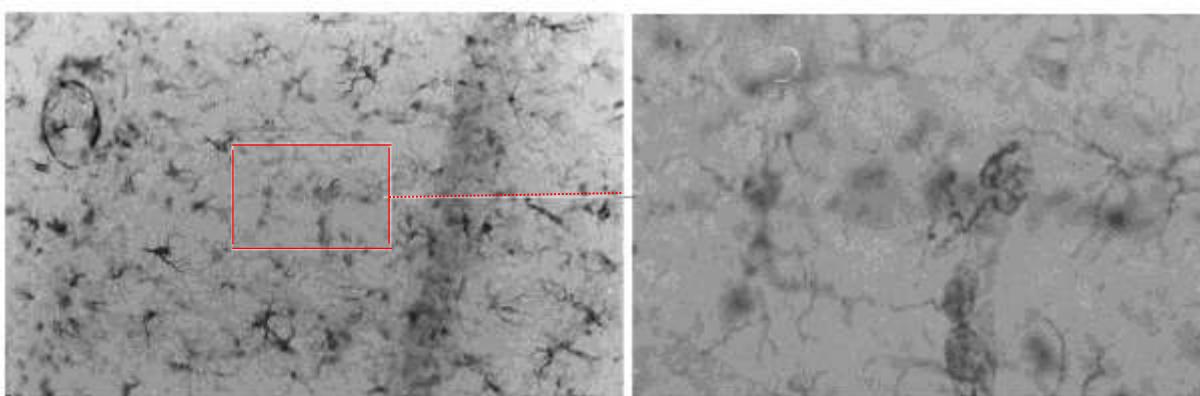
# AT-1 antagonism protects against pressure-induced oxidative stress in hippocampus



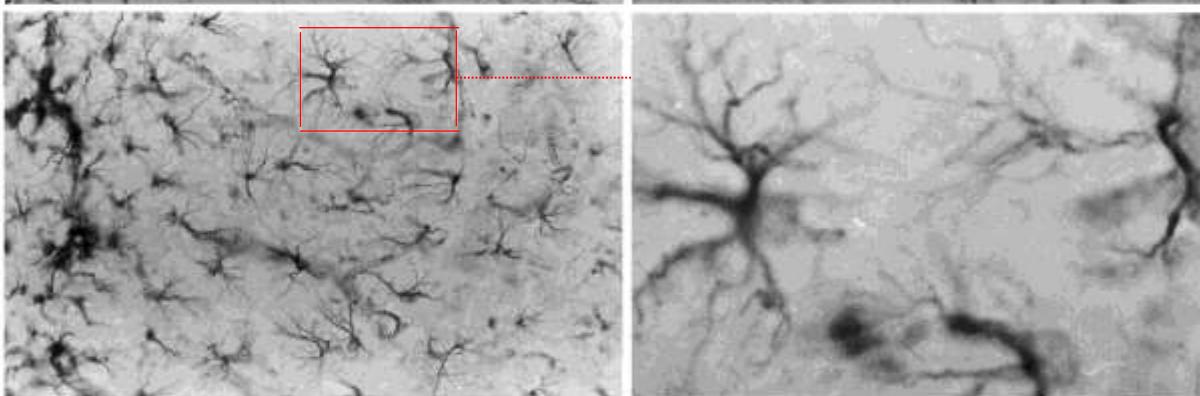
\* P < 0.05 vs. Sham

# Astrocytes in the hippocampus

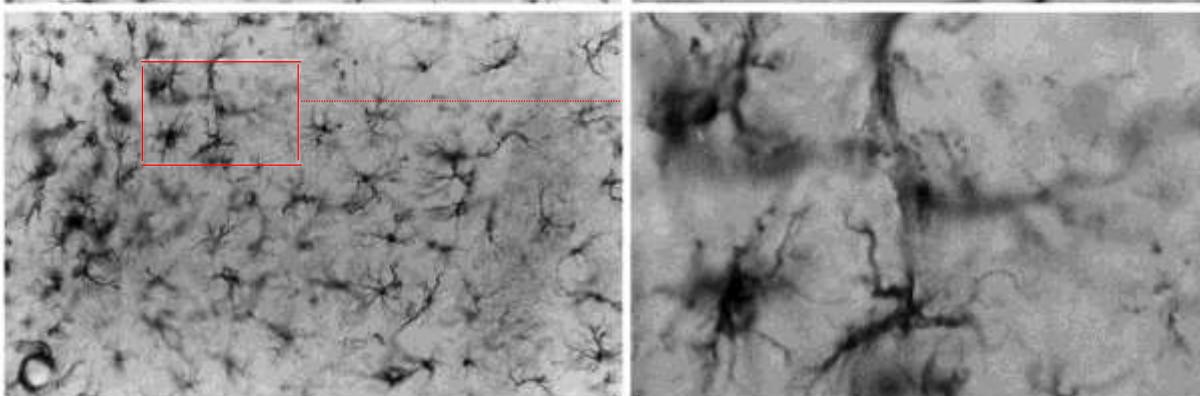
Sham



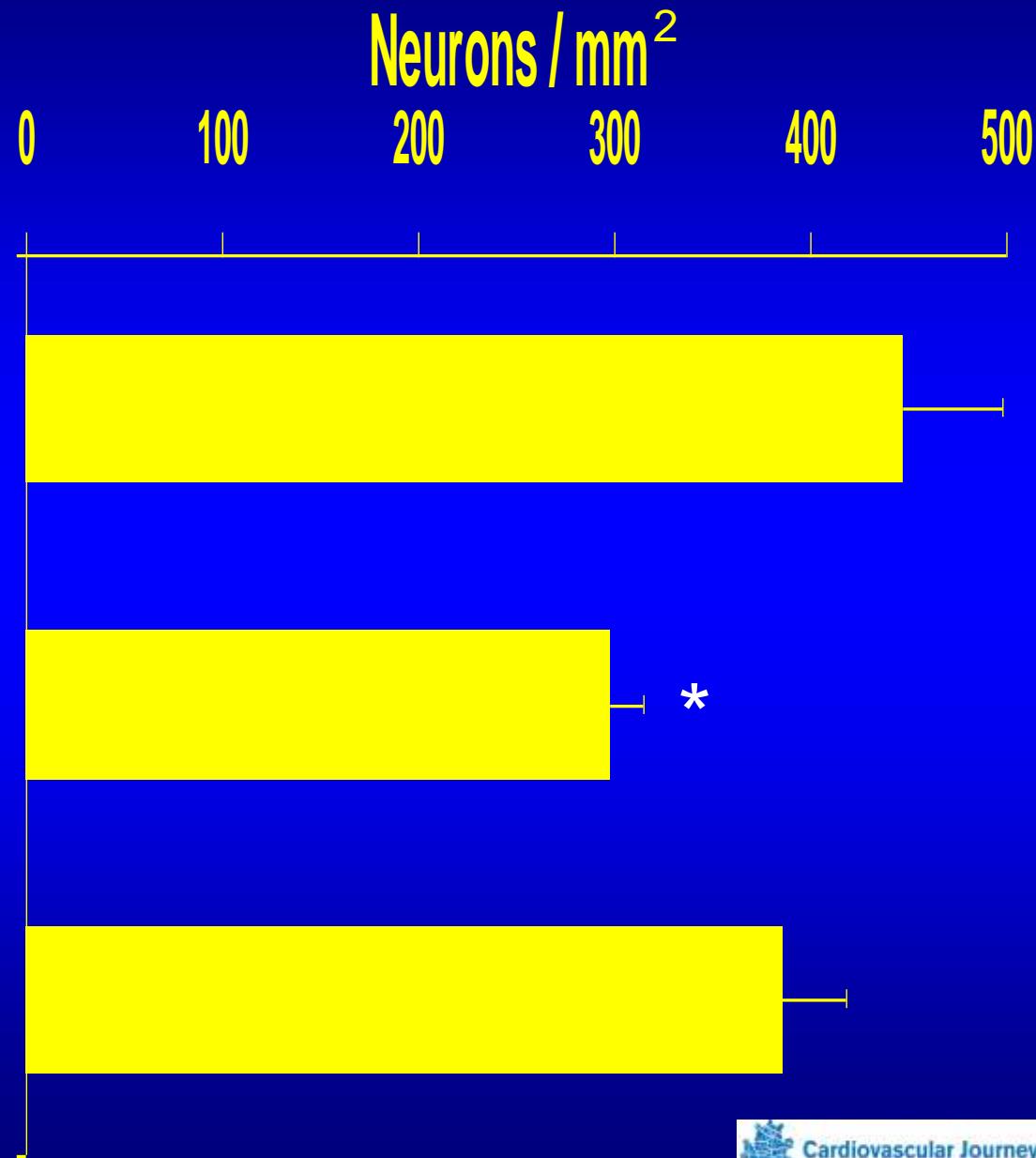
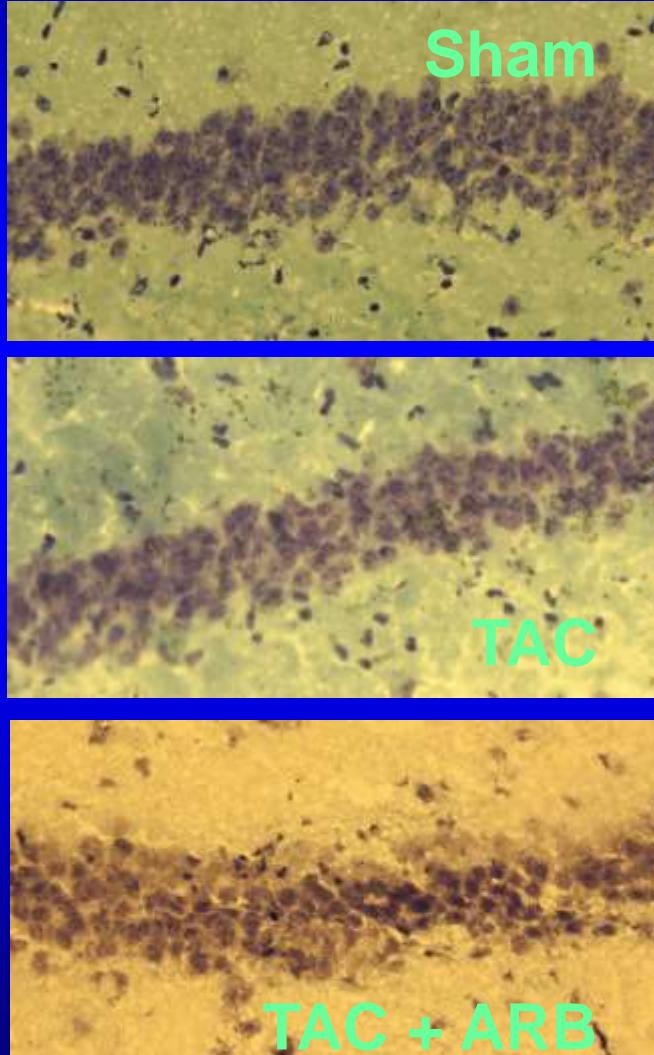
1 day  
TAC



7 days  
TAC

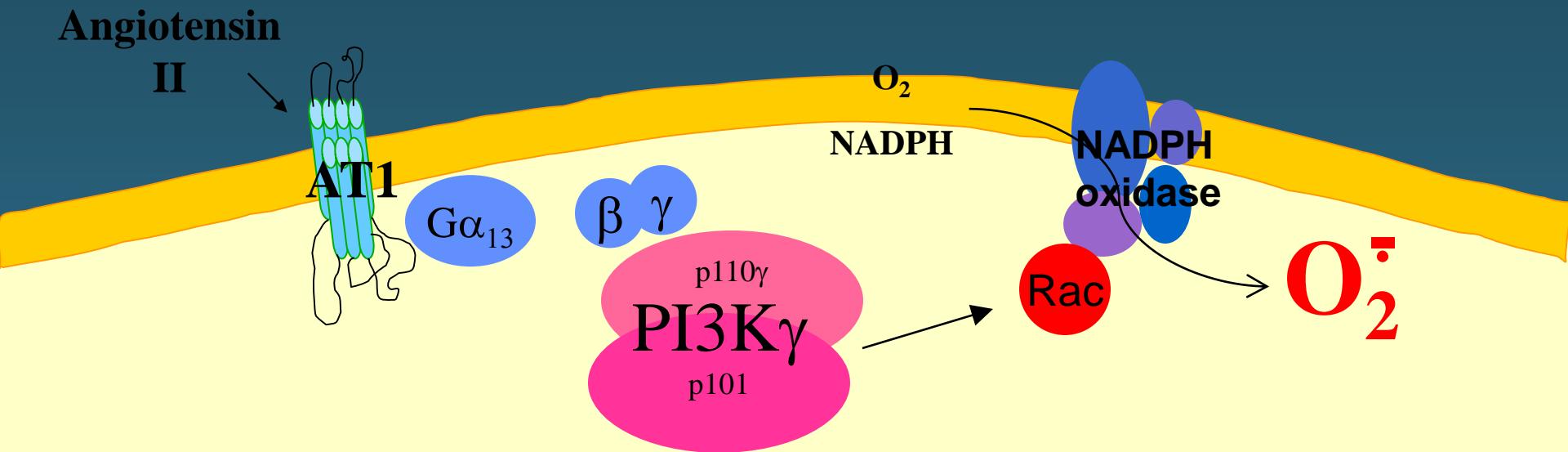


# Neuronal degeneration in hippocampus



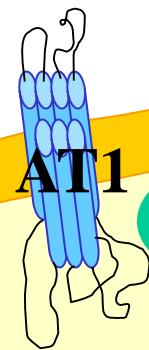
Cardiovascular Journey

2013



**Angiotensin**

**II**



**G $\alpha_{13}$**

$\beta$   $\gamma$

p110 $\gamma$   
**PI3K $\gamma$**   
p101

$O_2$

NADPH

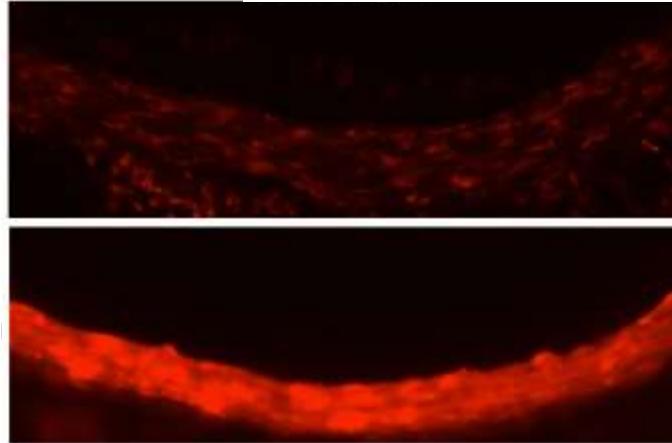
NADPH  
oxidase

Rac

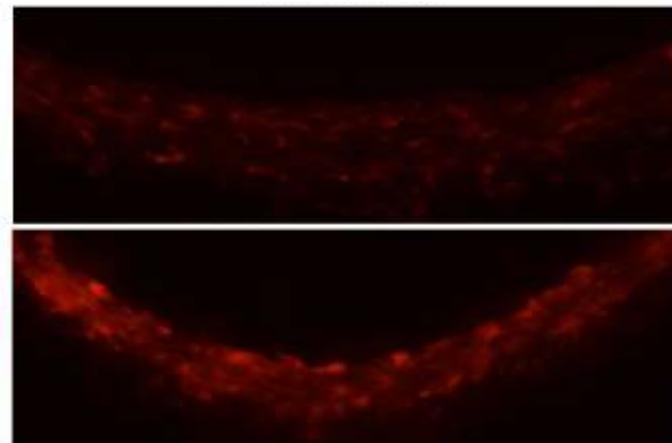
$O_2^{\cdot}$

# Angiotensin II-induced vascular oxidative stress

Dihydroethidium

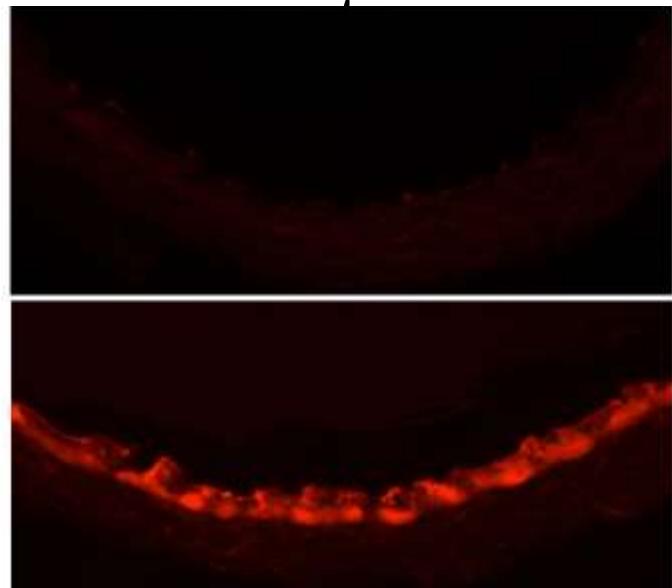


PI3K $\gamma$  +/+



PI3K $\gamma$  -/-

Nitrotyrosine



Ctr

Ang

Ctr

Ang

## **Subtle Post-Procedural Cognitive Dysfunction After Atrial Fibrillation Ablation**

Caroline Medi, BMED, PHD,\* Lisbeth Evered, BSc, MBIOSTATS,† Brendan Silbert, MBBS,†  
Andrew Teh, MBBS, PHD,\* Karen Halloran, RN,\* Joseph Morton, MBBS, PHD,\*  
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