FIBRILLAZIONE ATRIALE

Mauro Zennaro





GIANNI

67 aa

- Ipertensione arteriosa
- Sovrappeso

2014 intervento di chirurgia bariatrica

EPISODI RECIDIVANTI DI CARDIOPALMO

Recommendations for diagnostic workup of atrial fibrillation patients

	Recommendations	Class ^a	Level ^b
	ECG documentation is required to establish the diagnosis of AF.		B
	A full cardiovascular evaluation, including an accurate history, careful clinical examination, and assessment of concomitant conditions, is recommended in all AF patients.	1	U
	Transthoracic echocardiography is recommended in all AF patients to guide management.	1	C
\langle	Long-term ECG monitoring should be considered in selected patients to assess the adequacy of rate control in symptomatic patients and to relate symptoms with AF episodes.	lla	C





....FRANCO.....IL COGNATO DEL PAZIENTE (CHE NON PRESENTA SINTOMI)



	AF pattern	Definition
	First diagnosed AF	AF that has not been diagnosed before, irrespective of the duration of the arrhythmia or the presence and severity of AF-related symptoms.
AZIENTE	Paroxysmal AF	Self-terminating, in most cases within 48 hours. Some AF paroxysms may continue for up to 7 days. ^a AF episodes that are cardioverted within 7 days should be considered paroxysmal. ^a
OGNATO	Persistent AF	AF that lasts longer than 7 days, including episodes that are terminated by cardioversion, either with drugs or by direct current cardioversion, after 7 days or more.
	Long-standing persistent AF	Continuous AF lasting for ≥ 1 year when it is decided to adopt a rhythm control strategy.
ESC Guide	Permanent AF	AF that is accepted by the patient (and physician). Hence, rhythm control interventions are, by definition, not pursued in patients with permanent AF. Should a rhythm control strategy be adopted, the arrhythmia would be re-classified as 'long-standing persistent AF'.

IL PAZIENTE

IL COGNATO



FA parossistica FA persistente

Camm & Obel - Am J Cardiol 1996; 78: 3-11

Come si presenta

Studio ALFA

Sintomi	Popolazione totale % (n=756)	F A parossistica % (n=167)	F A cronica % (n=389)	FA di recente insorgenza % (n=200)
Palpitazioni	54,1	79,0	44,7	51,5
Dolore toracico	10,1	13,2	8,2	11,0
Dispnea	44,4	22,8	46,8	58,0
Sincope	10,4	17,4	8,0	9,5
Affaticabilità	14,3	12,6	13,1	18,0
Altro	0,9	0	1,8	0
lessuno	11,4	5,4	16,2	7,0



TIVI PER CARDIOLOGICA VALUTAZIONE URGENTE

GIANNI? FRANCO?

•		• • •	
ncal	cond	ifions	
in Cua	Conta		

Haemodynamic instability

Uncontrollable rate

Symptomatic bradycardia not amenable to reduction of rate control agents

Severe angina or worsening left ventricular function

Transient ischaemic attack or stroke

Cosa fare.

- A 12-lead ECG is recommended to establish a suspected diagnosis of AF, to determine rate in AF, and to screen for conduction defects, ischaemia, and signs of structural heart disease.
- Initial blood tests should evaluate thyroid and kidney function, as well as serum electrolytes and full blood count.
- Transthoracic echocardiography should be used to identify structural disease (e.g. valvular disease) and assess LV size and function (systolic and diastolic), atrial size, and right heart function.

GIANNI

- -Ventricolo sx non dilatato.
- -EF normale
- -Atrio sx ai limiti alti della norma
- -Non IM

FRANCO

- -Ventricolo sx non dilatato.
- -EF 50%
- -Atrio sx moderatamente dilatato -IM moderata

Rate or rhythm control

The acute management of patients with AF is driven by acute protection against thromboembolic events and acute improvement of cardiac function.

The severity of AF-related symptoms should drive the decision for acute restoration of sinus rhythm (in severely compromised patients) or acute management of the ventricular rate (in most other patients).



European Heart Journal (2010) **31**, 2369–2429 doi:10.1093/eurheartj/ehg278

Rate control should be the initial approach in elderly patients with AF and minor symptoms (EHRA score 1).	Т
Rate control should be continued throughout a rhythm control approach to ensure adequate control of the ventricular rate during recurrences of AF.	I
Rhythm control is recommended in patients with symptomatic (EHRA score <u>>2</u>) AF despite adequate rate control.	I
Rhythm control in patients with AF and AF-related heart failure should be considered for improvement of symptoms.	lla
Rhythm control as an initial approach should be considered in young symptomatic patients in whom catheter ablation treatment has not been ruled out.	lla

EUROPEAN SOCIETY OF CARDIOLOGY®

European Heart Journal (2010) **31**, 2369–2429 doi:10.1093/eurheartj/ehq278



Rate or Rhythm control

FAVOURING RHYTHM CONTROL
Paroxysmal AF or newly detected AF
More symptomatic
Age < 65 y
No hypertension
HF clearly exacerbated by AF
No previous failure of antiarrhythmic drug
Patient preference

Canadian CardioVascular Society – Atrial Fibrillation Guidelines 2010: rate and rhythm management

REVIEW

Rate versus rhythm control in atrial fibrillation and clinical outcomes: Updated systematic review and meta-analysis of randomized controlled trials

	Rate c		Rhythm		ART ALLER	Risk ratio				sk ra		
Study or subgroup	Events	Iotal	Events	lotal	Weight	M-H, Random, 95% CI	rear		м-н, ка	ndon	n, 95% Cl	
PIAF	2	125	2	127	0.3%	1.02 [0.15, 7.10]	2000		<u>.</u>	+		
AFFIRM	310	2027	356	2033	52.1%	0.87 [0.76, 1.00]	2002					
RACE	21	256	19	266	2.8%	1.15 [0.63, 2.09]	2002			+	-3	
STAF	8	100	4	100	0.7%	2.00 [0.62, 6.43]	2003		2	+	- K-	
HOT CAFE	1	101	3	104	0.2%	0.34 [0.04, 3.25]	2004		08.	+		
AF-CHF	228	694	217	682	43,2%	1,03 [0,89, 1,20]	2008			ŧ		
J-RHYTHM	3	404	4	419	0.5%	0.78 [0.18, 3.45]	2009		74	-		
CAFE-II	1	31	1	30	0.1%	0.97 [0.06, 14.78]	2009	03		+		-
Total (95% CI)	37	38	37	51	100.0%	0.95 [0.86, 1.05]				4		
Total events	574		606							1		
Heterogeneity : Tau ² :	= 0.00 ; C	hi² = 3.5	7, df = 7 (P	= 0.61);	² = 0%			.		+		- 1
Test for overall effect	: Z = 0.99) (P= 0.3	2)					0.05	0.2 Favours rate control	1	5 Favours rhythm contro	20 x i

Forest plot for all-cause mortality

Caldeira D. Archives of Cardiovasc Dis 2012; 105: 226-238



Antiarrhythmics for maintaining sinus rhythm after cardioversion of atrial fibrillation (Review)

Lafuente-Lafuente C, Valembois L, Bergmann JF, Belmin J

Several class IC (flecainide, propafenone) and III (amiodarone, dronedarone, sotalol) drugs significantly reduced recurrence of atrial fibrillation (OR 0.19 to 0.70, number needed to treat to beneft (NNTB) 3 to 16).

Betablockers (metoprolol) also significantly reduced atrial fibrillation recurrences (OR 0.62, 95% CI 0.44 to 0.88, NNTB 9).

Compared with controls, class IA drugs and sotalol were associated with increased all-cause mortality. Other antiarrhythmics did not seem to modify mortality.

All analysed drugs increased withdrawals due to adverse affects and all but amiodarone, dronedarone and propafenone increased proarrhythmia.

> Antiarrhythmics for maintaining sinus rhythm after cardioversion of atrial fibrillation (Review) Copyright © 2015 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.



Efficacia e Tollerabilità



rmaco	Metabolismo; dose	Tossicità non cardiovascolare	Tossicità cardiovascolare
pafenone	Epatico;150-300 x 3/die; rilascio prolung 225-425 x 2/die	ato Sapore metallico, vertigini	Flutter atriale 1:1, TV, smascheramento ST sopra in Brugada, non in ischemia miocardica
cainide	Renale/epatico CYP2D6. 50-100 x 2 (max 300-400/24h)	Vertigini, cefalea, visione sfocata	Flutter atriale 1:1, TV, smascheramento ST sopra in Brugada, non in ischemia miocardica
alolo	Renale. 80-120 x 2 (max 240 x 2)	Broncospasmo	Bradicardia, torsioni di punta
fetilide lo USA)	Renale/epatico CYP3A4; dose variabile p funzione renale (500-125 µg x 2)	er Nessuno	Torsioni di punta
iodarone	Epatico. Tempo di dimezzamento 50 gior Carico 10 g in 7-10 giorni, poi 400 mg p 3 settimane, poi 200 mg se FA. Dose rido se >QT, bradicardia. 150-300 bolo e.v., p 1 mg/min per 6h, poi 0.5 mg/min mantenimento	per ipersensibilità, infiltrati cronici otta interstiziali), epatite,	Bradicardia sinusale
tilide (e.v.)	Epatico CYP3A4. 1 mg e.v. in 10 min, ripetibile dopo 10 min	Nausea	Torsioni di punta
onedarone	Renale/epatico/gastrointestinale. 400 mg	x 2 Anoressia, nausea, epatotossicità	Bradicardia

Modificato da Zimetbaum P. Circulation 2012;125:381-9.



Canadian Journal of Cardiology 33 (2017) 965e976

Review

Contemporary Atrial Fibrillation Management: A Comparison of the Current AHA/ACC/HRS, CCS, and ESC Guidelines

Jason G. Andrade, MD,^{a,b} Laurent Macle, MD,^b Stanley Nattel, MD,^b

Atul Verma, MD,^c and John Cairns, MD

The recommendations of the ACC/AHA/HRS, CCS, and ESC are in general agreement, with each providing a strong recommendation for AF ablation for paroxysmal AF patients in whom an AAD has failed (strong recommendation for CCS, grade I for ESC, and ACCF/AHA/HRS).





Bisoprolol 1.25-20 mg once daily or split. Most common reported adverse symptoms are lethargy, headache, Carvedilol 3.125-50 mg twice daily. peripheral oedema, upper Metoprolol 100-200 mg total daily dose respiratory tract symptoms, (according to preparation). gastrointestinal upset and dizziness.Adverse effects include Nebivolol 2.5-10 mg once daily or split. bradycardia, atrioventricular block Esmolol and hypotension. Calcium-channe 60 mg 3 times daily up to Most common reported adverse Diltiazem 360 mg total daily dose symptoms are dizziness, malaise, (120-360 mg once daily lethargy, headache, hot flushes, gastrointestinal upset and modified release). oedema. Adverse effects include Verapamil 40-120 mg 3 times daily bradycardia, atrioventricular block (120-480 mg once daily and hypotension (prolonged modified release). hypotension possible with verapamil). Cardiac glycosid 0.0625-0.25 mg daily dose Most common reported adverse Digoxin symptoms are gastrointestinal upset, dizziness, blurred vision, headache and rash. In toxic states (serum levels >2 ng/ mL), digoxin is proarrhythmic and can aggravate heart failure, particularly with Digitoxin 0.05-0.3 mg daily dose. co-existent hypokalaemia. Specific indicatic 200 mg daily Hypotension, bradycardia, nausea, Amiodarone QT prolongation, pulmonary toxicity, skin discolouration, thyroid dysfunction, corneal deposits and cutaneous reaction with extravasation.

ate control therapy



Risk factor-based approach expressed as a point based scoring system, with the
acronym CHA ₂ DS ₂ -VASc

(NOTE: maximum score is 9 since age may contribute 0, 1, or 2 points)

Risk f	Score				
Congestive heart failure/LV dysfunct	1				
Hypertension		1			
Age ≥75		2			
Diabetes mellitus		1			
Stroke/TIA/thrombo-embolism		2			
Vascular disease*		1			
Age 65 to 74		1			
Sex category (ie, female sex)		1			
Maximum score	9				
Adjusted stroke rate according to CHA ₂ DS ₂ -VASc score					
CHA ₂ DS ₂ -VASc score	Patients (n = 7329)	Adjusted stroke rate (percent/year)*			
CHA ₂ DS ₂ -VASc score					
	(n = 7329)	(percent/year)•			
0	(n = 7329) 1	(percent/year)* 0 percent			
0	(n = 7329) 1 422	(percent/year)* 0 percent 1.3 percent			
0 1 2	(n = 7329) 1 422 1230	(percent/year)* 0 percent 1.3 percent 2.2 percent			
0 1 2 3	(n = 7329) 1 422 1230 1730	(percent/year)* 0 percent 1.3 percent 2.2 percent 3.2 percent			
0 1 2 3 4	(n = 7329) 1 422 1230 1730 1718	(percent/year)* 0 percent 1.3 percent 2.2 percent 3.2 percent 4.0 percent			
0 1 2 3 4 5	(n = 7329) 1 422 1230 1730 1718 1159	(percent/year)* 0 percent 1.3 percent 2.2 percent 3.2 percent 4.0 percent 6.7 percent			
0 1 2 3 4 5 6	(n = 7329) 1 422 1230 1730 1718 1159 679	(percent/year)*0 percent1.3 percent2.2 percent3.2 percent4.0 percent6.7 percent9.8 percent			

A TERAPIA ANTICOAGULANTE a scelta difficile



mparison of the efficacy and safety of new oral anticoagulants th warfarin in patients with atrial fibrillation: a meta-analysis of ndomised trials

42 411 participants received a new oral anticoagulant and 29 272 participants received warfarin

comparative efficacy of high-dose of NOACS and warfarin. Allocation to a new oral anticoagulant significantly reduced the composite of stroke or systemic embolic events by 19% compared with warfarin

NOAC (events) Warfarin (events) RR (95% CI) р E-LY⁵* 134/6076 199/6022 0.66 (0.53-0.82) 0.0001 OCKET AF6† 269/7081 306/7090 0.88 (0.75-1.03) 0.12 RISTOTLE7[‡] 265/9081 0.80 (0.67-0.95) 212/9120 0.012 NGAGE AF-TIMI 488 296/7035 337/7036 0.88 (0.75-1.02) 0.10 ombined (random) 0.81 (0.73-0.91) 911/29312 1107/29229 <0.0001 0.5 1.0 2.0 Favours NOAC Favours warfarin

C.T Ruff Dr Lancet, The, 2014-03-15



his study showed clear avid

that the outcomes of ruptured abdominal aortic aneurysm in Ecoland are score than in the US





benefit was mainly driven by a large reduction in haemorrhagic stroke.

v oral anticoagulants were also associated with a significant reduction in all-cause mortality

drugs were similar to warfarin in the prevention of ischaemic stroke and myocardial infarction



C.T Ruff Dr Lancet, The, 2014-03-15

Growing body of real-world experience from >650 000 patients



Reversal of dabigatran-associated bleeding using idarucizumab: review of the current evidence

Elective surgery

DICATION FOR USE OF THE ANTIDOTES

favor Against

fe-threatening bleeding (intracranial bleeding or uncontrollable hemorrhage)

leeding in a closed space or critical organ

ersistent major bleeding despite local hemostatic measures (or risk of recurrent bleeding because of delayed DOAC clearance or DOAC overdose)

eed for urgent intervention that is associated with a high risk of bleeding that cannot be delayed to allow for drug clearance

mergency surgery or intervention in patients at high risk for procedural bleeding

Gastrointestinal bleeds that respond to supportive measures

dTT (s)

High drug levels or excessive anticoagulation without associated bleeding

Need for surgery or intervention that can be delayed long enough to permit drug clearance



el mondo reale

studi di fase 3 possono essere tra loro non confrontabili. Idi 1:1 confronto diretto.

nsiderare il dosaggio adeguato al paziente e le interferenze farmacologiche

RE-LY ^a	ROCKET AF ^b	ARISTOTLE ^c	ENGAGE-AF ^d
Dabigatran	Rivaroxaban	Apixaban	Edoxaban *
 None US Regulators CrCl 15-30 mL/min: 75 mg BID Age > 80 years CrCl 30-50 mL/min + P-gp inhibitor, dronedarone, or ketoconazole 	 20 → 15 mg OD for Creatinine clearance < 30-49 mL/min 	 5 → 2.5 mg BID for ANY TWO of -Age ≥ 80 years -Body weight ≤ 60 kg -Serum creatinine ≥ 15 mg/dL US Regulators - Strong dual inhibitors of CYP3A4 and P-gp 	 60 → 30 mg OD or 30 → 15 mg OD for Creatinine clearance 30-50 mL/min Body weight ≤ 60 kg Use of quinidine, verapamil, or dronedarone

ACO/AIAC/SICI-GISE/SIC/SICCH Consensus Document: cutaneous occlusion of the left atrial appendage in -valvular atrial fibrillation patients: indications, ent selection, staff skills, organisation, and training

in Heart Journal Supplements (2017) 19 Supplement D

A/EAPCI44

alternative to OAT in patients intolerant of OAT ints with high risk of stroke and high risk of haemorrhage ints with thromboembolic events during OAT in the rapeutic range or during treatment with NOACs (when other origin of the bleeding can be identified) ints who can be treated with oral anticoagulants but where indication for left atrial appendage occlusion



- patients with non-valvular AF with high-thromboembolic risk and high-haemorrhagic risk (HAS-BLED ≥ 3);
- patients requiring triple antithrombotic therapy indefinitely;
- patients with tumours with increased risk of haemorrhage, underestimated by the HAS-BLED score;
- patients in whom OAT is ineffective in providing protection against cerebral ischaemic events probably correlated to thromboembolisms originating from the LAA;
- patients with kidney failure or undergoing dialysis, bearing in mind that all NOACs are contraindicated with creatinine clearance < 15 mL/min and that in these patients warfarin could increase tissue calcifica-

tion and the degree of atherosclerosis;

- patients with major bleeding of the urogenital or gastrointestinal system, or any other districts, such as the ocular area;
- frail patients (the very old, dementia, neurodegenerative diseases, malnutrition, etc.);
- patients with difficulty in managing oral therapies (e.g. mental illnesses, vision impairment); and
- patients who, after being suitably informed about the OAT/NOACs therapy, refuse it and demand a 'definitive' therapy. In this context, it should be underlined that the Watchman has had approval by the US regulatory authority as a valid alternative to warfarin in patients who refuse or prefer not to take OAT.

GIANNI E FRANCO VISSERO FELICI E CONTENTI GIANNI E FRANCO VISSERO FELICI E CONTENTI CON LA FIBRILLAZIONE ATRIALE.



Grazie per l'attenzione