Catheter Ablation of Atrial Fibrillation in the Elderly

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Projected number of persons with AF in the US assuming no further increase in age adjusted AF incidence and assuming a continued increase in incidence rate as evident in 1980 to 2000 (dotted curve) Study assessed community based trends in AF incidence from 1980-2000 and provided prevalence projections to 2050 (in Olmstead county)





Current Prob Cardiol, May 06 references 1 – 4, economic burden reference 5



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AF begets AF

> Animal models have demonstrated that the more frequently AF is induced and the longer it lasts, the more easily inducible and more permanent AF becomes (Wijffels MCEF. Circulation 1995)

> 25-40% of patients with PAF will progress to chronic AF over a 10-year period



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Management of Atrial Fibrillation

- > 1) Rate control and anti-coagulation
- > 2) Rhythm control with anti-arrhythmic agents (AADs) and anti-coagulation
- > 3) AVJ ablation and pacemaker placement
- > 4) Atrial fibrillation ablation





TABLE 1. BASE-LINE CHARACTERISTICS OF THE PATIENTS.*				
Characteristic	OVERALL (N=4060)	RATE-CONTROL GROUP (N=2027)	RHYTHM-CONTROL GROUP (N=2033)	P Value
Age — yr	69.7±9.0	69.8±8.9	69.7±9.0	0.82
Female sex — no. (%)	1594 (39.3)	823 (40.6)	771 (37.9)	0.08
Ethnic minority group - no. (%)	461 (11.4)	241 (11.9)	220 (10.8)	0.28
Predominant cardiac diagnosis — no. (%)	()	()	()	0.29
Coronary artery disease Cardiomyopathy Hypertension Valvular disease	1059 (26.1) 194 (4.8) 2063 (50.8) 198 (4.9)	497 (24.5) 99 (4.9) 1045 (51.6) 98 (4.8)	562 (27.6) 95 (4.7) 1018 (50.1) 100 (4.9)	
Other No apparent heart disease	42 (1.0) 504 (12.4)	23 (1.1) 265 (13.1)	$ 19 (0.9) \\ 239 (11.8) $	
History of congestive heart failure — no. (%)	939 (23.1)	475 (23.4)	464 (22.8)	0.64
Duration of qualifying atrial fibrillation ≥2 days — no. (%)	2808 (69.2)	1406 (69.4)	1402 (69.0)	0.80
First episode of atrial fibrillation (vs. recurrent episode) — no. (%)†	1391 (35.5)	700 (35.8)	691 (35.3)	0.74
Any prerandomization failure of an antiarrhythmic drug — no. (%)	713 (17.6)	364 (18.0)	349 (17.2)	0.51
Size of left atrium normal - no. (%)‡	1103 (35.3)	549 (35.3)	554 (35.3)	0.98
Left ventricular ejection fraction — %§	54.7±13.5	54.9±13.1	54.6±13.8	0.74
Normal left ventricular ejection fraction — no. (%)‡	2244 (74.0)	1131 (74.9)	1113 (73.2)	0.29



- > Did AFFIRM actually study rate control versus rhythm control in an elderly population?
- > Was there truly equivalence between the two strategies?

rate control. Radiofrequency ablation to modify or eliminate atrioventricular conduction was used in 105 (5.2 percent) of the patients in the rate-control group after drug failure. During the course of the study, 248 patients crossed over from the rate-control group to the rhythm-control group (actuarial rate of crossover, 7.8 percent, 11.6 percent, and 14.9 percent after one, three, and five years, respectively). Eighty-six of these patients had crossed back to the rate-control group by the end of the study. Uncontrolled symptoms due to atrial fibrillation and congestive heart failure were the most common reasons for the initial crossover to rhythm control in this group.

N Engl J Med, Vol. 347, No. 23 · December 5, 2002

The down side of rate control

- Rate control with AV nodal blocking agents was not satisfactory in 20% of patients
- > 5.2% required AVJ ablation with pacemaker placement
 - Invasive strategy making patients 100% pacemaker dependent
- > 14.9% (actuarial) crossover to AAD's at 5 years





On treatment analysis which was only published recently, presence of sinus rhythm was one of the most powerful independent predictors of survival along with use of warfarin. Patients in sinus rhythm wee half as likely to die than those with AF. This benefit however was offset by the use of AAD which increased the risk of death.

(Reduction in mortality with sinus rhythm has also been shown in DIAMOND and CHF-STAT trials.





Pharmacologic Approaches to Maintain Sinus Rhythm

- Low success rate
 - 1-year AF recurrence of approximately 50%
 - Regardless of anti-arrhythmic agent
 - Amiodarone best agent <u>~</u> 65% SR at 16 months
 - Side effects common
 - Amiodarone
 - Attrition rate of therapy due to many side effects
 - 18% at 16 months in CTAF trial
 - 11% at one year AFFIRM trial
 - Risk of pro-arrhythmia

Am J Cardiol 1991;68:335-41 NEJM 2000;342:913-20 Am J Geriatr Cardiol, 2002; 11: 370–375 Heart Rhythm, 2007; 4: 1577–1599.

The Elderly patient and AAD's

> Age related senescence alters the pharmacokinetics of AADs

- metabolism less predictable
- · increased predilection for side effects
- · Increased risk of pro-arrhythmias

Dayer MB, Hardman SMC, Special problems with antiarrhythmic drugs in the elderly: Safety, tolerability, and efficacy. Am JGenatr Cardiol, 2002; 11: 370–375. Curds AB, Rich MW, Atrial fibrillation in the elderly: Mechanisms and management. Heart Rhythm. 2007; 4: 1377–1399. Fang MC, Chen J, Rich MW, Atrial fibrillation in the elderly. AmJ Med, 2007; 120: 481–487. Meta-analysis of 1100 patients after AV junctional ablation and pacemaker placement for medically refractory atrial fibrillation, atrial flutter or atrial tachycardia: left ventricular function, healthcare use, and New York Heart Association (NYHA) functional classification



Catheter Ablation of the AV Junction

- > Advantages
 - Improved rate control
 - Improved QOL
 - Improved LVEF
 - No AADs
 - Less hospitalizations
- > Disadvantages
 - Pacemaker
 dependence
 - Procedure
 complications
 - Continued embolic risk
 - Progression of AF to more permanent





- Figure 1. Diagram of the Sites of 69 Foci Triggering Atrial Fibrillation in 45 Patients. Note the clustering in the pulmonary veins, particularly in both superior pulmonary veins. Numbers indicate the distribution of foci in the pulmonary veins.
- A single point of origin of ectopic beats was identified in 29 patients, two were identified in 9 patients, three were identified in 6 paitents and four in 1 patient. The venous confirmation of earliest ectopic activity was demonstrated in 23 patients by the radiographic position of the mapping catheter, which was superimposed on the lungs and was outside the cardiac silhouette and by confirationtory angiographic visuatization. The conduction time from the venous spike to the activation of the left atrium increased progressively as the coupling interval of the spike spontaneously shortened and when there were repetitive spike discharges.

Important Insights into Pulmonary Vein Architecture







AF is initiated by rapid discharges from one or several focal sources within the atria. 94% arise from PV. Extra PV sites may trigger AF in 6 – 10%. Rotors are high frequency micro reentrant circuits that perpetuate AF. Conduction becomes slower and less organized with increasing distance from rotors because of atrial structural remodeling resulting in fibrillatory conduction.. Interestingly the dominant rotors in AF are localized primarily in the junction between LA and PVs. Further vagal inputs may be important in both triggering and maintaining AF. We will discuss this later.



Figure 4. Similarity in location of the radiofrequency lesions produced by various groups' approaches to AF ablation. Top left, An outer view of a patient's LA as seen from the posterior aspect using 3D, multislice CT. Seen clearly are the tubular portions of each of the 4 PVs (individually labeled). The borders between the antra of the PVs and the posterior wall of the LA are indicated by small white arrows. Bottom left, A 3D electroanatomic map (CARTO, Biosense Webster Inc) of the LA (same patient as panel above) acquired during AF ablation guided by ICE. With ICE, the borders of the pulmonary venous antra can be accurately defined, and lesions can be placed to completely surround and electrically isolate the antra. Red dots represent the anatomic locations of these lesions produced by ICE-guided ablation. Top left, Location of lesions produced with a CARTO-guided approach described by Morady and colleagues. Reproduced from Oral et al,40 with permission. Bottom left, Location of lesions produced using another CARTO-guided approach described by Pappone and colleagues. Reproduced from Pappone et al,29 with permission. In all 3 cases, the location of the lesion sets is similar, encompassing the anterior and posterior borders of all 4 pulmonary venous antra. LSPV indicates left superior PV; LIPV, left inferior PV; RSPV, right superior PV; and RIPV, right inferior PV.





APAF

Trial Design: APAF was a randomized trial circumferential pulmonary vein ablation (CPVA) (n=99) compared with antiarrhythmic medical therapy (n=99) with flecainide (n=33), sotalol (n=33) or amiodarone (n=33) among patients with paroxysmal atrial fibrillation. Primary endpoint was freedom from recurrent atrial arrhythmias at 1 year.



Results

- Only preliminary data available in 150 patients At 9 months follow-up, freedom from recurrent AF and AT 1 in CPVA patients vs medical therapy (Figure)
 Of 8 patients who had recurrent AF in CPVA group,
- repeat ablation procedure performed in 3 patients. 1 of whom still had additional recurrent AF
- Of 52 patients in medical therapy group who had recurrent AF, 38 had CPVA performed, 4 of whom still had additional recurrent AF
- Significant decrease in left atrium diameter at 12 months in CPVA group (P<0.05) but no difference in medical therapy group
- Conclusions Among patients with paroxysmal atrial fibrillation, treatment with circumferential pulmonary vein ablation was associated with reduction in recurrent AF and AT compared with conventional antiarrhythmic medical therapy • Present trial one of first randomized trials of CPVA
- in paroxysmal atrial fibrillation
- · CPVA recently shown to be beneficial in maintaining sinus rhythm in chronic atrial fibrillation
- Presented at ACC 2006


Radiofrequency Ablation vs Antiarrhythmic Drugs as First-line Treatment of Symptomatic Atrial Fibrillation A Randomized Trial

Characteristics	Pulmonary Vein Isolation Group (n = 33)	Antiarrhythmic Drug Group (n = 37)
Age, mean (SD), y	53 (8)	54 (8)
Left atrial size, mean (SD), cm	4.1 (0.8)	4.2 (0.7)
Duration of atrial fibrillation, mean (SD), mo	5 (2.0)	5 (2.5)
Atrial fibrillation Paroxysmal	32 (97)	35 (95)
Persistent	1 (3)	2 (5)
Structural heart disease and hypertension	8 (25)	10 (28)
Left ventricular ejection fraction, mean (SD), %	53 (5)	54 (6)
Use of β-blocker therapy	19 (57)	23 (62)
*Data are presented as No. (%) unless otherwise specified.		
		JAMA. 2005;293:2634-2640





There is a disconnect between the AF ablation population and the real AF population!!!





Characteristics of Patients Undergoing Atrial Fibrillation Ablation: Trends Over a Seven-Year Period 1999–2005

EDWARD P. GERSTENFELD, M.D., DAVID CALLANS, M.D., SANJAY DIXIT, M.D., DAVID LIN, M.D., JOSHUA COOPER, M.D., ANDREA M. RUSSO, M.D., RALPH VERDINO, M.D., MARK WEINER, M.D., ERICA ZADO, P.A.C., and FRANCIS E. MARCHLINSKI, M.D.

TABLE 2 Patient Characteristics by Year 1999 2000 2001 2002 2003 2004 2005 Characteristic (N = 29) (N = 109) (N = 187) (N = 200) (N = 2005 Characteristic (N = 29) (N = 109) (N = 187) (N = 200) (N = 221) (N = 265) P								
								Р
Age (years)	47.1	52.4	53.5	55.1	54.8	56.8	55.7	< 0.01
Gender (% male)	90	68	80	77	79	75	77	NS
Prior AF duration (years)	7.6	7.7	5.7	7.6	7.1	6.7	7.4	NS
# Prior AA drugs	3.9	3.6	3.3	3.0	3.1	2.6	2.0	< 0.0
Left atrial size (cm)	4.0	4.2	4.1	4.5	4.4	4.5	4.4	< 0.0
LV ejection fraction (%)	55	54	56	60	59	59	58	NS
Prior stroke/TIA (%)	3.4	2.1	3.6	3.7	6	6.2	10	NS
Prior cardiomyopathy (%)	0	0	0.9	5.3	5.5	7.7	16	< 0.0
Obstructive sleep apnea (%)	10.3	12.8	9.2	7.5	10.5	11.8	12	NS
Body mass index	31.2	29.6	29.1	30.8	29.1	30.7	28.6	NS

J Cardiovasc Electrophysiol, 2007; 18: 23-38

AF ablation in the Elderly Review of the Literature

There are <u>no</u> prospective randomized controlled trials comparing the safety and efficacy of catheter ablation for paroxysmal or persistent AF in the elderly to best medical therapy or alternative strategies such as AV node ablation plus pacemaker placement





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TABLE 2 Procedural Acute Comp	lications
Number of procedures, n	194 (*)
Embolic TIA/stroke, n (%)	1 (0.5%)
Hemothorax, $n(\%)$	1(0.5%)
Groin hematomas, n (%) (*) 194 ablations = 174 first ablations + 20 s	3 (1.5%) second ablations.

Long-Term Clinical Efficacy and Risk of Catheter Ablation for Atrial Fibrillation in the Elderly

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From the Cardiovascular Division, Department of Medicine, Hospital of the University of Pennsylvania, Philadelphia, Pennsylvania, USA

Clinical Characteristics				
	<65 Years	65-74 Years	≥75 Years	P-Value
Number of patients	948	185	32	NS
Number of procedures	1,244	228	34	NS
Mean procedures/patient	1.3 ± 0.55	1.2 ± 0.47	1.1 ± 0.25	NS
Mean age (yrs)	52 ± 9	68 ± 3	77 ± 2	NS
Age range (yrs)	16 - 64	65 - 74	75 - 82	NS
#Women (%)	185 (20%)	62 (34%)	18 (56%)	< 0.00
#PAF (%)	614 (65%)	115 (62%)	17 (53%)	NS
LA size (cm)	4.4 ± 0.7	4.5 ± 0.7	4.4 ± 1.0	NS
#with LVEF <50% (%)	114 (12%)	13 (7%)	5 (16%)	NS
#HTN and/or SHD (%)	533 (56%)	126 (68%)	28 (88%)	< 0.001
$\#$ CHADS2 score ≥ 2 (%)	111 (12%)	30 (30%)	23 (72%)	< 0.001

 $LA = left atrial; LVEF = left ventricular ejection fraction; HTN = hypertension; PAF = paroxysmal atrial fibrillation; SHD = structural heart disease; CHADS2 = 1 point each for Congestive heart failure, Hypertension, Age <math>\geq$ 75, Diabetes, and 2 points for Stroke or transient ischemic attack.

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Catheter ablation of atrial fibrillation in the elderly: Where do we stand?

Darren Traub, James P. Daubert, Scott McNitt, Wojciech Zaręba, Burr Hall Cardiology Division, University of Rochester Medical Center, Rochester, NY, USA

Clinical variables	Age < 70 (n = 45)	Age ≥ 70 (n = 15)	P
Age	52 ± 11	74 ± 2	< 0.001
Female	29%	20%	0.738
Ejection fraction (%)	54 ± 8	57±6	0.105
Ejection fraction $\leq 45\%$	13%	7%	0.668
Left atrial size [mm]	41 ± 6	43±5	0.151
Left atrial size ≥ 50 mm	4%	13%	0.258
AF duration > 60 months	24%	40%	0.324
Comorbidites			
Coronary artery disease	13%	20%	0.678
Hypertension	42%	40%	1.000
Diabetes	4%	7%	1.000
Medications at baseline			
ACE-inhibitors	27%	40%	0.347
Beta-blockers	40%	87%	0.002
Statins	27%	40%	0.347
Flecainide	24%	13%	0.485
Tikosyn	11%	47%	0.006
Propafenone	33%	20%	0.517
Sotalol	18%	7%	0.427
Amiodarone	9%	0	0.564
None	4%	13%	0.258 (Cardiol J 2009; 16, 2

Catheter ablation of atrial fibrillation in the elderly: Where do we stand?

Darren Traub, James P. Daubert, Scott McNitt, Wojciech Zaręba, Burr Hall

Cardiology Division, University of Rochester Medical Center, Rochester, NY, USA

	Age $< 70 (n = 45)$	Age ≥ 70 (n = 15)	Р
NSR in the absence of symptoms at 12 months	36 (80.0%)	9 (60.0%)	0.169
NSR or symptomatic improvement	42 (93.3%)	12 (80.0%)	0.159
Remaining on anti-arrhythmic therapy (%)	7 (15.6%)	9 (60.0%)	0.002
With successful ablations who remained on anti-arrhythmic therapy (%)	3/36 (8%)	3/9 (33%)	< 0.05
Remaining on warfarin (%)	10 (22.2%)	12 (80.0%)	< 0.001
Complications	2 (4.4%)	1 (6.7%)	1.000
NSR — normal sinus rhythm			

(Cardiol J 2009; 16, 2: 113-120)

	Inc DAVID D. SPRAGG, M.D.,* DAI DANEL SCHERR, M.D.*,* CHARLES A. HEN RONALD D. BERR arr	theter Ablation for At idence and Predictors RSHAN DALAL, M.D., M.P.H., ' KARUNA CHILUKURI, M.D., ' KIKSON, M.D., 'JOSEPH E. MA IKSON, M.D., 'JOL PODG, J di HUGH CALKINS, M.D.' Middame. Alsen Bigen Unsersty is belof. Bepartness of Molece. Medical Usersity of	AAMIR CHEEMA, M.D.,* LAN CHENG, M.D.,* KINS, M.D.,* A.D., Ph.D.,* 4 Molicue, Balimore, Mayland, UNA;	
Clinical Characteristic	TABLE 1 s of Patients Undergoing A	Ablation of AF		
Age (y) Male gender (%) AF burden Paroxysmal (%) Persistent (%) Ejection fraction (%) Left atrial diameter (cm) Hypertension (%) Prior cardioversions Prior antiarthythmics		$57.2 \pm 11 \\77.7 \\53.6 \\46.4 \\57.0 \pm 9.2 \\4.7 \pm 0.7 \\42.0 \\1.3 \pm 2.0 \\1.6 \pm 0.9$		
		Major Compli	TABLE 2 cations in Patients Undergoing Abl.	ation of AF
I Cardiovasc Electrophysiol, Vol	. 19, pp. 627-631, June 2008.)	Stroke Tamponade Vascular injury Pulmonary vein occl Hemothorax Heart block Acute lung injury Mitral valve injury	usion	7 8 11 2 1 1 1





Can we use the same AF ablation strategies in younger and older patients?







CHANGE OF DEMOGRAPHICS, CLINICAL CHARACTERISTICS AND OUTCOMES IN PATIENTS REFERRED FOR ATRIAL FIBRILLATION ABLATION

Freddy Del-Carpio Munoz, MD, Gregory A. Cogert, MD, Kristi H. Monahan, RN, Janice M. Haroldson, RN, Laura A. Peterson, Yong M. Cha, MD, Paul A. Friedman, MD, Thomas M. Munger, MD, Samuel J. Asirvatham, MD and Douglas L. Packer, MD. Mayo Clinic, Rochester, MN

	Early Period (1998-2002) n=275	Intermediate Period (2003-2005) n=515	Early Period (2006-2008) n=624	p value
Age (mean ± SD)	52.1 ± 10	55.6 ± 11	58.1 ± 11	0.001
Number of failed AAs	2.5 ± 1.3	1.9 ± 1.3	2.5 ± 1.1	0.001
PAF (n, %)	193 (70%)	261 (51%)	227 (37%)	0.0001
Persistent AF (n, %)	70 (25%)	192 (37%)	318 (52%)	0.0001
Permanent AF (n, %)	12 (5%)	60 (12%)	67 (11%)	0.0002
Drganic Heart Disease n, %)	44 (16%)	126 (25%)	194 (31%)	0.0001
Normal LA size (n, %)	95 (35%)	161 (32%)	88 (16%)	0.0001
Repeat procedure (n, %)	15 (5%)	55 (11%)	80 (13%)	0.003
F Elimination/Control	48/59%	61/71%	69/82%	0.0001

Heart Rhythm Vol 6, No.5 May Supplement 2009

	Darren Ti	raub, James P. Dau	bert, Scott N	re do we stand IcNitt, Wojciech Zaręba, Medical Center, Rochester, N	Burr Hall	
able 3. Summ Author	Patients	fibrillation (AF) ablation AF type (paroxysmal,	Mean-follow	Outcome	Remaining on	Major
	(age range in years)	persistent, chronic)	(months)		AAD's	complication
Hseih et al. Corrado et al. (see Fig. 1)	37 (72 ± 4) 174 (> 75)	Paroxysmal Paroxysmal 55% Persistent 45%	52 ± 6 20 ± 14	81% in SR 88% (127/143) SR off AAD's after 1 th procedure 10.8% (16/143) SR off AAD's after 2 nd procedure	11% 13%	0%
Zado et al. (see Fig. 2)	635 (< 65) 124 (65–74) 22 (≥ 75)	65% paroxysmal** 62% paroxysmal 53% paroxysmal	27.6 ± 13.8 27.7 ± 13.6 23.8 ± 11.3	89% AF control* 84% AF control* 87% AF control*	20% 29% 37%	1.6% 1.7% 2.9%
Hall et al.	15 (73.6)	100% paroxysmal	56	60% in SR at 12 months 80% in SR or with symptomatic improvement	60%	6.6%
Santinelli et al.	172 (> 80, mean age 83 ± 2)	35% paroxysmal 29% persistent 36% paroxysmal	18 ± 5	Overall success rate of 75% 90% paroxysmal 76% persistent 60% permanent	Not reported	5%

I

Conclusion

- Catheter ablation of AF can be safely performed in an elderly population.
 - There may be a slightly increased risk of peri-procedural complications
- Catheter ablation appears to be efficacious in an elderly population
 - All data based on retrospective analyses
 - More elderly remain on AADs post-ablation
 This may be an acceptable endpoint for an elderly patient with
 symptomatic AF
- The next step is to perform large scale, randomized trials like AFFIRM to more definitively establish the role of catheter ablation in treating elderly patients with atrial fibrillation