#### Case 1: 62-Year-Old with Persistent AF with Cardiomyopathy and EF of 32%



#### Mélèze HOCINI

Cardiac Electrophysiology and Modeling Institute LIRYC – University of Bordeaux

## **CASE PRESENTATION**

- 63 years old woman
- Former smoker
- Rhythmic cardiomyopathy discovered in December 2018
  - NYHA III
  - TTE: LVEF 33% w/ global hypokinesia, mild RV hypokinesia, no valvular disease.
  - Coronary angiography: normal
  - Cardiac MRI: LVEF 30%, no gadolinium late enhancement
  - BNP: 626 pg/mL
  - 1 CHF episodes in the last 6 Mo

#### Therapeutic management

- Apixaban 5mg bd
- B-bloquers + ACE inhibitors + Diuretics
- AF
  - History of AF for 5 years
  - Persistent AF for 12 months
  - Amiodarone failed



		1.1.1.1	••••	. j				1	••••		( · · · ·	• • • •	1001		<u> </u>			••••		(		1	• • • • • • •				···· ] ;		1		<u> //</u>			11		- <b>1</b>		
		مسببهم			have	~~~.		h		: hermany	mar	ليسه	 سبيه	****	han		/}		÷,		******	June	:: جورمنده		: 	إلىب	4	لىورىمە	hin	مر دیا می	EL	بمسم	: 4.4	لمسالم		-Ali-	: جىسمىد	: :
:11:	:				:: ::				::::	:		: :::				:		::::	:	1	:::::	1	::: ::::: :		::::		::::				~~~~	<u>/</u>	:				: :::: :	:
	:		.				11	ĥ.	ما . ا	: :	ĥ÷.	~			⊫  }: .		ΙÅ		: ;	::  \ :: _		Å	:: : ::	- A		.			1		27		:	1		k		: ::
		::::::	•	The second s	1. a.a.	77	-	1	1	$\uparrow \uparrow$	( free	1	الرينينية 		معمدنة ( 111111)			**************************************		1 100000	**************************************	() :N.,			····	:::::::	******		مند الم المنالية	· · · · · · · · · · · · · · · · · · ·	11	.); 				~~		
. 111	:									:	1				J.	:			:			1		<u> </u>		1					17		:					
		him		in y	مسها	-in-ra	~~~	have	<u>~~</u>	in	Juni	$\sim \gamma$	مينيها	~~~ <b>~</b> ~	lhim	uman	mγ		نمسم	him		Yhne		-n/r	~		m.	~~~	l		Ëf	ښې ا	n	1		-16-	~~~~	
av	R									::::::		: :::																					: ::::					
~~~		ä	,		,	منعم		m	ممر	: بيمية	, min	<u> </u>	<u> </u>	يسررته وارد	ŝ	بىمىسى	-		: پېرىمىن	1 mar		:: 17			~~~		$\sim$		vär	<b></b>	h	يبسر	: •••••••	1 de		-		
a∖	L.							1							1		¥					1				···· {					1			1		1		
				Ĵ				Å			1	. İ			li.		LÌ.			λü.		I				. /			1		£					J.		
		1									1		<u> </u>		1			••••				1				1			<b>F</b>		<u> </u>	$\int$		1				
av	1 <del>F</del>									1					l:				:					A		A					ii /	1 :	1			ŀ	-	
	-			man	مسعبه					~~~	hun	لابد	ميبيه	المنتخذ	-	~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~	ten (			low		1	*****			rend.		~~~·`	1	ج	i (		-	h		alter	~~~~	
V1	-									-									:												1	1	ł					d i
÷	ليسببه	$\sim$	منطلح		$\sim$		سم	1	2.00	أستنج		إنبه	~	استنجب	1	ليتبنين		~	لنحب	1		1/		~1	$\sim$		7	~~~	m		<u>}</u>	منبيها		1	منهبه	-10	-	÷
V2								Į.			1	Ì			16				-	1		1		- II					18		Į.	Ŧ						d.
		يعنذ	مرين	4				-	Sill			æ	min	ليتنق	1	أنشلت			لتعنينا	منتر <b>ا</b>	لينتك	من ا		A	أمتر	<b>.</b>	م	لنشنك	سنزلا	<b>N</b>	N			ليرار	ينتنك		nu i	i.
		1						1			V II				16		171			1		Ĭ.		ľ		-	1		Ye.		$\mathcal{V}$			W.		1		
										i												l								A		1						
	in and	1	معرد الإيد 	~~1		2.4		r	Par la	لسبح	1	$\sim$	1	محم	1	Landres		-		1		1	$\sim$	~//		~~	٢	المرسيمة	V	hered	1	يمر.		71	- And and a second	1		~~
V4		1								i					Æ					í ::		i			!	1	[]		£		11	1	: 				!	
 rm		مسب	 مریدامیده	-	بمسم	,		~~	****	m		ليسم		ليعرب	<b>.</b>			~~~	أسعمهم		****	-	~				,	لعيسي			17-	C		1		-	****	
V5		1			1			ľ		1	1	[			ľ.				1	1		ľ		ľ		- 4			1		ų.		1			ľ		:
::::	::::::						: :::		::::	:::::		: ::	:::::			:			:::::::::::::::::::::::::::::::::::::::		:::::		: :::		::::		:::		1	: :::: :	1	1					-	
			**********					<b>I</b> ~~	r			~~~~		~~~~	1 m	جمود المرجم :	t-M			1	****	*		~~{h	•••••†	~~~~	r	ليعمدهم	<b> </b>	~~~~~	r	~		1	~~~~	Abm		
Vt		1::::									(i :::	: :::			le :::		::: \			1						II					1	.:;		周			: :::: ;	
-	himm	Lin	-		ن ا			11	m		مننسا	~~~	مننيا	*	مسنال	manin	fund {		-	منذا	·		mi		اد		L.		Niin		M.L	mit		JUJ	man	mll.		

### **LESIONS SET: MARSHALL Plan for PsAF**





**Pambrun T**, Hocini M et al MARSHALL bundles elimination, PVI and Lines completion for ANatomical ablation of PsAF: JCE 2019 Jan;30(1):7-15

## **VOM VARIABILITY**



#### Injection 1 no OH

#### OH 3cc



- Long sheath
- IMA catheter
- 0,014 Whisper angioplasty wire
- 8x2.0 OTW angioplasty balloon

### **MARSHALL Plan for PsAF**







### **MARSHALL Plan for PsAF**



#### **Mitral isthmus Roof line**



## AF ablation: after full lesions set

AF Ablation Dec 22th 2018 Initial AF CL: 136 ms in LAA Cardioversion



## ECHOCARDIOGRAMME

• Dec 27 th post Abl, SR



LV EF: 32% (Simpson) MR : II to III

# FOLLOW UP

- Jan 2019: Recurrence of arrhythmia:
- Atypical Flutter
- No CHF episode
- Cardioversion



## Second procedure Feb 2019: Ablation of 1 AT from RSPV





## **FOLLOW UP AT 9 MONTHS**



@1mth LVEDD: 77 LVESD: 65 EF: 32%



@3mths LVEDD: 65 LVESD: 47 EF: 52%

# **AF begets HF: HF begets AF**



# 2017 HRS/EHRA/ECAS/APHRS/SOLAECE consensus statement on AF ablation in HF

Indications for AF ablation in populations of patients are not well represented in clinical trials

patients with HF as in	Congestive Heart Failure	It is reasonable to use similar indications for AF ablation in selected patients with HF as in patients without HF	lla
------------------------	-----------------------------	-----------------------------------------------------------------------------------------------------------------------------	-----

Calkins et al, Heart rhythm 2017

#### Registries evaluating AF Ablation vs medical ttt: Consistent improvement in outcomes

	AF RFA (n)	Mortality HR (95% CI)	Stroke HR (95% CI)	Follow-up (yrs)
Reynolds CCQO 2012	801	NA	.62 (.4486)	3
Chang CAE 2014	846	.88 (.62-1.23)	.57 (.3594)	3.5
Karasoy EHJ 2015	4050	NA	.53 (.4365)	3.4
Friberg EHJ 2016	2836	.5 (.3762)	.69 (.5193)	4.4
Saliba HR 2017	969	.57 (.4766)	.62 (.4782)	NA
Srivatsa CAE 2018	4169	.59 (.45–.77)	.68 (.4797)	3.6
Noseworthy EHJ 2019	6907	.60 (.53–.69)	.56 (.4373)	2.3

# AF ablation in CHF Pts: Impact on EF in controlled studies

	Pts n	RFA % Success	EF%	EF increase	Pe AF	RF	F/up (mths)
Khan NEJM 2008	40	88%	27%	+8%	51%	PVI+ vs AVJ+BiV	6
MacDonald Heart 2011	20	50%	36%	+4.5% (ns)	100%	Stepwise	6
Jones, JACC 2013	52	88%	21%	+10.9%	100%	Stepwise	12
Hunter CAE 2014	50	73%	32%	+8%	100%	Stepwise vs Rate	12
DiBiase Circ 2016	203	70%	29%	+8%	100%	Stepwise vs Amio	24
Prabhu JACC 2017	66	75%	32%	+18%	100%	PVI+Post' LA vs Rate	6
Marrouche NEJM 2018	263	50%	33%	+8%	70%	PVI+ vs Rate/Rhythm	60

#### AF ablation vs rate control for AF/CHF pts CAMTAF trial: 50 pts



Hunter et al, Circ AE 2014

CA vs rate control in AF an systolic dysfunction CAMERA-MRI: multicentre randomised trial

Aims:

- To determine whether
  - CA is superior to medical rate control in PsAF and idiopathic cardiomyopathy
  - The absence of ventricular fibrosis on cardiac MRI predicts LV recovery with CA

Prabhu et al, JACC 2017

#### **CAMERA-MRI: Baseline Characteristics**

Characteristics	Catheter ablation	Medical Rate
	(n=33)	Control (n=33)
Age (years)	59 ± 11	62 ± 9.4
Male (%)	94% (31)	88% (29)
CHA2DS2VASc score	2.42 ± 0.87	2.36 ± 0.96
LVEF (MRI)	32 ± 9.4%	34 ± 7.8%
Late gadolinium present (%)	36% (12)	36% (12)
Mean duration continuous AF(mths)	23 ± 18	21 ± 15
24hr average HR (bpm)	86 ± 14	85 ± 17
DCR attempts per patient	2.1 ± 0.8	2.0 ± 0.7
Amiodarone failed or C/I (%)	91% (30)	82% (27)
ACE inhibitor or ARB (%)	94% (31)	94% (31)
Beta-blocker (%)	97% (32)	97% (32)

#### **CAMERA-MRI: Primary Endpoint**



Mean difference=14% 95% CI(8,5%, 19,5%)

#### LV EF in CA groupe: LGE status



#### **Reverse atrial remodeling at 24 months**



Sugumar et al, JACC 2019

#### **Conclusions: CAMERA MRI**

- AF is an important under appreciated reversible cause of cardiomyopathy
  - Despite adequate rate control
- Restoration of SR with CA improves LVEF
- Absence of LGE on CMR identified "super responders'

#### Catheter ablation for AF and heart failure: CASTLE AF

• Hypothesis:

AF ablation improves mortality and hospitalization rates in pts with LV dysfunction & AF compared to conventional treatment

• Primary Endpoint:

Composite of death from any cause or heart failure hospitalization

Marrouche et al NEJM 2018

#### Catheter ablation for AF and heart failure: CASTLE AF

- Design :
  - Multicenter studies (33 centers EU, USA et AUS)
  - Randomisation 1:1
- Patients :
  - PAF or PsAF
  - NYHA II, III ou IV
  - LVEF< 35%
  - Indication for ICD due to primary prevention
  - Dual chamber ICD or CRT-D already implanted

#### **FLOW CHART: CASTLE AF**



#### **PATIENTS CHARACTERISTICS**

Characteristic	Treatment of A	trial Fibrillation					
	Ablation group	Pharmacological group		Initi	al Procedure*	Chronic Re (after bla of 1	peat Ablations Inking period 2 weeks)
New York Heart Association class	(179 patients)	(104 patients)		Ablation group	Pharmacological group	Ablation group	Pharmacolog
l – no. (%)	0 (0% vs. 11%*)	0 (0% vs. 11%*)		151 pts	18 pts	37 pts	5 pts
II – no. (%) III – no. (%) IV – no. (%)	107 (60% vs. 58%*) 66 (37% vs. 29%*) 6 (3% vs. 2%*)	107 (58% vs 61%*) 75 (41% vs. 27%*) 2 (1%)	Time of initial procedure since baseline – days Mean ± SD Median (IQR) Minimum-maximum	6 ± 24 1 (0-1) 0-256 <sup>†</sup>	268 ± 270 212 (62-372) 0-1107		
<sup>†</sup> Body-mass index – kg/m <sup>2</sup> <sup>†</sup> Left ventricular ejection fraction – %	28.7 (25.7-32.3) 29.0 (25.0-32.0)	29.1 (25.8-32.2) 30.0 (25.0-32.0)	Time between initial procedure and 1 <sup>st</sup> repeat ablation – days				
Medication ACE-inhibitor or ARB – no. (%) Beta-blocker – no. (%)	n=179 168 (94%) 164 (92%)	n=183 166 (91%) 174 (95%)	Mean ± SD Median (IQR) Minimum-maximum			427 ± 354 336 (134-630) 95-1345	233 ± 133 260 (111-273 98-421
Diuretics including spironolactone – no. (%) Digitalis – no. (%) Antiarrhythmic drug (class Ia, Ic, or III) – no. (%) Amiodarone – no. (%)	170 (95%, vs. 93%*) 36 (20%, vs. 18%*) <sup>‡</sup> 51 (29%, vs. 32%*) 50 (28%, vs. 31%*)	168 (92%, vs. 93%*) 56 (31%) <sup>‡</sup> 51 (28%, vs. 31%*) 46 (25%), n=182, (vs. 26%*)	Patients with repeat procedures – no. of pts One repeat procedure – no. of pts Two repeat procedures – no. of pts Three repeat procedures – no. of pts			34 3 0	3 2 0 7
Current type of atrial fibrillation			PVI only – no. of pts	74	8	16	3
Paroxysmal – no. (%) Persistent – no. (%) Long-standing persistent (>1-year duration) – no.	66 (33) 134 (67) 55	72 (37) 125 (63) 56	PVI + additional lesions – no. of pts	77	10	21	2
†Left atrial diameter – mm	Not available	Not available					
†Left ventricular ejection fraction – %	29.0 (25.0-32.0)	30.0 (25.0-32.0)					

#### Survival Free of Death from any cause or admission for worsening heart failure



#### **Conclusions: CASTLE AF**

- CASTLE AF supports and expands upon findings from multiple previous trials
- AF ablation in CHF pts results in more SR, improved EF, functional status and QoL
- Now adds mortality benefit
- Establish a new paradigm for management of CHF pts with HF

# Ablation of AF in HF pts: additional outcomes of the CABANA trial

• Purpose:

To determine the impact of AF ablation compared to drug therapy in pts with HF in the CABANAA trial

• Primary Endpoint:

Composite of death from any cause or heart failure hospitalization

#### **HF** patient characteristics

	Non-HF Subjects N=1422	HF Subjects N=778	P-Value
Age, Median (Q1, Q3)	68 (62,72)	68 (62,73)	0.185
<65 yrs	480/1422 (33.8%)	284/778 (36.5%)	
65 to <75yrs	765/1422 (53.8%)	363/778 (46.7%)	
≥ 75	177/1422 (12.4%)	131/778 (16.8%)	
Sex (Female)	472/1422 (33.2%)	345/778 (44.3%)	<.001
Minority	164/1418 (11.6%)	61/778 (7.8%)	0.006
BMI, Median (Q1, Q3)	30 (26,34)	31 (27,35)	0.002

#### AF type at enrollment



#### **Risk of all cause mortality ITT: impact of HF**



#### **Conclusions: CASTLE AF**

- 60% of HF patients have persistent AF
- Reduction of mortality in HF patients with ablation therapy
- AF recurrences is also reduced in HF patients with ablation therapy

#### FINAL CONCLUSION: AF and HF

#### 6.3.4. Catheter Ablation in HF

	Recommendation for Catheter Ablation in HF									
Re	Referenced studies that support the new recommendation are summarized in Online Data									
	Supplement 7.									
COR	LOE	Recommendation								
llb	B-R	<ol> <li>AF catheter ablation may be reasonable in selected patients with symptomatic AF and HF with reduced left ventricular (LV) ejection fraction (HFrEF) to potentially lower mortality rate and reduce hospitalization for HF (S6.3.4-1, S6.3.4-2).</li> <li>NEW: New evidence, including data on improved mortality rate, has been published for AF catheter ablation compared with medical therapy in patients with HF.</li> </ol>								