

ELECTRA



18^{èmes} journées françaises
pratiques de rythmologie
& de stimulation cardiaque

5-6 DÉCEMBRE 2024

HOTEL VILLA MASSALIA,
MARSEILLE | FRANCE

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Stratégies d'ablation de la FA persistante: Plan-MARSHALL



N Derval
Bordeaux



- Consulting and lecture fees from Biosense Webster
- Study Grant (IIS) from Biosense Webster



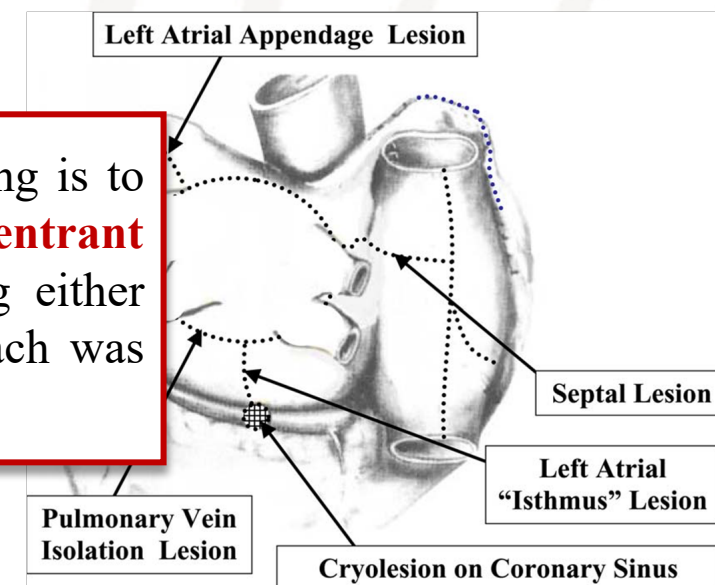
ANATOMICAL APPROACH

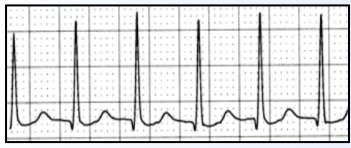
hope that a map-guided surgical procedure could be developed for the treatment of the entire spectrum of these arrhythmias. The electrophysiologic data suggested that atrial fibrillation might have the same relationship to atrial flutter as ventricular fibrillation has to ventricular tachycardia. Some types of atrial fibrillation might be preceded by atrial flutter, during which the entire atrium is driven by a single "macro-reentrant wave" that degenerates into the multiple wavelets characteristic of atrial fibrillation. Much as ventricular tachycardia is often prevented by surgical procedures directed at the bundle branch, atrial tachycardia, we thought it possible that atrial fibrillation might be prevented by surgical procedures directed at atrial flutter. We still are not certain what relationship exists between atrial flutter and atrial fibrillation. However, if this hypothesis proves to be true in the future, intraoperative mapping of atrial flutter might eventually be used

to guide surgical intervention. However, our present studies indicate that once complex atrial fibrillation caused by multiple reentrant flutter waves has developed, the changing patterns of

... perhaps the only way to prevent the atrium from fibrillating is to **interrupt *all* of the potential pathways for atrial macroreentrant circuits** that have been identified by intraoperative mapping either experimentally or clinically, and our eventual surgical approach was based on this principle...

for atrial macroreentrant circuits that have been identified by intraoperative mapping either experimentally or clinically, and our eventual surgical approach was based on this principle. The





SVT

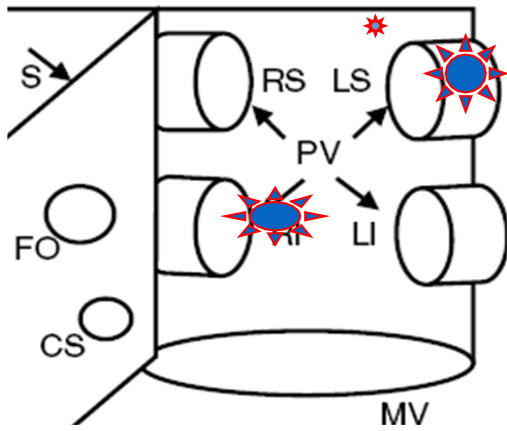




SVT



Parox AF



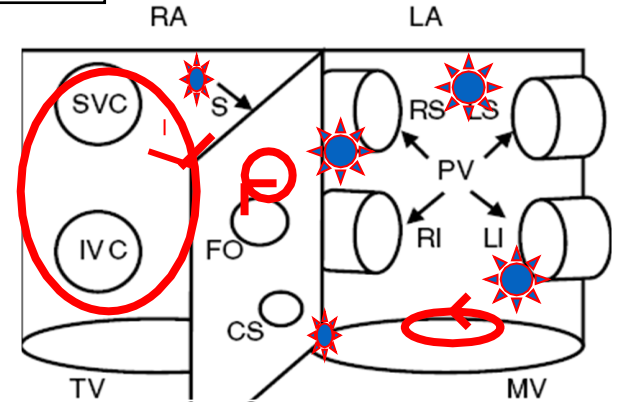
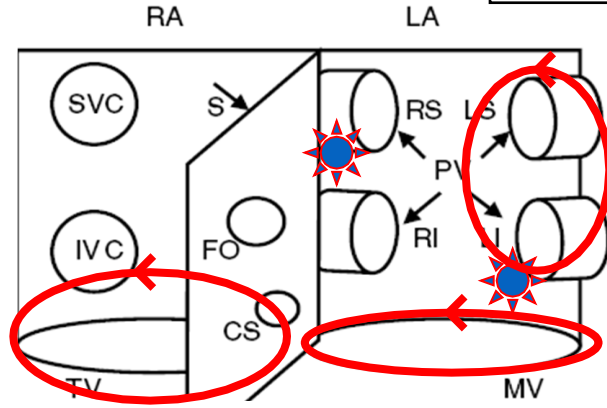
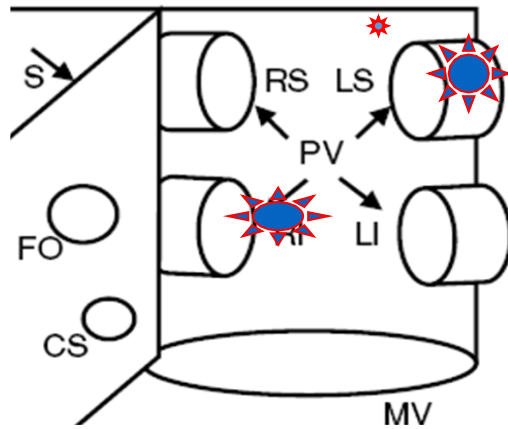
Trigger to "start-up"
Induction phase



SVT

Parox AF

Persistent AF



Trigger to "start-up"
Induction phase



Primer to "warm-up"
Escalation phase



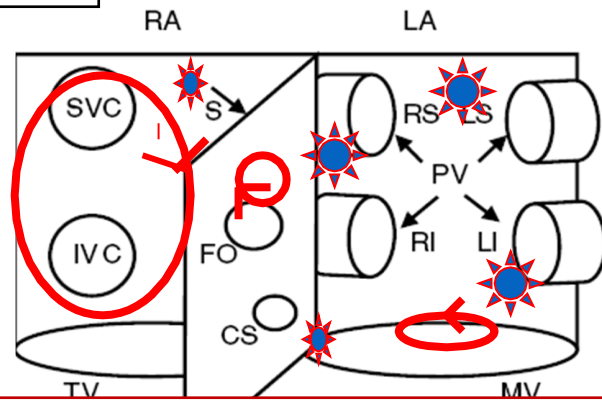
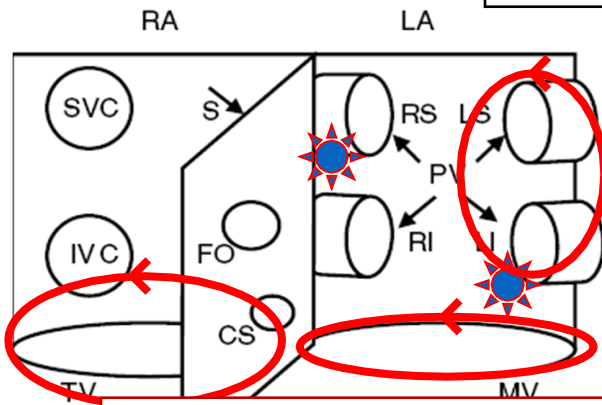
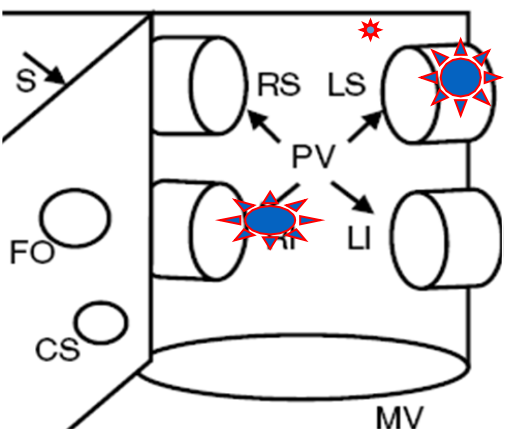
Driver to "keep-up"
Perpetuation phase



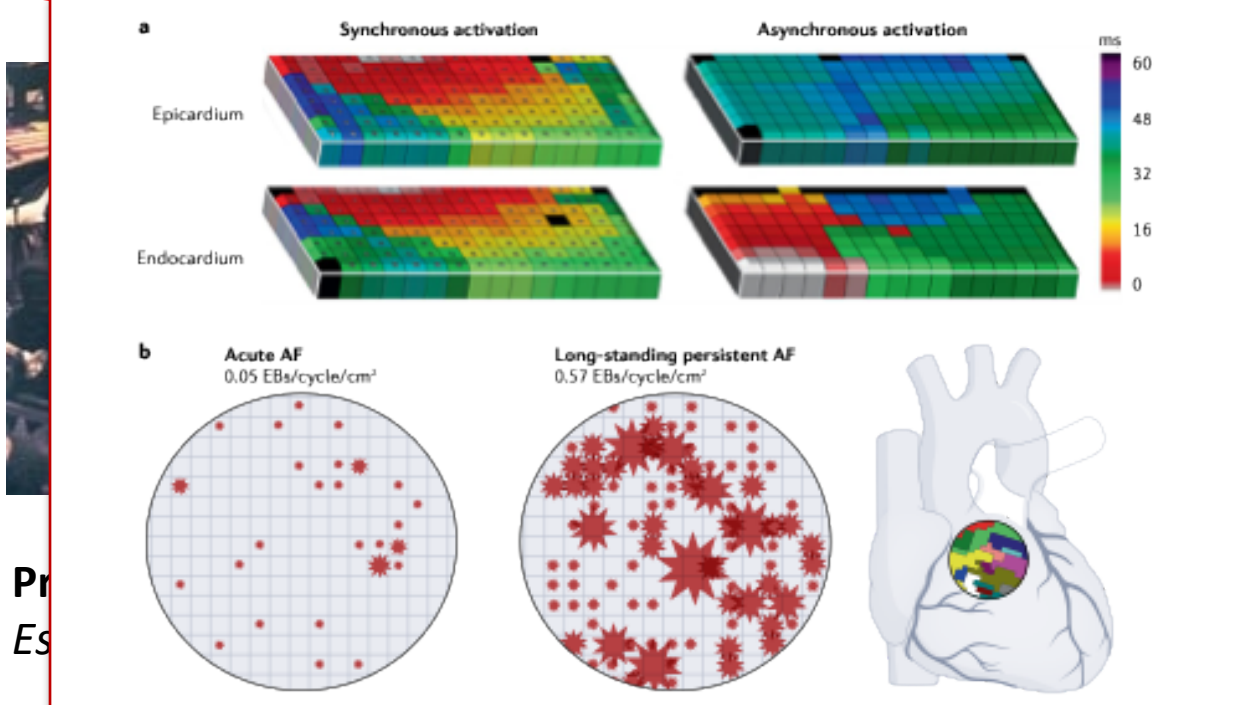
SVT

Parox AF

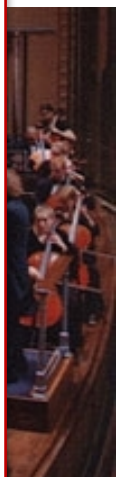
Persistent AF



Trigger to "start-up"
Induction phase



Pr
Es

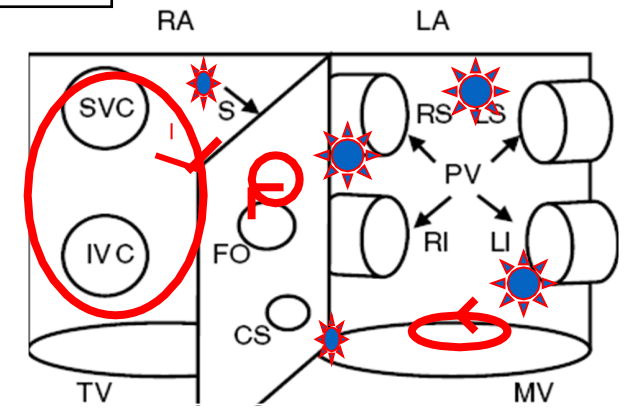
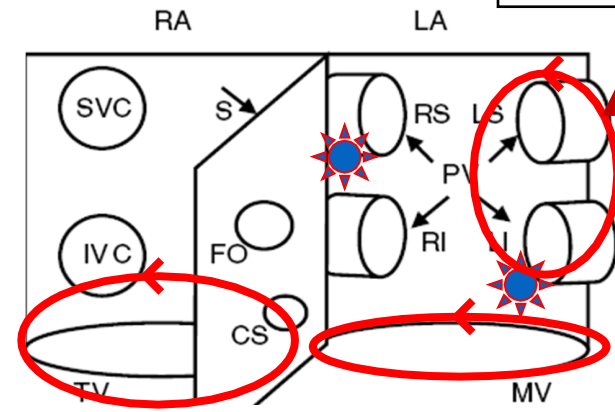
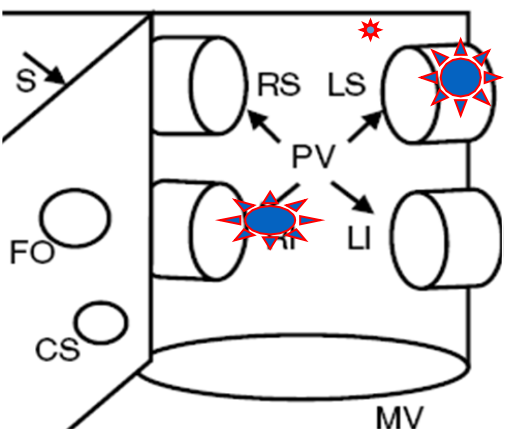




SVT

Parox AF

Persistent AF




Anatomical approach target the main persistent AF "primers"



PLAN-MARSHALL STRATEGY

Journal of Cardiovascular Electrophysiology

ORIGINAL ARTICLE |  Full Access

MARSHALL bundles elimination, Pulmonary veins isolation and Lines completion for ANatomical ablation of persistent atrial fibrillation: MARSHALL-PLAN case series

Thomas Pambrun MD✉, Arnaud Denis MD, Josselin Duchateau MD, Frédéric Sacher MD, PhD, Méleze Hocini MD, Pierre Jais MD, PhD, Michel Haïssaguerre MD, Nicolas Derval MD

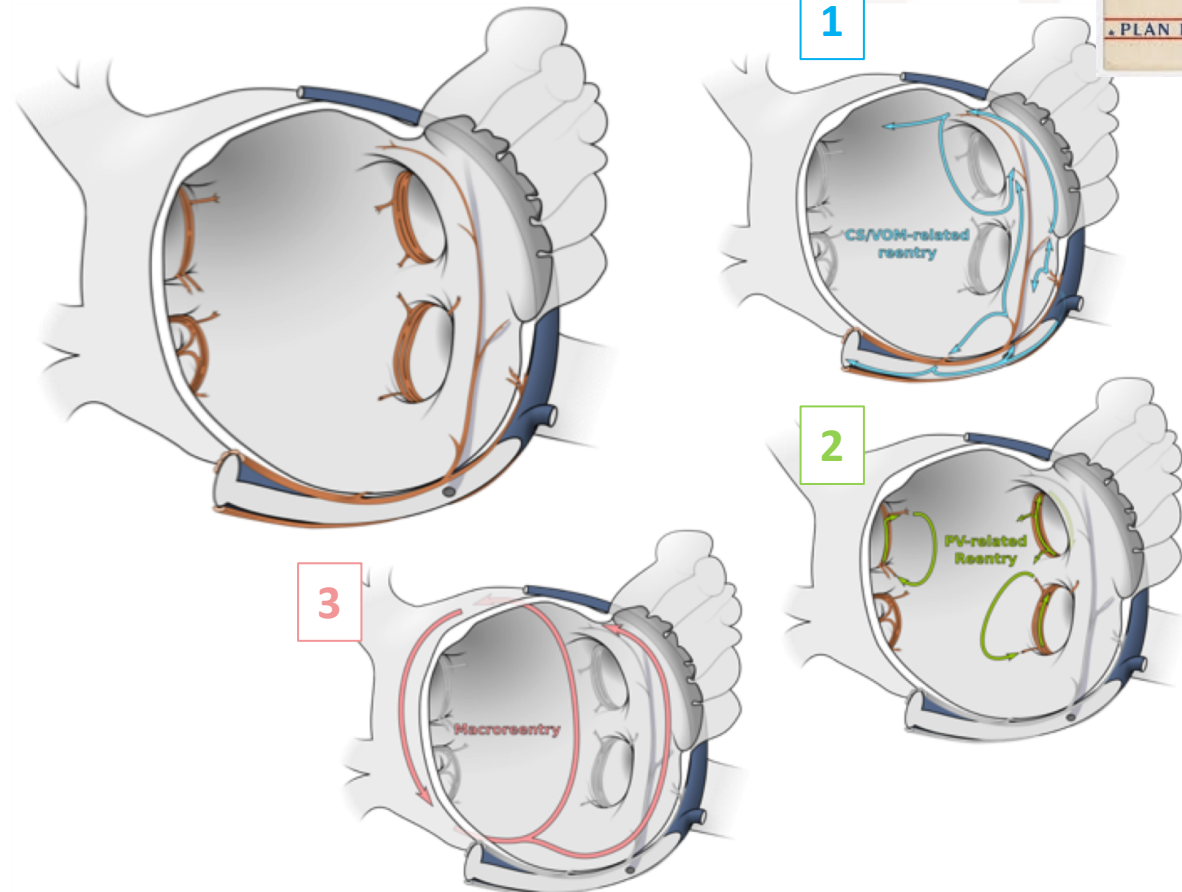
First published: 21 November 2018 | <https://doi.org/10.1111/jce.13797>

Marshall bundle elimination, Pulmonary vein isolation, and Line completion for ANatomical ablation of persistent atrial fibrillation (Marshall-PLAN): Prospective, single-center study

Nicolas Derval, MD,^{*†} Josselin Duchateau, MD, PhD,^{*††} Arnaud Denis, MD,^{*†} F. Daniel Ramirez, MD,^{*†} Saagar Mahida, MD,[‡] Clémentine André, MD,^{*†} Philipp Krisai, MD,^{*†} Yosuke Nakatani, MD,^{*†} Takeshi Kitamura, MD,^{*†} Masateru Takigawa, MD,^{*†} Remi Chauvel, MD,^{*†} Romain Tixier, MD,^{*†} Xavier Pillois, PhD,^{*††} Frédéric Sacher, MD, PhD,^{*††} Méleze Hocini, MD,^{*††} Michel Haïssaguerre, MD,^{*††} Pierre Jais, MD, PhD,^{*††} Thomas Pambrun, MD^{*†}


- **Target:** Muscular network highly propitious for reentry
- **Endpoint:** complete lesion set

ATRIAL BODY MYOCARDIUM + ADJACENT VENOUS MUSCULATURE



PLAN-MARSHALL STRATEGY

Journal of Cardiovascular Electrophysiology

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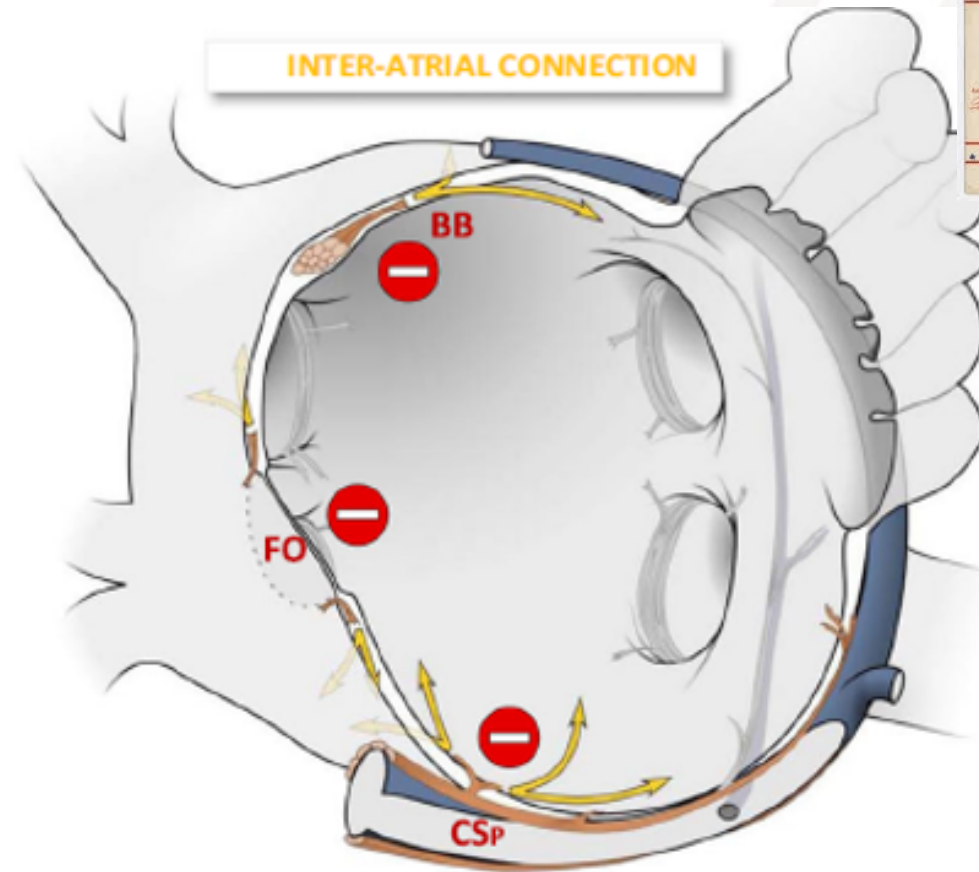
Thomas Pambrun MD✉, Arnaud Denis MD, Josselin Duchateau MD, Frédéric Sacher MD, PhD, Méleze Hocini MD, Pierre Jais MD, PhD, Michel Haïssaguerre MD, Nicolas Derval MD

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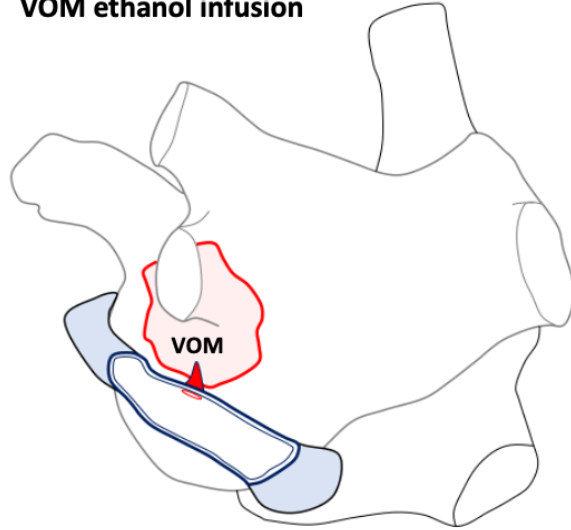


SPARING Inter-Atrial Connections

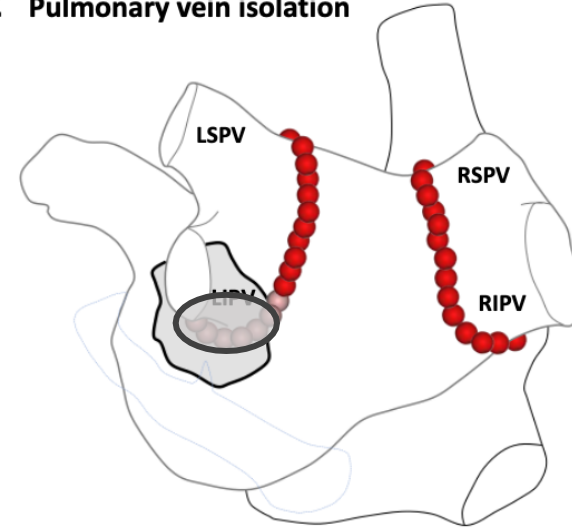


PLAN-MARSHALL STRATEGY

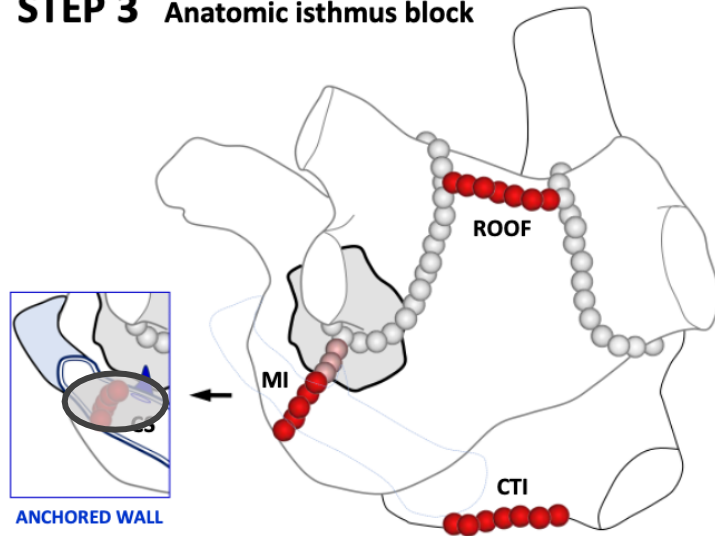
STEP 1 VOM ethanol infusion



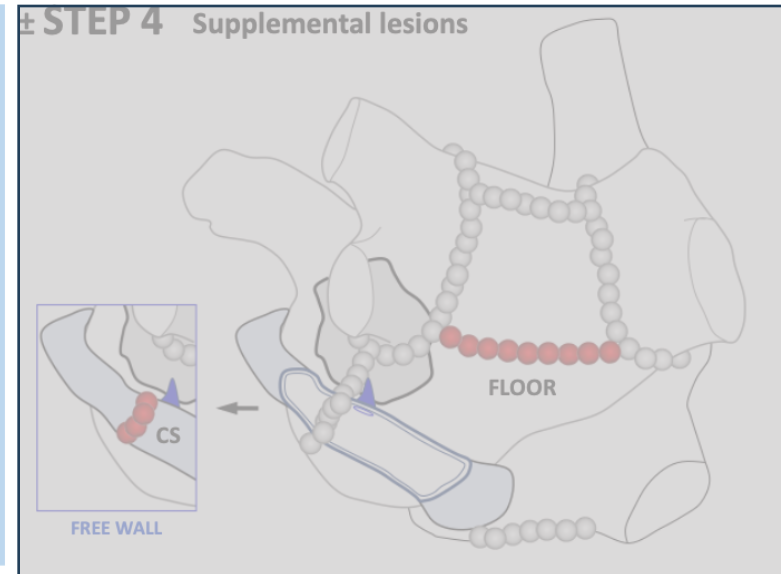
STEP 2 Pulmonary vein isolation



STEP 3 Anatomic isthmus block



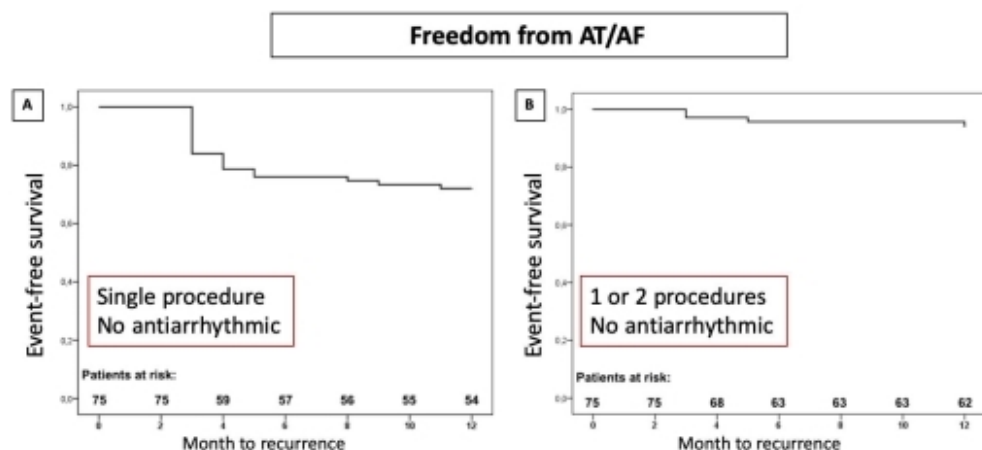
± STEP 4 Supplemental lesions



RESULTS OF "PLAN-MARSHALL" ABLATION

Prospective monocentric
75 patients with pers AF
VOM OH success rate: 92%

2020



Single procedure, off AAD, 12 m: **72%**

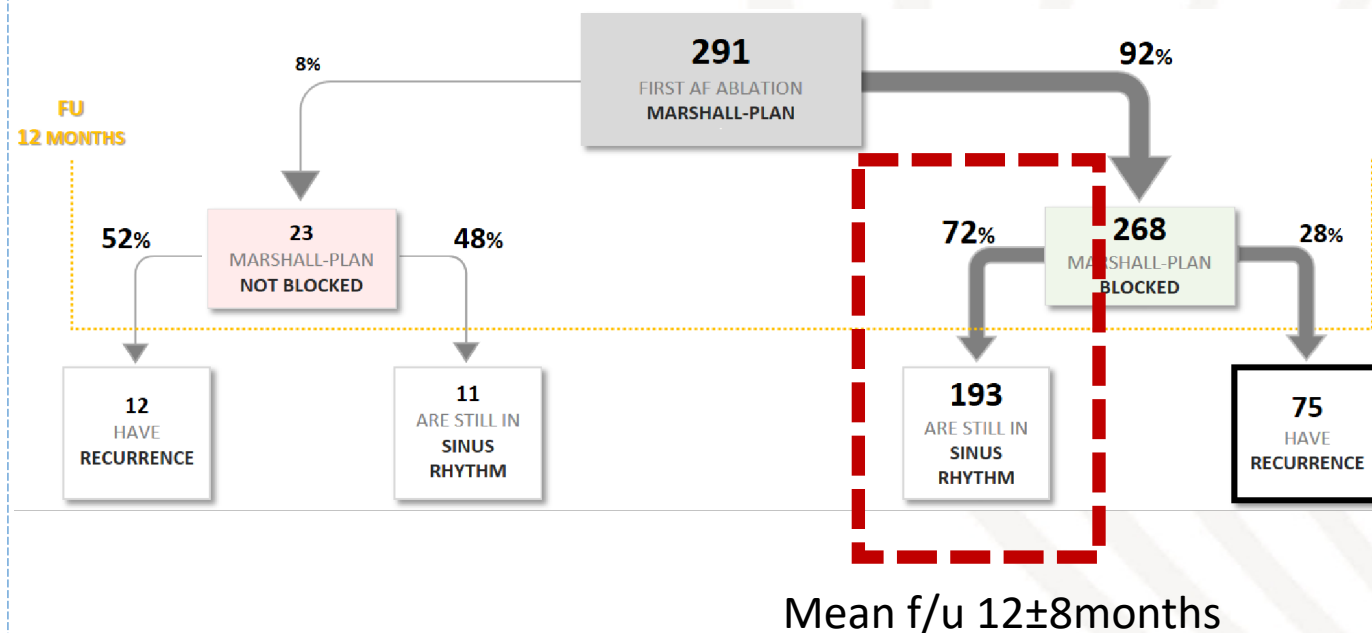
Single procedure, off AAD, 12m, VOM+: **79%**

Multiple procedure (1,28/pt), off AAD: **89%**

Derval et al. Heart Rhythm 2021

Retrospective monocentric
291 patients with pers AF
VOM success rate 97%

2022

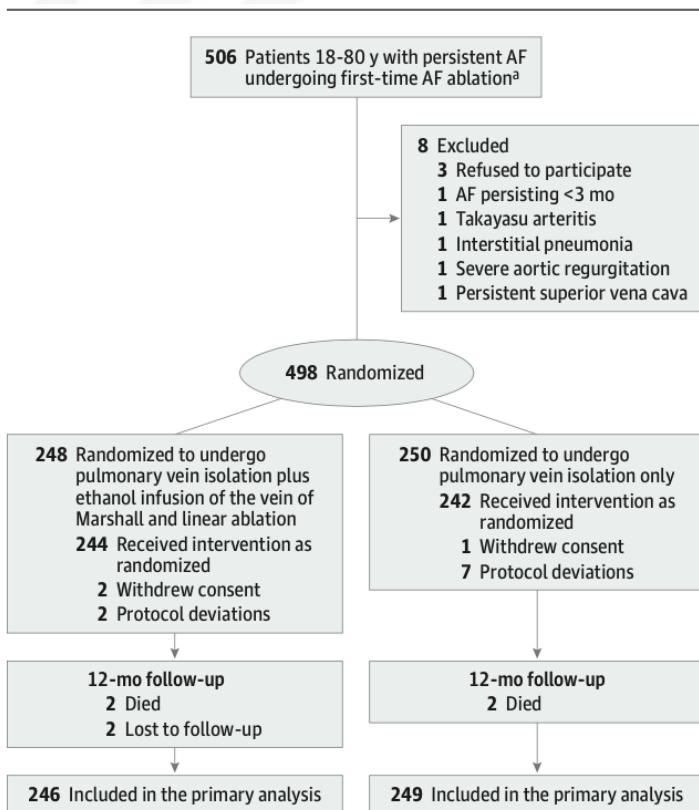


Takagi et al. Heart Rhythm 2022

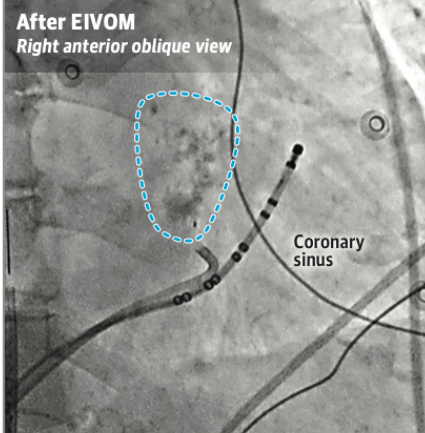
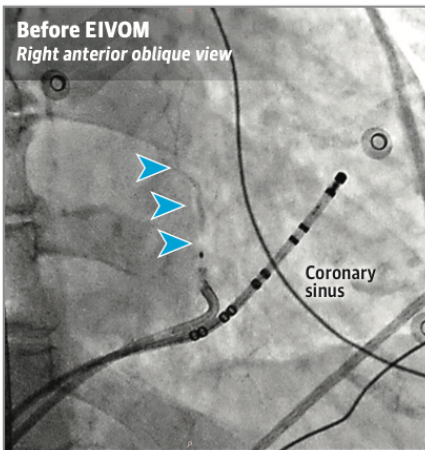
Pulmonary Vein Isolation With Optimized Linear Ablation vs Pulmonary Vein Isolation Alone for Persistent AF

The PROMPT-AF Randomized Clinical Trial

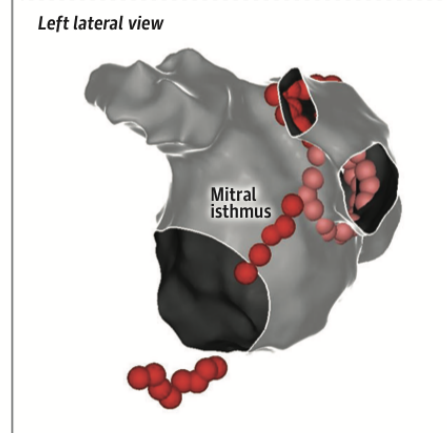
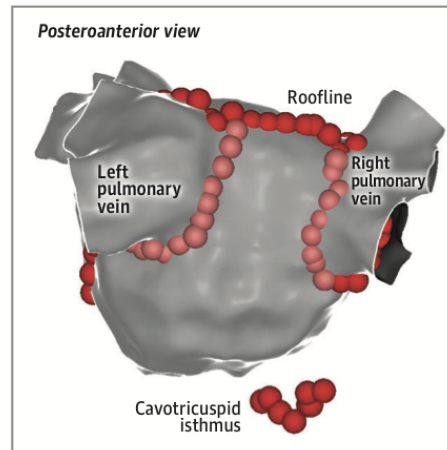
Caihua Sang, MD; Qiang Liu, MD; Yiwei Lai, MD; Shijun Xia, MD; Ruhong Jiang, MD; Songnan Li, MD; Qi Guo, MD; Qifan Li, MD; Mingyang Gao, MD; Xueyuan Guo, MD; Lihong Huang, MD; Nian Liu, MD; Chenxi Jiang, MD; Song Zuo, MD; Xiaoxia Liu, MD; Mengmeng Li, MD; Weili Ge, MD; Shangming Song, MD; Lianghua Chen, MD; Shuanglun Xie, MD; Jiangang Zou, MD; Ke Chen, MD; Xiangfei Liu, MD; Hesheng Hu, MD; Xinhua Wang, MD; Jinlin Zhang, MD; Zhaojun Wang, MD; Chi Wang, MPH; Liu He, PhD; Chao Jiang, MD; Ribo Tang, MD; Ning Zhou, MD; Yunlong Wang, MD; Deyong Long, MD; Xin Du, MD; Chenyang Jiang, MD; Laurent Macle, MD; Jianzeng Dong, MD; Changsheng Ma, MD; for the PROMPT-AF investigators



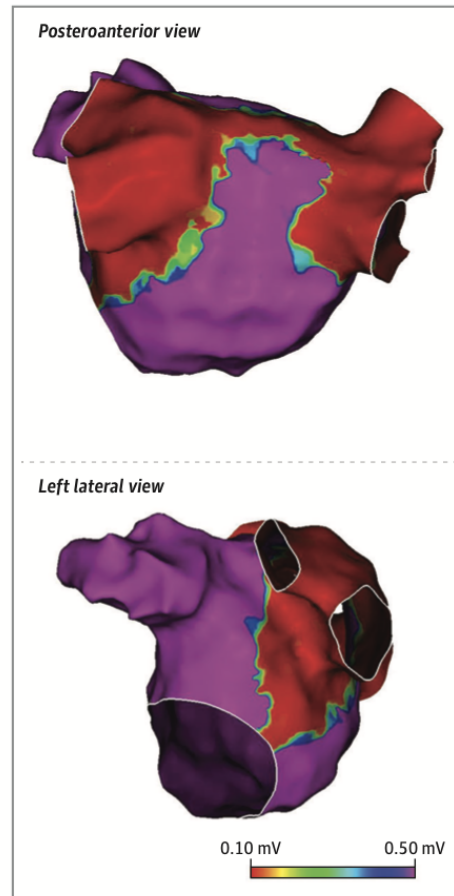
A Ethanol infusion via vein of Marshall (EIVOM)



B Bilateral pulmonary vein isolation and linear ablation



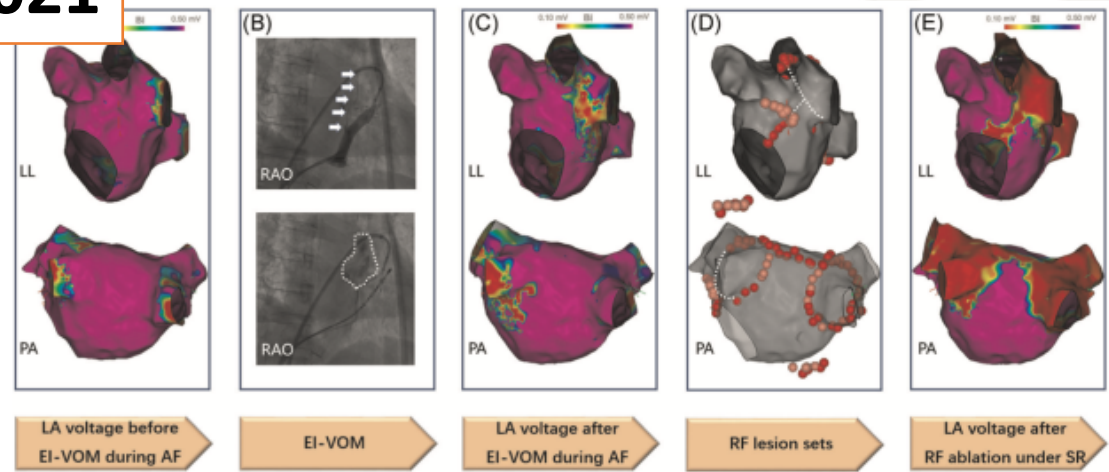
C Voltage map at end of procedure



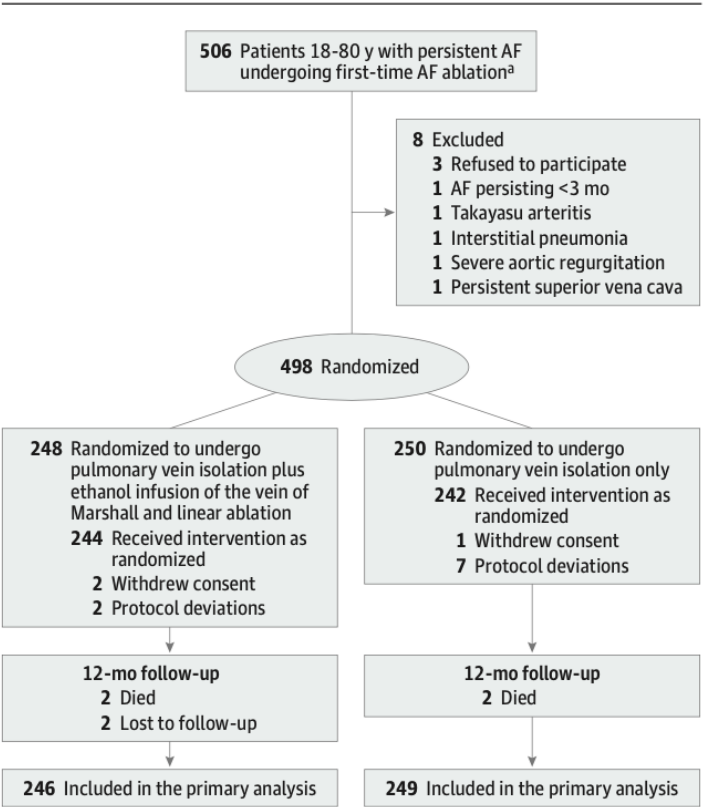
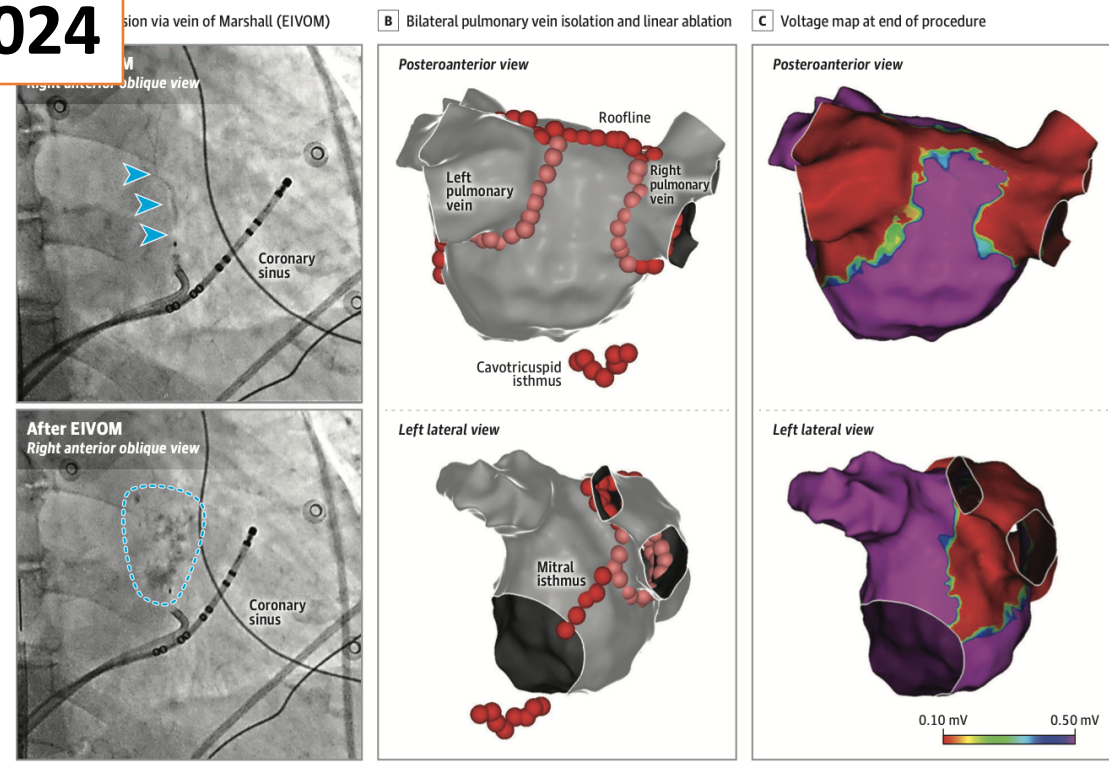
Pulmonary Vein Isolation With Optimized Linear Ablation vs Pulmonary Vein Isolation Alone for Persistent AF The PROMPT-AF Randomized Clinical Trial

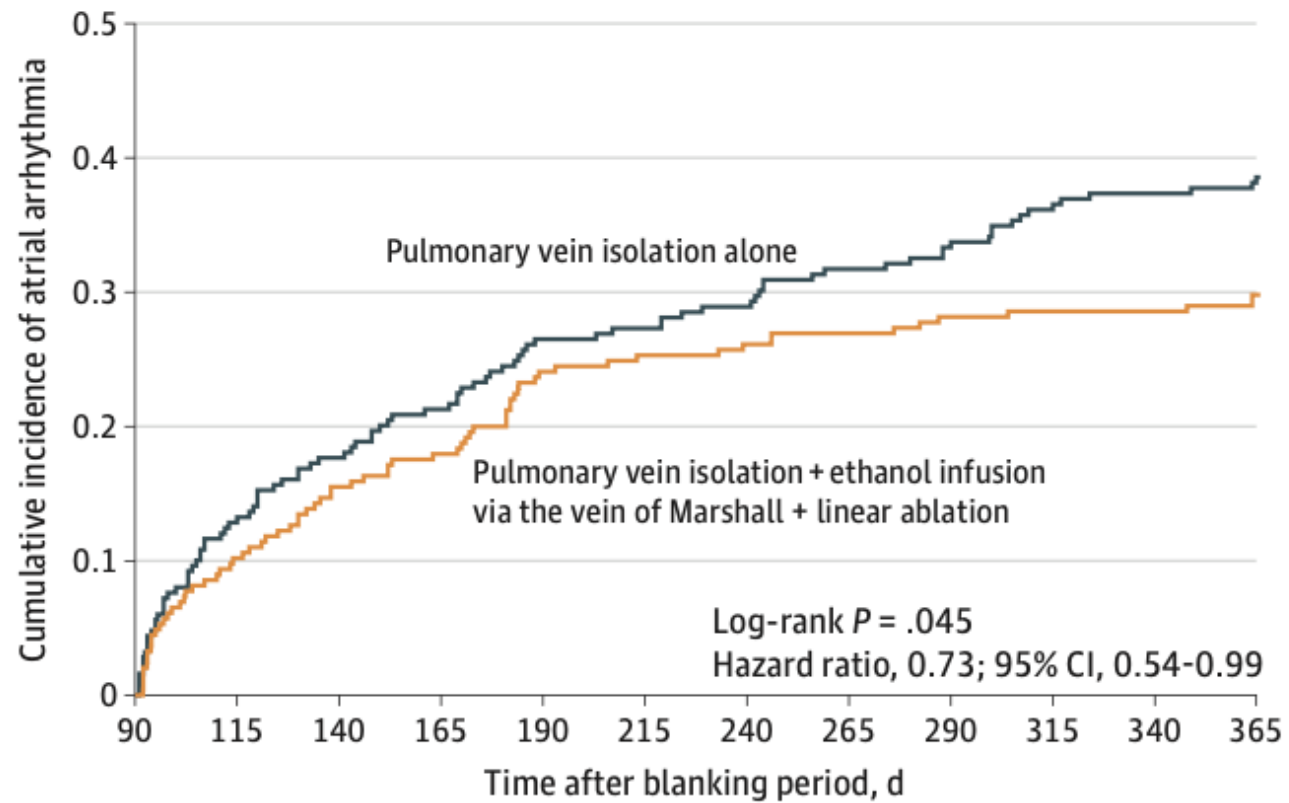
Caihua Sang, MD; Qiang Liu, MD; Yiwei Lai, MD; Shijun Xia, MD; Ruhong Jiang, MD; Songnan Li, MD; Qi Guo, MD; Qifan Li, MD; Mingyang Gao, MD; Xueyuan Guo, MD; Lihong Huang, MD; Nian Liu, MD; Chenxi Jiang, MD; Song Zuo, MD; Xiaoxia Liu, MD; Mengmeng Li, MD; Weili Ge, MD; Shangming Song, MD; Lianghua Chen, MD; Shuanglun Xie, MD; Jiangang Zou, MD; Ke Chen, MD; Xiangfei Liu, MD; Hesheng Hu, MD; Xinhua Wang, MD; Jinlin Zhang, MD; Zhaojun Wang, MD; Chi Wang, MPH; Liu He, PhD; Chao Jiang, MD; Ribo Tang, MD; Ning Zhou, MD; Yunlong Wang, MD; Deyong Long, MD; Xin Du, MD; Chenyang Jiang, MD; Laurent Macle, MD; Jianzeng Dong, MD; Changsheng Ma, MD; for the PROMPT-AF investigators

2021



2024





Cumulative No.

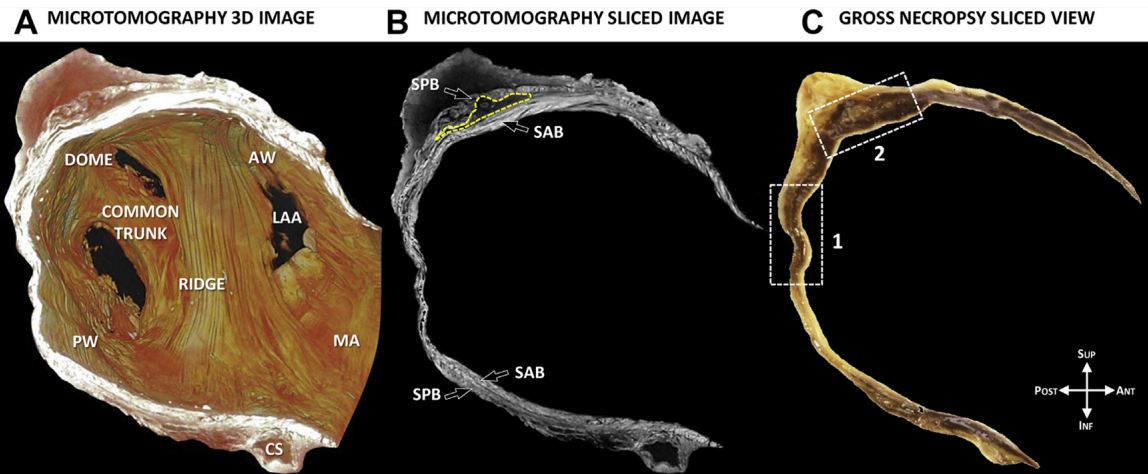
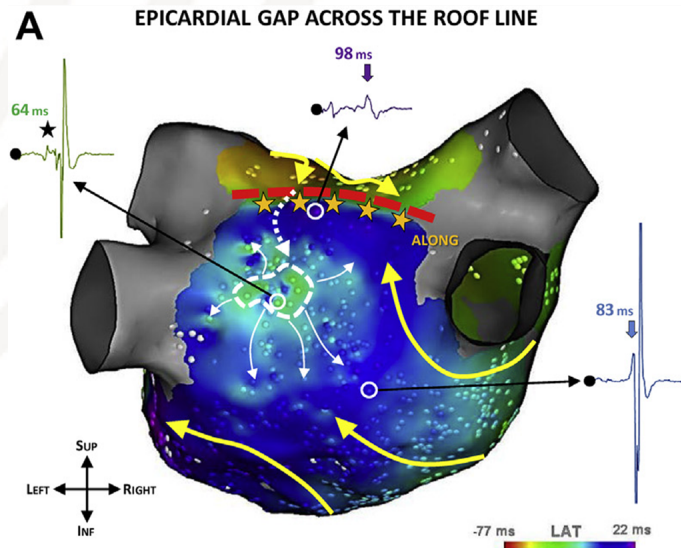
Pulmonary vein isolation + ethanol infusion via the vein of Marshall + linear ablation

Event	0	25	38	44	59	62	64	66	69	70	70	72
Death	0	0	0	0	0	0	0	0	0	0	0	2
Atrial fibrillation	0	23	36	41	47	50	51	53	56	56	57	58
Redo procedure	0	1	5	6	7	8	9	10	10	10	11	11

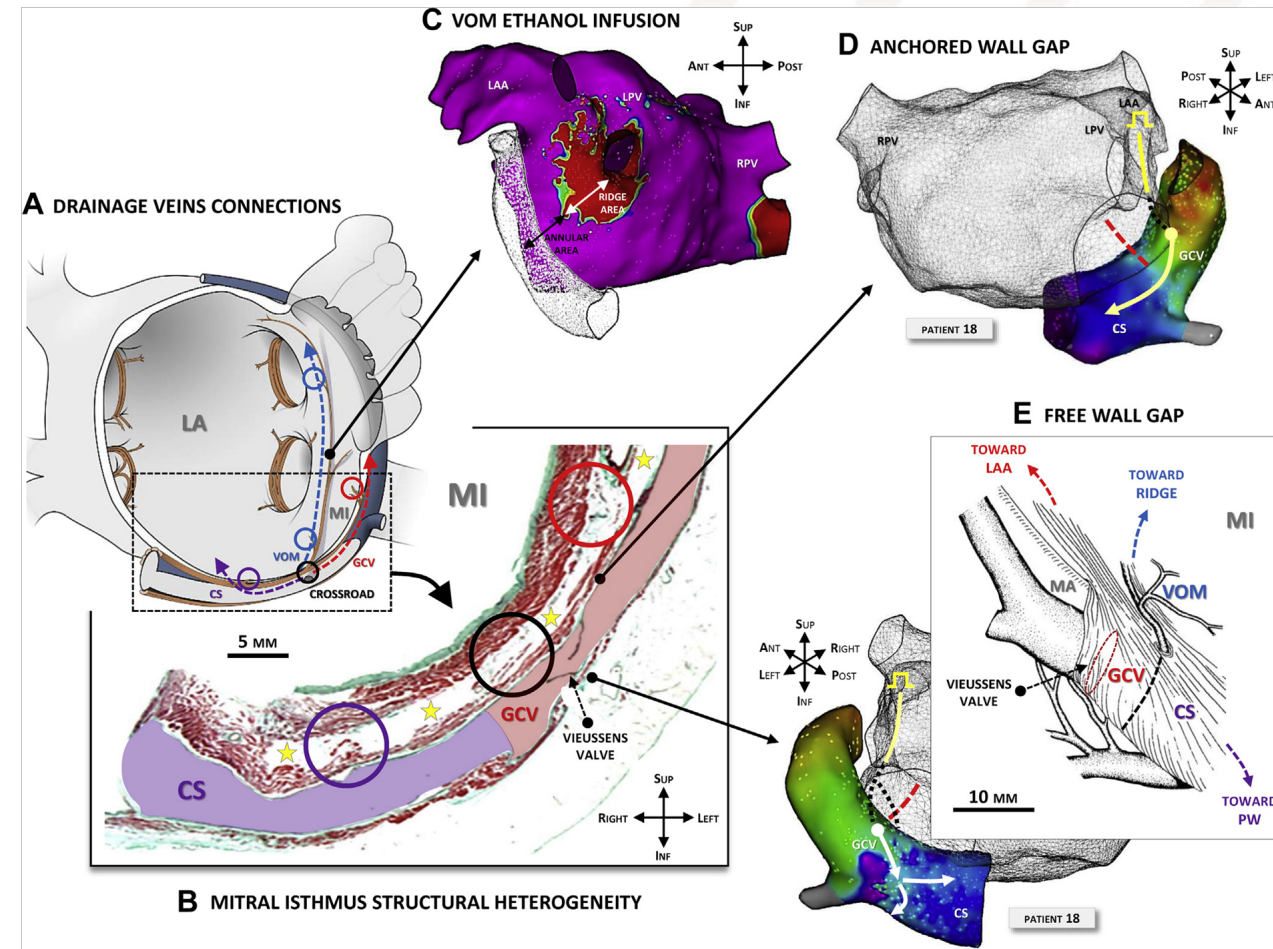
Pulmonary vein isolation alone

Event	0	33	44	53	66	68	72	79	84	91	93	96
Death	0	0	0	0	0	1	2	2	2	2	2	2
Atrial fibrillation	0	27	34	42	48	49	54	62	64	70	74	75
Redo procedure	0	7	7	9	9	9	10	16	17	17	18	20

Epicardial course of the septopulmonary bundle: Anatomical considerations and clinical implications for roof line completion



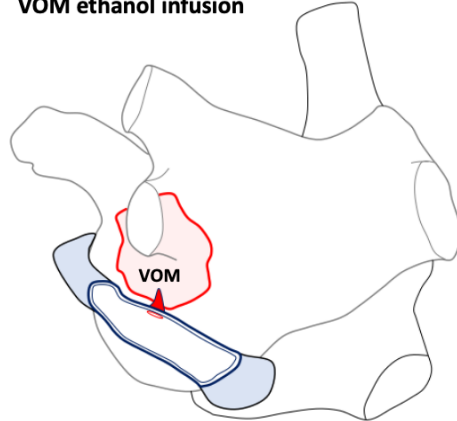
Epicardial course of the musculature related to the great cardiac vein: Anatomical considerations and clinical implications for mitral isthmus block after vein of Marshall ethanol infusion



PLAN-MARSHALL STRATEGY

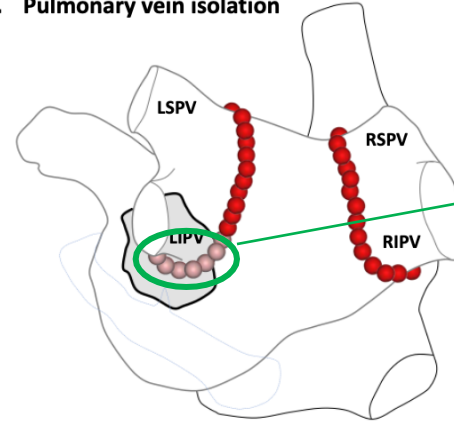


STEP 1 VOM ethanol infusion



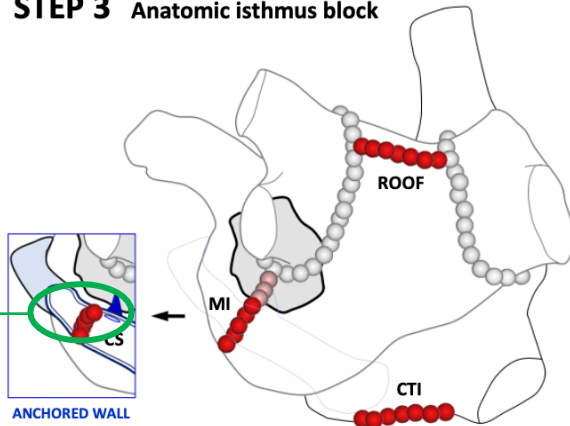
↗ success rate
↘ dissection

STEP 2 Pulmonary vein isolation



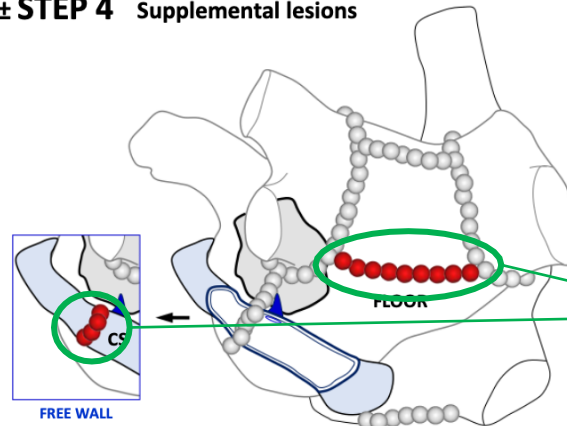
Systematic ablation

STEP 3 Anatomic isthmus block

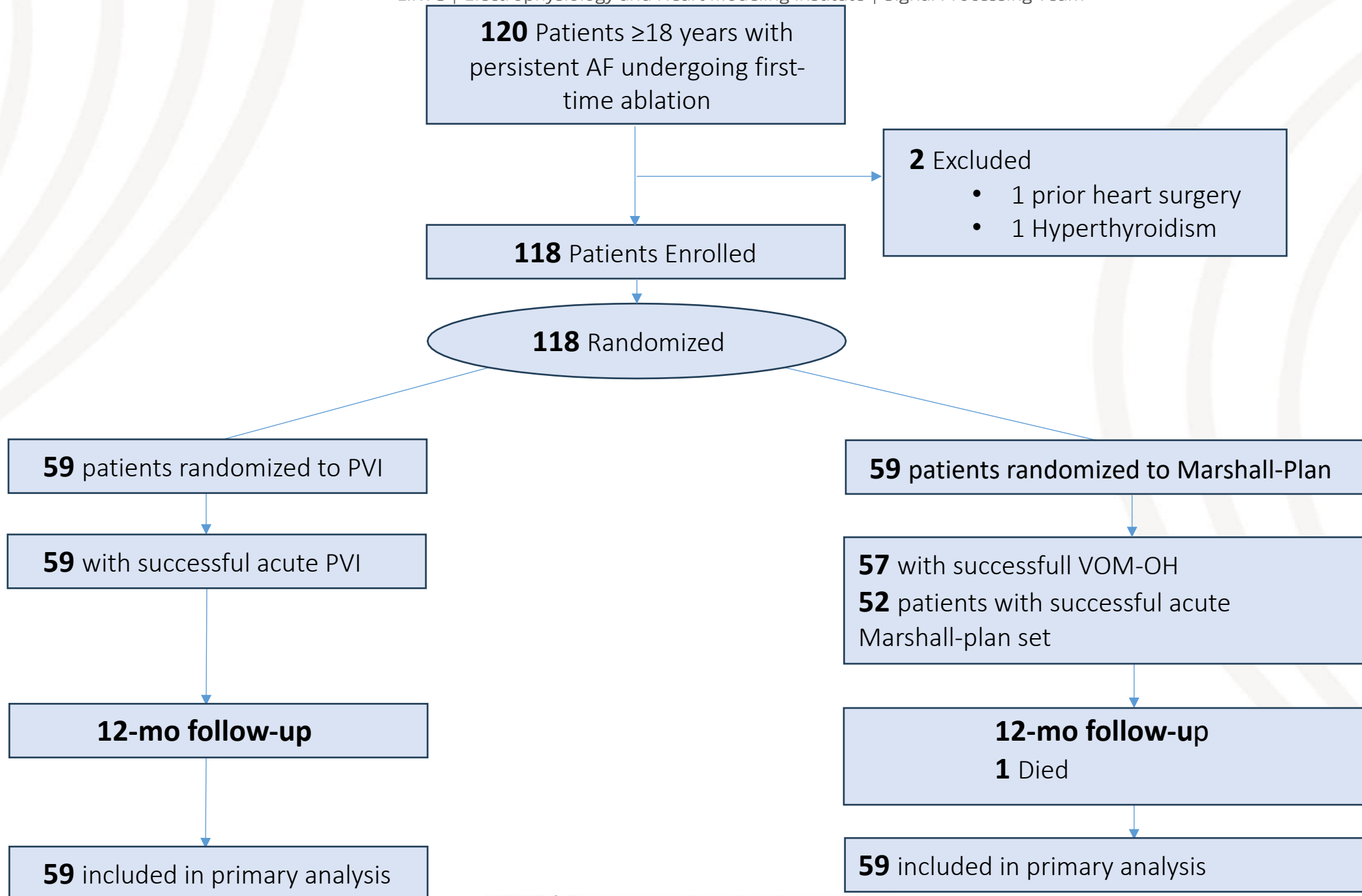


Systematic ablation

± STEP 4 Supplemental lesions



Better understanding





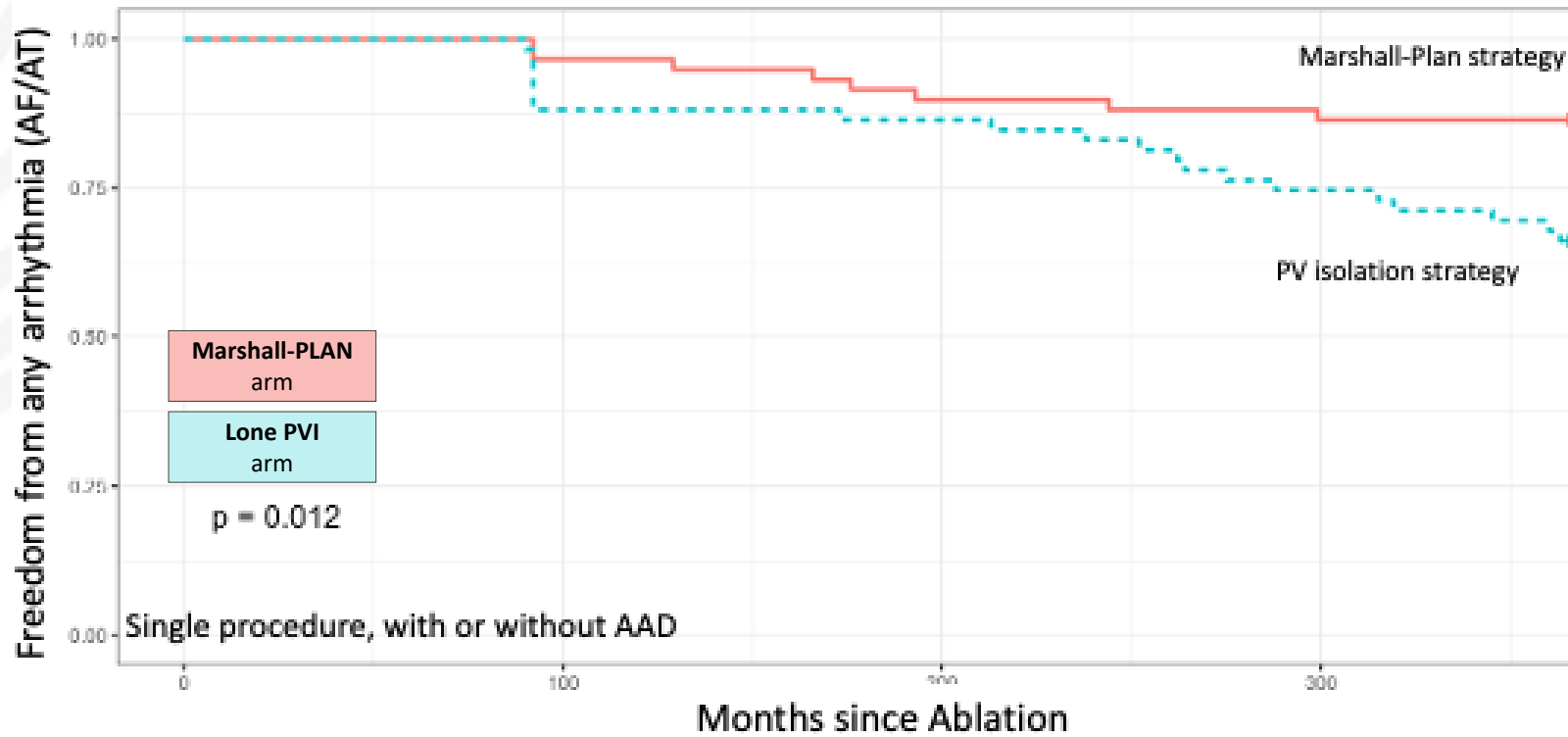
Characteristics	Marshall-Plan N=60	PV Isolation N=60	p-value
Demographics			
Age (y)	66±8	65±8	0.21
Gender			0.47
F	12 (20%)	9 (15%)	
M	48 (80%)	51 (85%)	
CHA ₂ DS ₂ -Vasc	2±1	2±1	0.04
Hypertension	36 (60%)	25 (42%)	0.04
Diabetes	9 (15%)	3 (5%)	0.07
Previous Stroke	5 (8.3%)	2 (3.3%)	0.44
History of Amiodarone	54 (90%)	49 (82%)	0.19
LVEF (%)	51±12	56±10	0.12
SHD	6 (10%)	6 (10%)	0.99
AF Characteristics			
Maximum AF length (m)	10±18	7±6	0.86
Current AF length (m)	9±19	6±7	0.77
Long-standing AF>1y	11 (18%)	11 (18%)	1.00
History of DCC	1±1	1±1	0.32
Rhythm at inclusion			
AF	32 (53%)	39 (65%)	0.19
SR	28 (47%)	21 (35%)	
Left atrial volume (ml)	187±53	192±53	0.31

	Marshall-Plan N=59	PV isolation N=59	p-value
Rhythm in EP lab			0.19
AF	32 (54%)	39 (66%)	
SR	27 (46%)	20 (34%)	
LAA cycle length (if AF) (ms)	206±138	176±26	0.88
LA volume (ml)	182±52	192±53	0.22
PVI	59	59	1.00
Total RF LPV (min)	9.5±4.3	13.0±3.7	0.001
Total RF RPV (min)	13.4±5.1	15.8±5.5	0.012
Total RF PVs (min)	23±8	29±8	0.001
VOM-OH	57 (97%)	-	-
OH volume (ml)	10±2	-	-
Time for OH (min)	22.5±15.4	-	-
Xray time for OH (min)	6.3±6.8	-	-
Mitral isthmus line attempt	56 (95%)	-	-
Mitral isthmus block (% per attempt; %total)	55 (98%; 93%)	-	-
RF mitral line (min)	7.2±6.2	-	-
Posterior wall line attempt	56 (95%)	-	-
Posterior wall block (% per attempt; %total)	54 (96%; 92%)	-	-
Roof line block (% per attempt; %total)	30 (54%; 51%)	-	-
RF roof line (min)	6.3±4.4	-	-
Floor line attempt	26 (46%)	-	-
Floor line block (% per attempt; %total)	24 (92%; 43%)	-	-
RF floor line (min)	4.8±2.6	-	-
CTI line attempt	56 (95%)	-	-
CTI line block(% per attempt; %total)	55 (93%; 98%)	-	-
RF CTI line (min)	5.9±4.5	-	-
AF termination	1 (2%)	3 (5%)	0.43
Ablation set complete	52 (88%)	59 (100%)	0.058
Total Procedure time (min)	157±53	125±31	0.001
Total RF time (min)	36.8±16.0	29.6±8.1	0.001
Total X-Ray time (min)	21±16	11±6	0.001

Intention-to-treat analysis

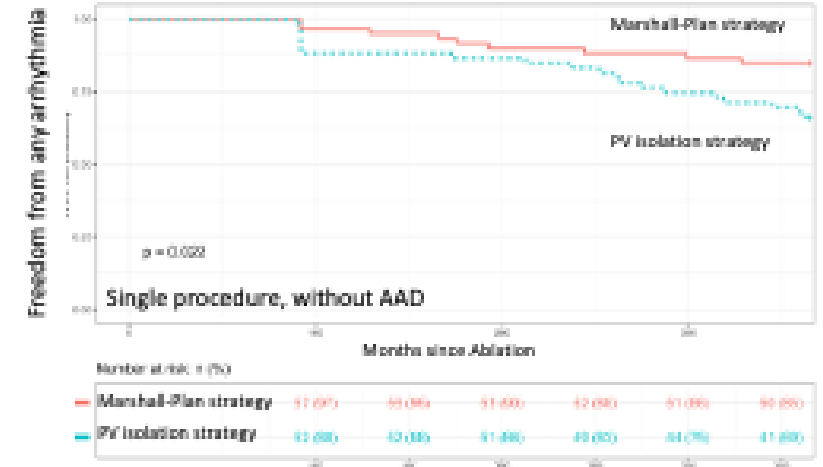
Primary Outcome

Secondary Outcome



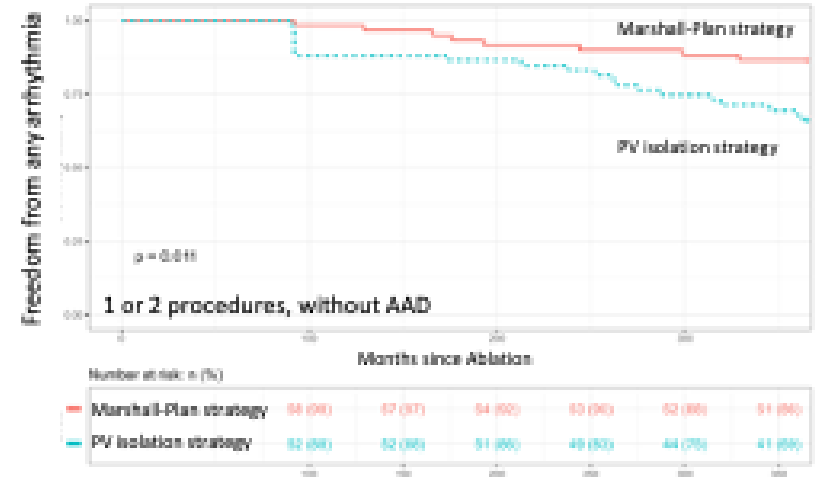
Number at risk: n (%)

Marshall-Plan strategy	57 (97)	56 (95)	53 (90)	52 (88)	51 (86)	51 (86)
PV isolation strategy	52 (88)	52 (88)	51 (86)	49 (83)	44 (75)	41 (69)



Number at risk: n (%)

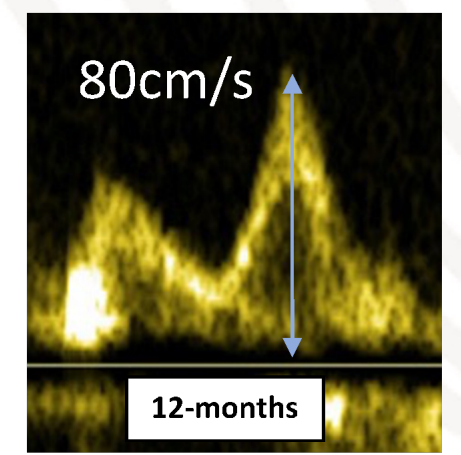
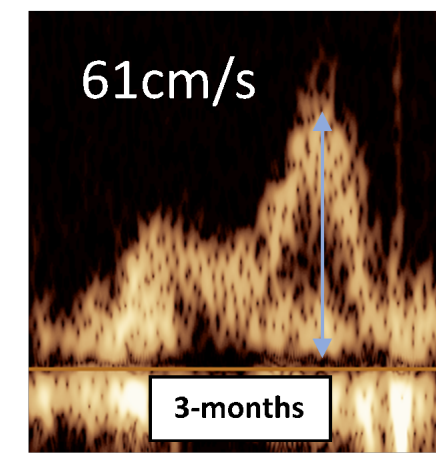
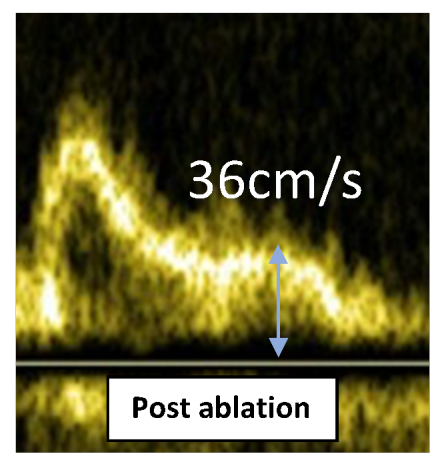
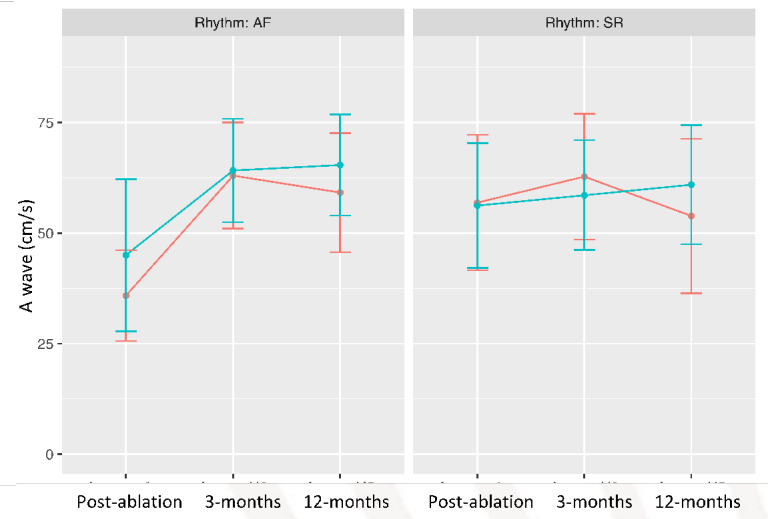
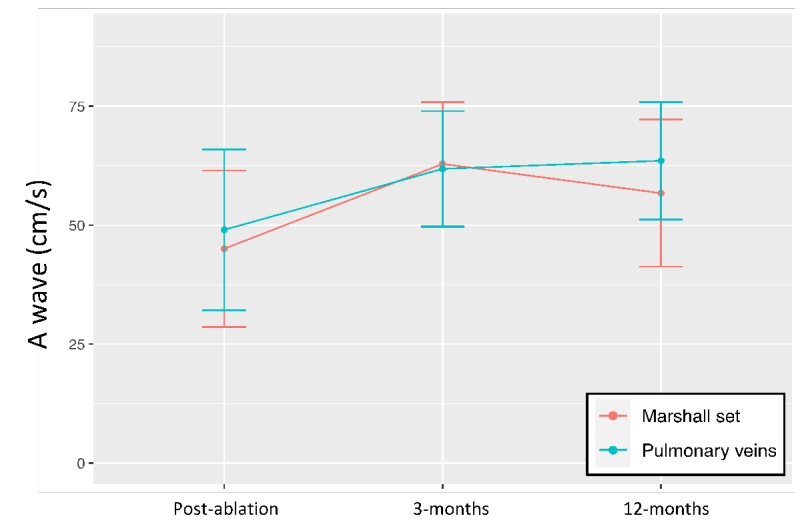
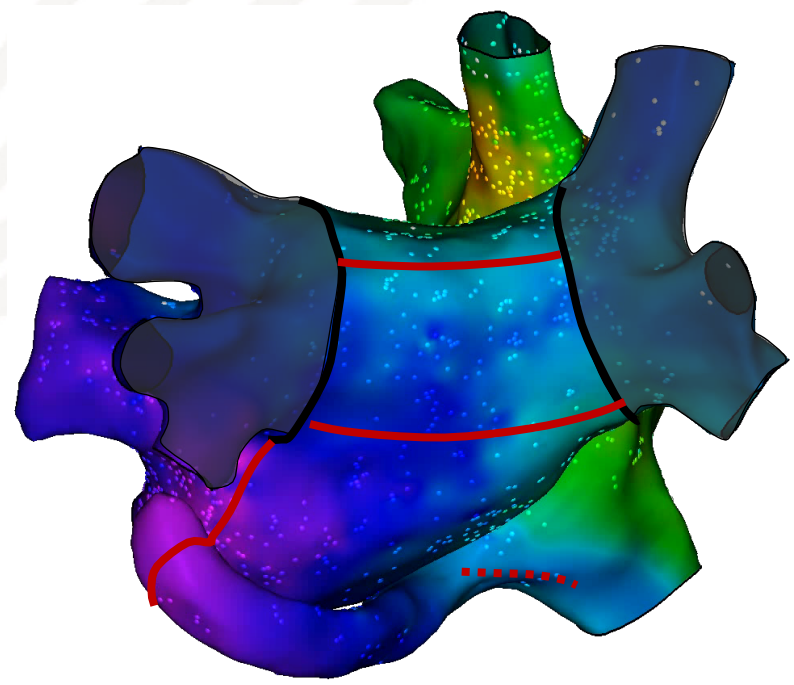
Marshall-Plan strategy	57 (97)	56 (95)	53 (90)	52 (88)	51 (86)	50 (86)
PV isolation strategy	52 (88)	52 (88)	51 (86)	49 (83)	44 (75)	41 (69)



Number at risk: n (%)

Marshall-Plan strategy	54 (96)	57 (97)	54 (92)	53 (90)	52 (88)	51 (86)
PV isolation strategy	52 (88)	52 (88)	51 (86)	49 (83)	44 (75)	41 (69)

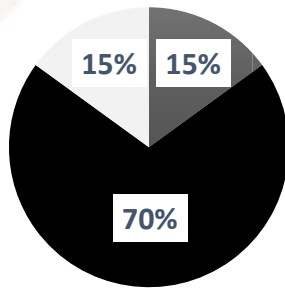
Preservation of LA function



RESULTS: FOLLOW-UP



PV Isolation 20 patients with clinical recurrence



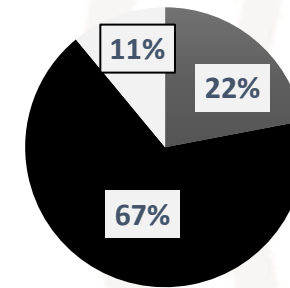
■ parox AF ■ pers AF ■ AT

7 patients with redo (during f/u period)

- All 4 PV isolated in 100% (cross-over to Marshall lesion set)

1 oeso-pericardial fistula

"Marshall Plan" 9 patients with clinical recurrence



■ parox AF ■ pers AF ■ AT

4 patients with redo (during f/u period)

- No patients with complete set
 - Mitral line: 4 (epi, CS)
 - Roof: 2
 - CTI: 3

1 severe Vascular complication

Recurrence type

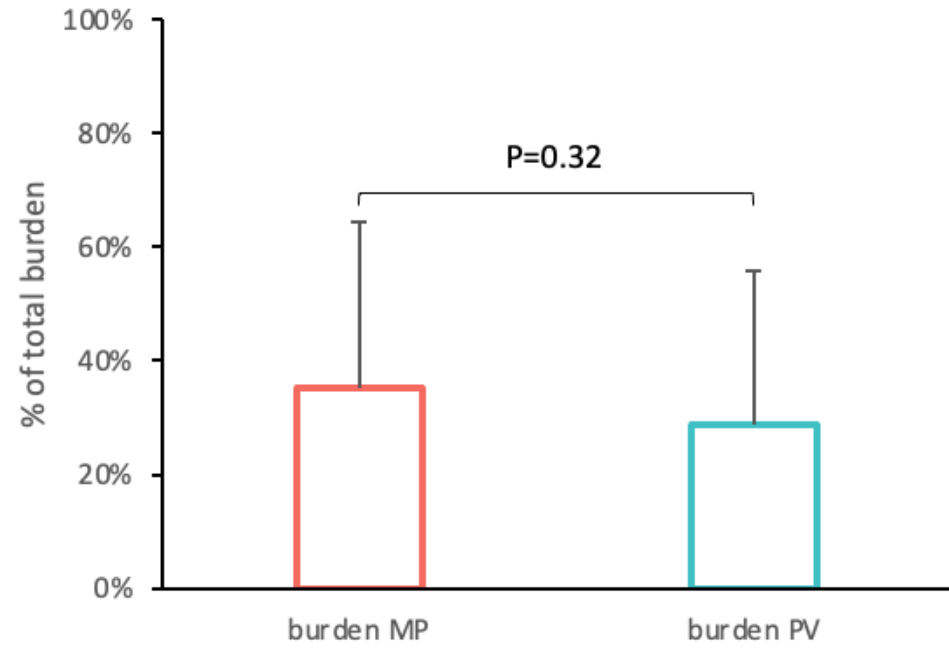
Redo during F/U

complications

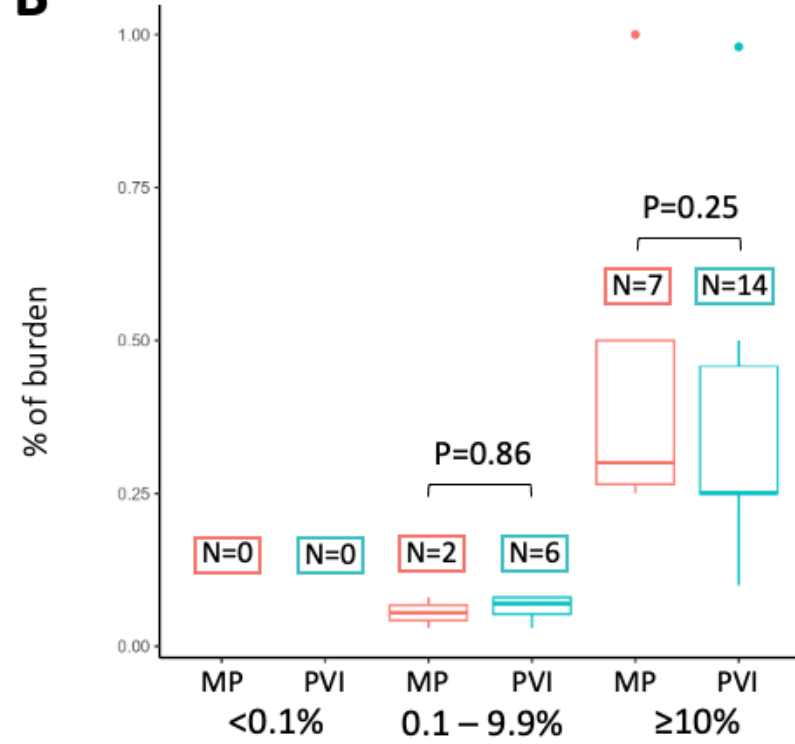


AF burden in patients with clinical recurrence

A



B



7 p

1 oeso-pericardial fistula

complications

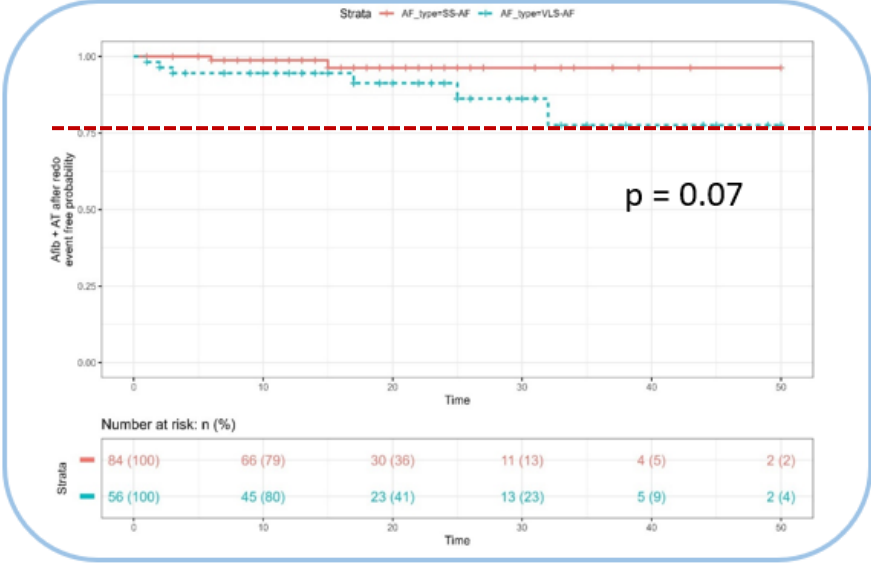
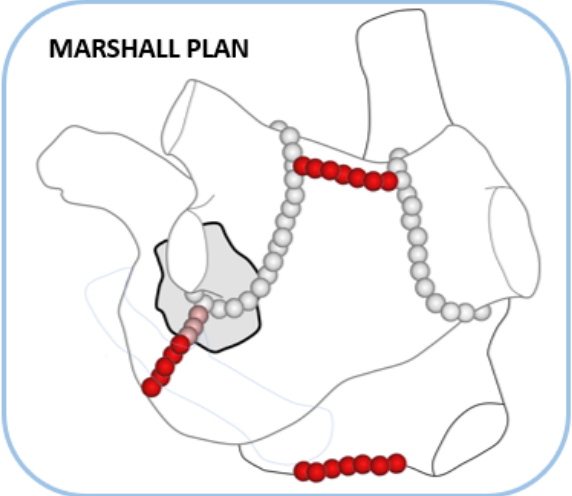
1 Vascular complication

VERY-LONG STANDING PERS AF



**>2years
continuous AF**

160 patients
Very long standing AF
and short-standing AF
Mean follow-up 18.6
+/- 11.52 months



76%

NUMBER NEEDED TO TREAT (NNT)

Blood pressure medicines for 5 years to prevent Death, Heart Attacks and Strokes:

- 125 pts (prevent death)
- 67 pts (prevent strokes)

Warfarin for AF to prevent Stroke

- 25 pts (prevent stroke)

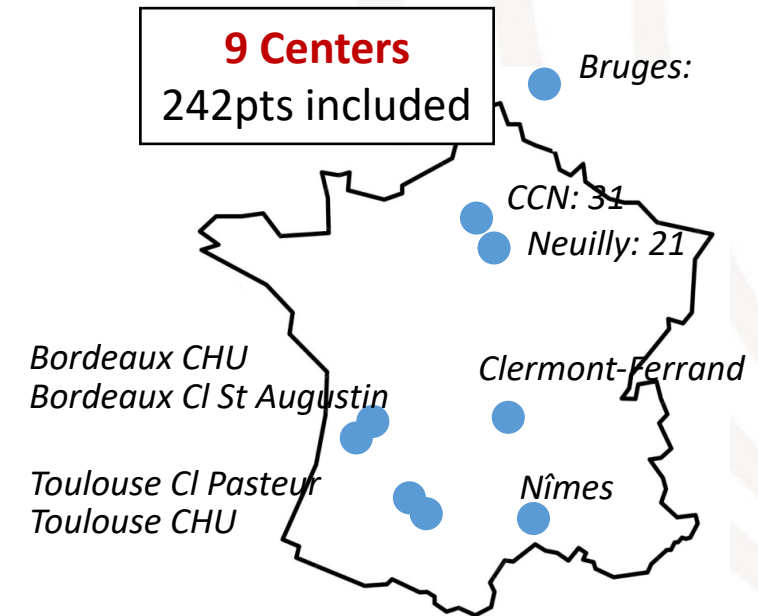
RE-LY: 167pts (prevent stroke)

- PROMPT-AF: 10,9 pts
- Marshall-Plan: 4,9 pts



- The Marshall-plan is a **comprehensive** ablation strategy for persistent AF based on **anatomical** principles.
- **Aim** is a **complete** lesion set performed **systematically** in all persistent AF patients.
- Accumulating evidence supporting superiority against PV isolation:
 - Marshall-plan mono
 - PROMPT-AF
- Marshall-plan strategy respect LA function
- This clear and predefined strategy should:
 - Help to obtained reproducible results
 - Stimulate industry to facilitate achievement

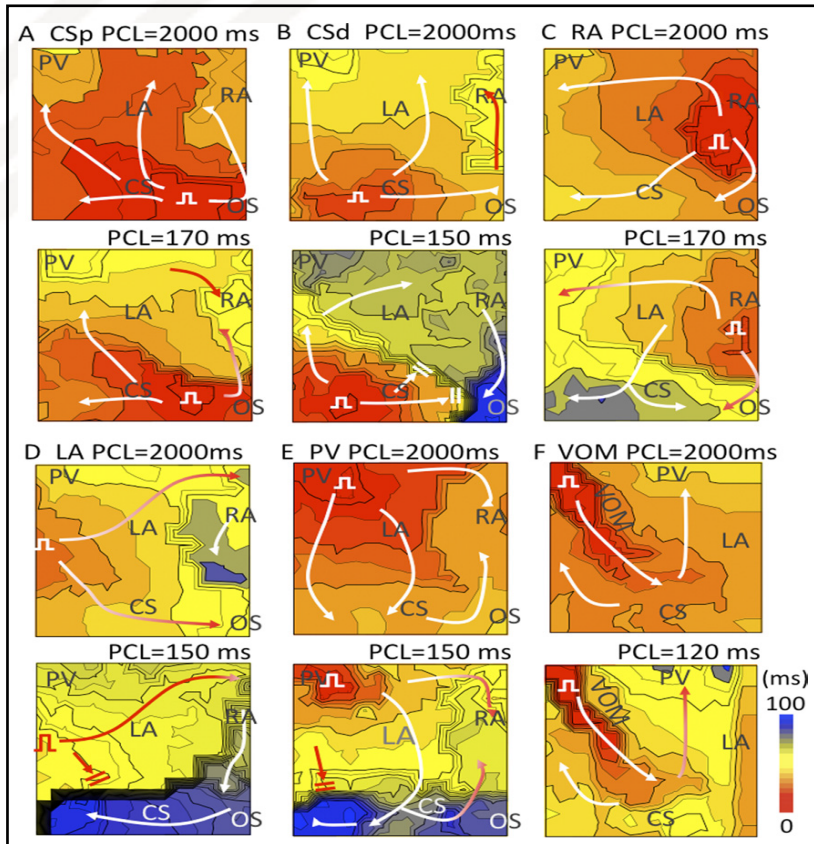
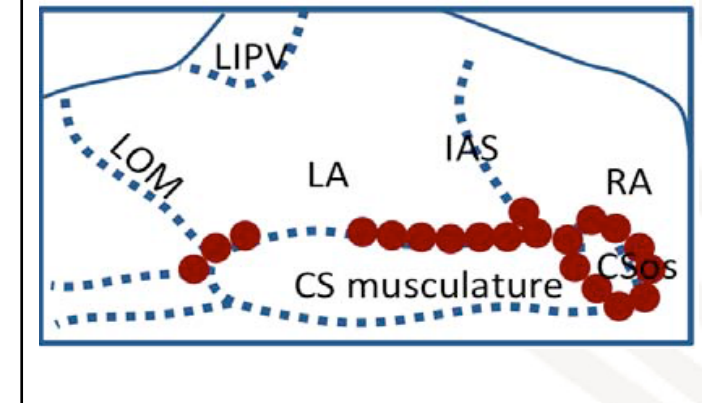
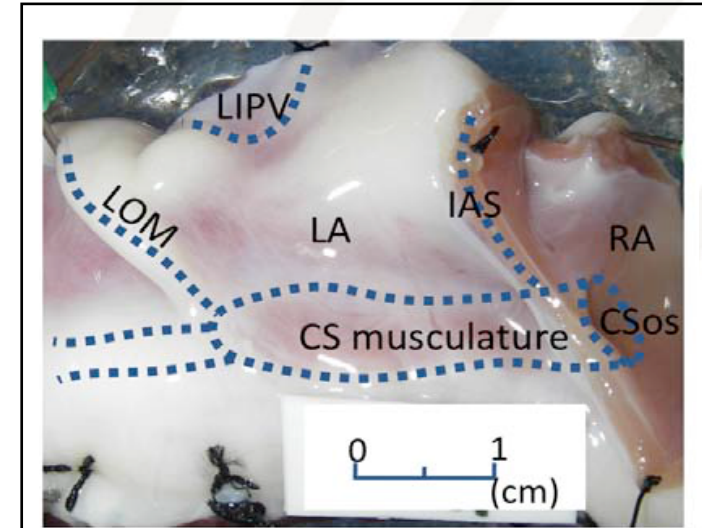
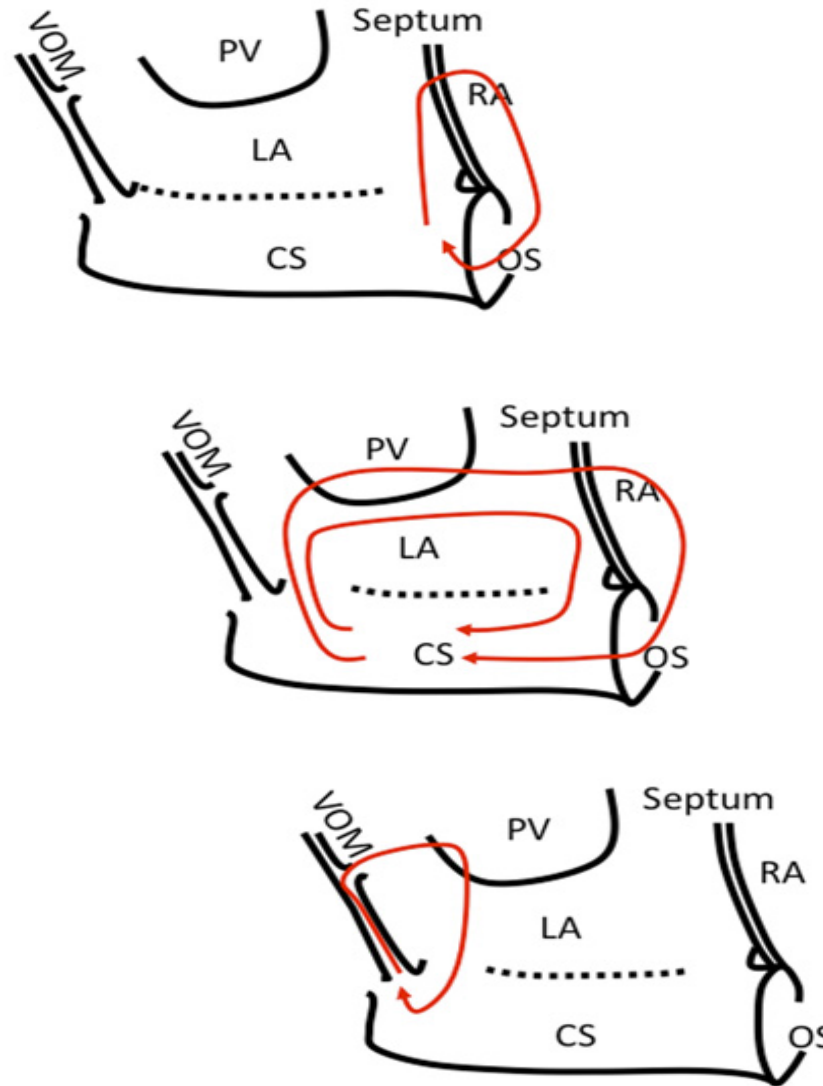
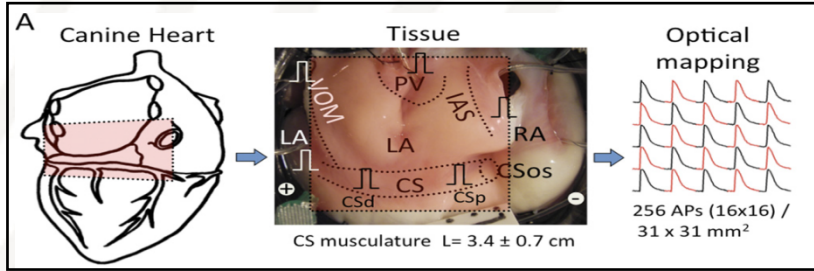
- Multicentric study
- "Marshall"-Plan ablation Vs. PV isolation (262pts)
- Single procedure, off AAD
- 24m f/u
- *NCT: 04681872*





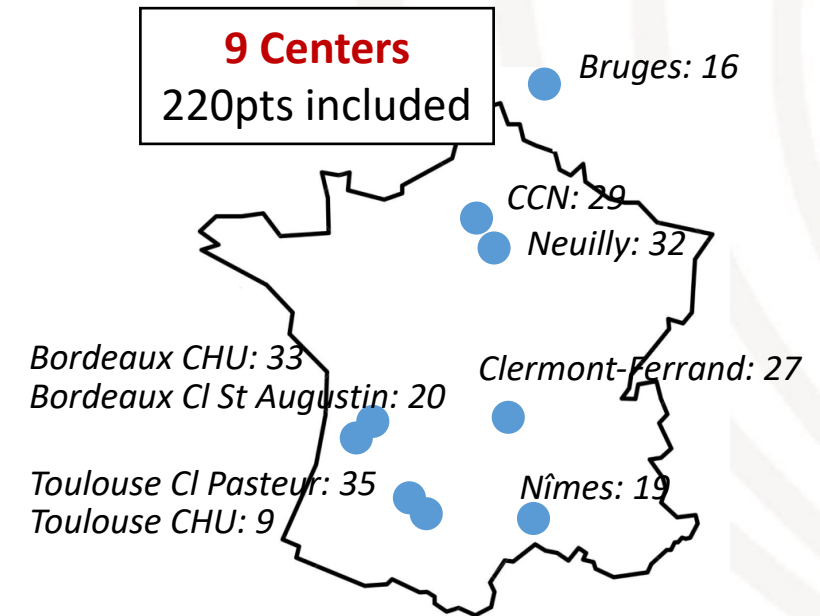
	No (%)			P value
	Marshall-Plan (n=59)	PV Isolation (n=59)	Absolute difference	
Primary outcome				
➤ Freedom from any arrhythmias (AF/AT), after a single ablation procedure with or without antiarrhythmic medication at 12 mo	51 (86.4)	39 (66.1)	+ 12 (+ 20.3)	.012
Secondary outcomes				
➤ Freedom from any arrhythmias (AF/AT), after a single ablation procedure without antiarrhythmic medication at 12 mo	50 (84.7)	39 (66.1)	+ 11 (+ 18.6)	.022
➤ Freedom from any arrhythmias (AF/AT), after 1 or 2 ablation procedures without antiarrhythmic medication at 12 mo	51 (86.4)	39 (66.1)	+ 12 (+ 20.3)	.011
➤ Freedom from any arrhythmias (AF/AT), after 1 or 2 ablation procedures with or without antiarrhythmic medication at 12 mo	52 (88.1)	39 (66.1)	+ 13 (+ 22.0)	.005

ROLE OF THESE EPICARDIAL STRUCTURE IN AF MECHANISM





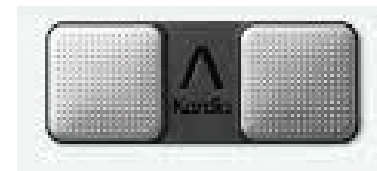
- Multicentric study
- "Marshall"-Plan ablation Vs. PV isolation (262pts)
- Single procedure, off AAD
- 24m f/u
- *NCT: 04681872*



- Prospective, randomized, parallel-group, monocentric trial of superiority
- Intention-to-treat analysis
- **Inclusion criteria**
 - >18 yo
 - Symptomatic Persistent AF (>1month) with documented ECG
- **Primary endpoint**
 - Freedom from **any arrhythmias** (AF/AT), after a **single** ablation procedure **without** antiarrhythmic medication at **12 months**

- **Follow-up protocol**

- Weekly ECG transmission with Alivecor® system
- Visit at 3, 6, 9 and 12 months with 24 hours holter monitoring, TTE at 3 and 12 months





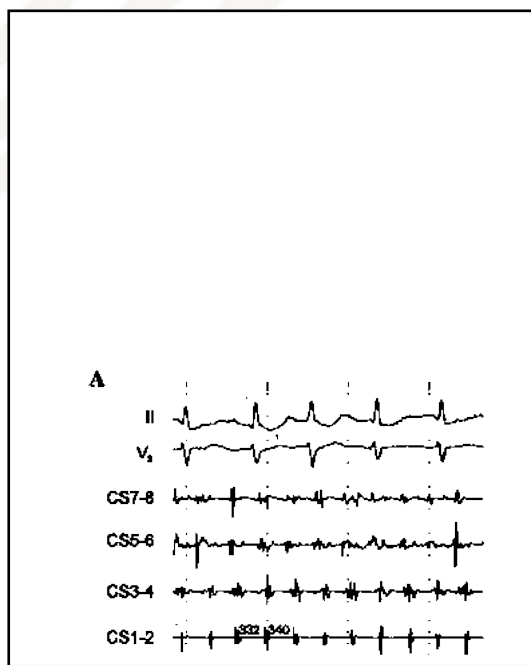
Characteristics	Marshall-Plan N=60	PV Isolation N=60	p-value
Demographics			
Age (y)	66±8	65±8	0.21
Gender			0.47
F	12 (20%)	9 (15%)	
M	48 (80%)	51 (85%)	
CHA ₂ DS ₂ -Vasc	2±1	2±1	0.04
Hypertension	36 (60%)	25 (42%)	0.04
Diabetes	9 (15%)	3 (5%)	0.07
Previous Stroke	5 (8.3%)	2 (3.3%)	0.44
History of Amiodarone	54 (90%)	49 (82%)	0.19
LVEF (%)	51±12	56±10	0.12
SHD	6 (10%)	6 (10%)	0.99
AF Characteristics			
Maximum AF length (m)	10±18	7±6	0.86
Current AF length (m)	9±19	6±7	0.77
Long-standing AF>1y	11 (18%)	11 (18%)	1.00
History of DCC	1±1	1±1	0.32
Rhythm at inclusion			
AF	32 (53%)	39 (65%)	0.19
SR	28 (47%)	21 (35%)	
Left atrial volume (ml)	187±53	192±53	0.31

	Marshall-Plan N=59	PV isolation N=59	p-value
Rhythm in EP lab			0.19
AF	32 (54%)	39 (66%)	
SR	27 (46%)	20 (34%)	
LAA cycle length (if AF) (ms)	206±138	176±26	0.88
LA volume (ml)	182±52	192±53	0.22
PVI	59	59	1.00
Total RF LPV (min)	9.5±4.3	13.0±3.7	0.001
Total RF RPV (min)	13.4±5.1	15.8±5.5	0.012
Total RF PVs (min)	23±8	29±8	0.001
VOM-OH	57 (97%)	-	-
OH volume (ml)	10±2	-	-
Time for OH (min)	22.5±15.4	-	-
Xray time for OH (min)	6.3±6±8	-	-
Mitral isthmus line attempt	56 (95%)	-	-
Mitral isthmus block (% per attempt; %total)	55 (98%; 93%)	-	-
RF mitral line (min)	7.2±6.2	-	-
Posterior wall line attempt	56 (95%)	-	-
Posterior wall block (% per attempt; %total)	54 (96%; 92%)	-	-
Roof line block (% per attempt; %total)	30 (54%; 51%)	-	-
RF roof line (min)	6.3±4.4	-	-
Floor line attempt	26 (46%)	-	-
Floor line block (% per attempt; %total)	24 (92%; 43%)	-	-
RF floor line (min)	4.8±2.6	-	-
CTI line attempt	56 (95%)	-	-
CTI line block(% per attempt; %total)	55 (93%; 98%)	-	-
RF CTI line (min)	5.9±4.5	-	-
AF termination	1 (2%)	3 (5%)	0.43
Ablation set complete	52 (88%)	59 (100%)	0.058
Total Procedure time (min)	157±53	125±31	0.001
Total RF time (min)	36.8±16.0	29.6±8.1	0.001
Total X-Ray time (min)	21±16	11±6	0.001

- Electrograms with 2 or more deflections and/or baseline perturbation with continuous activity over 10sec

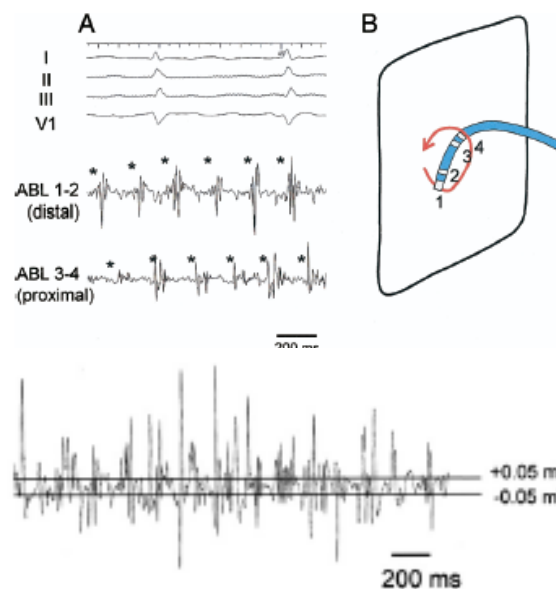
And/or

- Electrograms with a mean CL < 120ms over 10sec.



Nademane et al. JACC 2004

- Temporal activation gradient
- EGMs with continuous electrical activity
- Fast activities



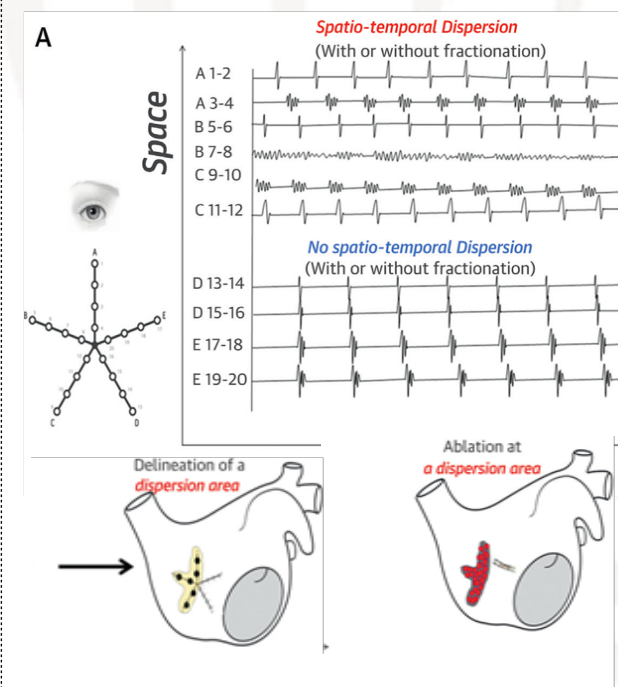
Takahashi Y et al. JACC 2008

- Classification of Fractionated atrial electrograms (CFAE AF trial)

Grade	Electrogram Criteria
1	Uninterrupted fractionated activity Fractionated activity (defined as continuous deflections without pause at the isoelectric line for ≥ 70 ms) occupying $\geq 70\%$ of sample, and at least 1 uninterrupted episode of fractionated activity lasting ≥ 1 second
2	Interrupted fractionated activity Fractionated activity occupying $\geq 70\%$ of sample
3	Intermittent fractionated activity Fractionated activity occupying 30–70% of sample
4	Complex electrograms Discrete electrograms (<70 ms) and complex (≥ 5 direction changes), with any fractionated activity occupying <30% of sample (otherwise grade 3)
5	Normal electrogram Discrete electrograms (<70 ms) and simple (≤ 4 direction changes)
6	Scar No discernible deflections

Hunter RJ et al. CirCAE 2011

- Spacio-temporal dispersion:** Dispersion areas were defined as clusters of electrograms, either fractionated or nonfractionated, that displayed interelectrode time and space dispersion at a minimum of 3 adjacent bipoles such that activation spread over all the AFCL



Seitz J et al. JACC 2017

Volume 101
Number 4
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... perhaps the only way to prevent the atrium from fibrillating is to **interrupt *all* of the potential pathways for atrial macroreentrant circuits** that have been identified by intraoperative mapping either experimentally or clinically, and our eventual surgical approach was based on this principle...

