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**When local impedance meets contact force: preliminary experience from the CHARISMA registry**

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1 **Title: When local impedance meets contact force: preliminary experience from the**  
2 **CHARISMA registry**

3

4 **Short Title:** Correlation between local impedance and contact force during PVI in AF

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6 Francesco Solimene<sup>1</sup>, MD; Valerio De Sanctis<sup>2</sup>, MD; Ruggero Maggio<sup>3</sup>, MD; Maurizio  
7 Malacrida<sup>4</sup>, MSc; Luca Segreti<sup>5</sup>, MD; Matteo Anselmino<sup>6</sup>, MD; Vincenzo Schillaci<sup>1</sup>, MD;  
8 Massimo Mantica<sup>2</sup>, MD; Marco Scaglione<sup>7</sup>, MD; Antonio Dello Russo<sup>8</sup>, MD; Filippo Maria  
9 Cauti<sup>9</sup>, MD; Gianluca Zingarini<sup>10</sup>, MD; Claudio Pandozi<sup>11</sup>, MD; Marco Cavaiani<sup>4</sup>, MSc; Anna  
10 Ferraro<sup>3</sup>, MD; Giampiero Maglia<sup>12</sup>, MD; Giuseppe Stabile<sup>1, 13</sup>, MD.

11

12 <sup>1</sup> Clinica Montevergine, Mercogliano, Avellino, Italy

13 <sup>2</sup> Istituto Clinico Sant'Ambrogio, Milan, Italy

14 <sup>3</sup> Infermi Hospital, Rivoli, Italy

15 <sup>4</sup> Boston Scientific, Milan, Italy

16 <sup>5</sup> Second Division of Cardiology, Cardiac-Thoracic-Vascular Department, New Santa Chiara  
17 Hospital, Azienda Ospedaliero Universitaria Pisana, Pisa, Italy

18 <sup>6</sup> Division of Cardiology, "Città della Salute e della Scienza di Torino" Hospital, Department of  
19 Medical Sciences, University of Turin, Turin

20 <sup>7</sup> Cardinal Massaia Hospital, Asti, Italy

21 <sup>8</sup> Cardiology and Arrhythmology Clinic, Marche Polytechnic University, Ancona, Italy

22 <sup>9</sup> Arrhythmology Unit, Ospedale San Giovanni Calibita, Fatebenefratelli, Isola Tiberina, Rome,  
23 Italy

24 <sup>10</sup> Ospedale Santa Maria della Misericordia, Perugia

25 <sup>11</sup> Division of Cardiology, San Filippo Neri Hospital, Rome, Italy

26 <sup>12</sup> Azienda Ospedaliera Pugliese Ciaccio, Catanzaro, Italy

27 <sup>13</sup> Anthea Hospital, Bari, Italy

28

29

30 **Corresponding author:**

31 Valerio De Sanctis,

32 Department of Cardiac Electrophysiology and Pacing,

33 Istituto Clinico Sant'Ambrogio,

34 Via Luigi Giuseppe Faravelli, 16, 20149 Milano, Italy.

35 [valeriodesanctis@yahoo.it](mailto:valeriodesanctis@yahoo.it)

36

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41 Formation, PVI

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46

47 **Abstract**

48 **Purpose.** Highly localized impedance (LI) measurements during atrial fibrillation (AF) ablation  
49 have emerged as a viable real-time indicator of tissue characteristics and the consequent  
50 durability of the lesions created. We investigated the impact of catheter-tissue contact force (CF)  
51 on LI behavior during pulmonary vein isolation (PVI).

52 **Methods:** Forty-five consecutive patients of the CHARISMA registry undergoing *de novo* AF  
53 radiofrequency (RF) catheter ablation with a novel open-irrigated-tip catheter endowed with CF  
54 and LI measurement capabilities (Stablepoint™ catheter, Boston Scientific) were included.

55 **Results:** A total of 2895 point-by-point RF applications were analyzed (RF delivery time  
56 (DT)=8.7±4s, CF=13±8g, LI drop=23±7Ω). All PVs were successfully isolated in an overall  
57 procedure time of 118±34min (fluoroscopy time=13±8min). The magnitude of LI drop weakly  
58 correlated with CF ( $r=0.13$ , 95% confidence interval (CI): 0.09 to 0.16,  $p<0.0001$ ), whereas both  
59 CF and LI drop inversely correlated with DT ( $r=-0.26$ , 95%CI: -0.29 to -0.22,  $p<0.0001$  for CF;  
60  $r=-0.36$ , 95%CI: -0.39 to -0.33,  $p<0.0001$  for LI). For each 10 grams of CF, LI drop markedly  
61 increased from 22.4±7Ω to 24.0±8Ω at 5 to 25g CF intervals (5-14 grams of CF vs 15-24 grams  
62 of CF,  $p<0.0001$ ), whereas it showed smooth transition over 25g (24.8±7Ω at ≥25g CF intervals,  
63  $p=0.0606$  vs 15-24 g of CF). No major complications occurred during the procedures or within  
64 30 days.

65 **Conclusions:** CF significantly affects LI drop and probable consequent lesion formation during  
66 RF PVI. The benefit of higher contact (>25g) between the catheter and the tissue appears to have  
67 less impact on LI drop.

68

## 69 INTRODUCTION

70 Catheter ablation aimed at pulmonary vein (PV) isolation is the most effective treatment in  
71 patients with atrial fibrillation (AF), and is now recommended as the first-line therapy (1, 2).  
72 Despite acute safety and efficacy, a considerable number of recurrences are observed during  
73 long-term follow-up, mainly as a result of PV reconnection (1-3). Several strategies have been  
74 proposed in order to achieve durable, transmural lesions, thereby improving the efficiency of  
75 catheter ablation (4). Highly localized impedance (LI) measurements during AF ablation have  
76 emerged as a viable real-time indicator of tissue characteristics and the consequent durability of  
77 the lesions created (5-7). A recently released catheter has combined CF detection with LI  
78 assessment in a single catheter-tissue contact force (CF)-LI catheter (8). In swine and *in vitro*,  
79 the addition of LI to CF has provided feedback on both electrical and mechanical loads and  
80 allows the evaluation of tissue resistivity, and thus of the type of tissue with which the catheter is  
81 in contact. It has also provided feedback on whether volumetric tissue heating is inadequate,  
82 sufficient, or excessive. In addition, in a point-by-point workflow with consistent CF, the  
83 visualization of LI significantly reduced RF time (8). We investigated the impact of CF on LI  
84 behavior during PV isolation.

## 85 METHODS

### 86 *Patient population and study design*

87 CHARISMA was a prospective, multi-center cohort study designed to describe Italian clinical  
88 practice regarding the approach to ablation of various arrhythmias. The study complied with the  
89 Declaration of Helsinki, the locally appointed ethics committee approved the research protocol,  
90 and informed consent was obtained from all patients. From January 2021 to July 2021, 45  
91 consecutive patients indicated for AF ablation who were undergoing their first high-resolution

92 mapping and ablation procedure with a novel CF- and LI-featured catheter in 9 Italian centers  
93 were included in our analysis. All patients were followed up at the same hospital, from the time  
94 of first **ablation** to the last follow-up visit.

#### 95 *Ablation procedure*

96 After completion of the baseline evaluation, patients underwent ablation in accordance with  
97 standard clinical practice guidelines [1]. All procedures were performed under conscious  
98 sedation or general anesthesia. Vitamin K antagonist treatment was not interrupted, while non-  
99 Vitamin K anticoagulants were skipped on the morning of the procedure. A decapolar catheter  
100 (e.g. Dynamic XT™, Boston Scientific, Marlborough, MA, USA) was used to cannulate the  
101 coronary sinus. After single or double transseptal punctures under fluoroscopic guidance,  
102 intravenous unfractionated heparin boluses were administered, in order to maintain an activated  
103 clotting time of >300 seconds. **Intracardiac echocardiography probe was not used in any**  
104 **procedure.** The basket mapping catheter (Orion™, Boston Scientific, Marlborough, MA, USA)  
105 and the ablation catheter (Stablepoint™ catheter, Boston Scientific, Marlborough, MA, USA)  
106 were then inserted. **A standard, non-steerable sheath was used.** The Orion™ catheter was used in  
107 combination with the Rhythmia™ HDx mapping system (Rhythmia™, Boston Scientific,  
108 Marlborough, MA, USA) to create a 3-dimensional electro-anatomical voltage and activation  
109 map of the left atrium. Mapping and ablation were primarily carried out in sinus rhythm; in  
110 patients in AF, electrical cardioversion was attempted in order to restore sinus rhythm, at the  
111 beginning of the procedure and before re-mapping. Point-by-point RF delivery was performed in  
112 such a way as to create contiguous ablation spots encircling the PVs. CF settings were at the  
113 individual operator's discretion, within the range of 5 to 40 g. Ablation was guided by the  
114 magnitude and time-course of impedance drop during RF delivery. RF applications were targeted

115 to a minimum LI drop of  $15 \Omega$  within 15 seconds and were stopped when a maximum cutoff LI  
116 drop of  $\geq 40 \Omega$  was observed. We aimed to reach an LI drop of 20–30 Ohms, on the basis of  
117 previous experimental data (8). Radiofrequency energy was applied in the power controlled  
118 mode (45–50 W) with a temperature limit of  $43^\circ\text{C}$ . The irrigation rate was 30 ml/min during  
119 applications and 2 ml/ min during mapping. A normal saline solution (NaCl 0.9%) was used. The  
120 recommended maximum distance between adjacent ablation spots (center-to-center) was  $\leq 6$  mm.  
121 The ablation points were marked automatically with 6 mm diameter, numbered AutoTags<sup>TM</sup>. The  
122 starting impedance, initial CF, LI drop during RF, and average force applied were recorded. The  
123 endpoint of ablation was PV isolation, as assessed on the basis of entry and exit block by means  
124 of the 64-pole Orion<sup>TM</sup> catheter placed sequentially in each of the PVs. In the absence of first-  
125 pass PV isolation (i.e. no isolation upon completion of the encirclement of ipsilateral veins), PV  
126 isolation was accomplished by means of additional RF applications at the investigator's  
127 discretion.

### 128 *Local impedance*

129 A 3-electrode method with separate circuits for field creation and measurement was used to  
130 measure LI. As previously described, non-stimulatory alternating current was delivered between  
131 the tip electrode and the proximal ring; voltage was passively measured between the tip electrode  
132 and the distal ring (9). As the catheter used does not have mini-electrodes, the resulting voltage  
133 was measured from the catheter tip. Impedance was calculated by dividing the voltage by the  
134 stimulatory current. To measure the baseline reference impedance of the blood pool, once the  
135 reference map had been completed, the ablation catheter was positioned in the blood pool for 10  
136 s and the value was calculated when no EGM recordings from the ablation catheter were present.  
137 Baseline tissue impedance and impedance drop for each ablation lesion were measured. To

138 analyze the impedance information, the isolation line around each pair of PVs was divided into  
139 seven distinct sections (Figure 1A) in accordance with the literature (10). Videos of the ablation  
140 procedures were exported from the mapping system, to display the procedure in real time. RF  
141 current applications were then retrospectively analyzed.

#### 142 *Contact force*

143 The ablation catheter used in the current study has the ability to measure both real-time LI  
144 calculated from a local electric field generated at the tip of the catheter, and CF. The force  
145 applied to the tip electrodes transferred to inductive sensors via a spring. The signal change  
146 measured by the inductive sensors is then converted to a 3-dimensional force vector by means of  
147 known spring dynamics. The target CF was 5-40 g, at the operator's discretion. We collected the  
148 following data on each first-pass ablation point: power, minimum CF, maximum CF, mean CF,  
149 duration of application, baseline LI and LI drop. In addition, the CF range during the applications  
150 was calculated by subtracting the minimum CF from the maximum CF of the ablation point. All  
151 numbered AutoTag™ points were exported from the system for off-line analysis. An example of  
152 visualization of CF values and the DirectSense™ tool on the Rhythmia™ mapping system  
153 during ablation is depicted in figure 1B, C.

#### 154 *Follow-up*

155 Complications were reported on the case report form and collected during follow-up. After  
156 ablation, anticoagulation and antiarrhythmic drugs therapy were continued. At 3 months,  
157 anticoagulation was continued according to the stroke risk, whereas antiarrhythmic drugs were  
158 continued at the discretion of the treating physician. Clinical evaluation and ECG were  
159 performed at 1, 3, 6, and 12 months. Holter ECG was performed at 3, 6 and 12 months post-



160 ablation or in the case of symptoms. For the purpose of this study, data were collected during the  
161 index procedure and during an ambulatory visit 30 days after the procedure.

### 162 ***Statistical analysis***

163 Descriptive statistics are reported as means±SD for normally distributed continuous variables, or  
164 medians with 25th to 75th percentiles in the case of skewed distribution. Normality of  
165 distribution was tested by means of the non-parametric Kolmogorov–Smirnov test. Differences  
166 between mean data were compared by means of a t-test for Gaussian variables, and the F-test  
167 was used to check the hypothesis of equality of variance. The Mann-Whitney non-parametric test  
168 was used to compare non-Gaussian variables. Differences in proportions were compared by  
169 applying  $\chi^2$  analysis or Fisher’s exact test, as appropriate. Linear regression analysis was  
170 performed to determine relationships between LI drop, CF and RF delivery time (DT). A *p* value  
171 <0.05 was considered significant for all tests. All statistical analyses were performed with  
172 STATISTICA software, version 7.1 (StatSoft, Inc., Tulsa, OK).

## 173 **RESULTS**

### 174 ***Study population and procedural parameters***

175 The demographic and procedural data of the 45 consecutive *de novo* PV isolation patients are  
176 reported in Table 1. Almost two-thirds of the patients suffered from paroxysmal AF (n=26, 58%)  
177 whereas 19 (42%) had a history of persistent AF. The mean procedure duration and fluoroscopy  
178 times were 107.4±39 min and 11.1±8 min, respectively. A total of 3196 RF applications were  
179 delivered, with a mean number of 64±31 ablation spots during a mean RF delivery time of 8.7±4  
180 sec, without any steam popping.

### 181 ***Local tissue impedance values***

182 High-quality data were available on 2895 (91%) RF applications performed around PVs. The  
183 baseline LI was  $157.9\pm 17\Omega$  prior to ablation and  $136.9\pm 14\Omega$  after ablation ( $p<0.0001$ , absolute  
184 LI drop of  $23.0\pm 7\Omega$ ) with an LI drop rate of  $3.5\pm 2\Omega/s$ . The mean blood-pool impedance was  
185  $152.7\pm 10\Omega$  ( $p<0.0001$  vs baseline LI). The magnitude of the impedance drop was predicted by  
186 the baseline LI (correlation coefficient  $r=0.61$ , 95% confidence interval (CI): 0.59–0.63,  
187  $p<0.0001$ ). Regarding AF type, no difference in baseline LI was found ( $158\pm 17\Omega$  for paroxysmal  
188 AF vs  $157.9\pm 17\Omega$   $157.7\pm 17\Omega$  for persistent AF,  $p=0.3878$ ), whereas LI drops were larger in  
189 paroxysmal AF cases ( $23.3\pm 7\Omega$ ) than in persistent AF cases ( $22.7\pm 7\Omega$ ,  $p=0.0097$ ). On  
190 considering the underlying rhythm, no differences were found in terms of either baseline LI or LI  
191 drop (baseline LI:  $157.2\pm 17\Omega$  for sinus rhythm vs  $158.4\pm 17\Omega$  for AF,  $p=0.0518$ ; LI drop:  
192  $22.8\pm 7\Omega$  for sinus rhythm vs  $22.9\pm 7\Omega$  for AF,  $p=0.8606$ ).

### 193 *Correlation between local impedance and key procedural parameters*

194 The mean RF delivery time was  $8.7\pm 4s$  and the mean CF was  $13.0\pm 8g$ . On assessing the various  
195 key ablation parameters, the magnitude of LI drop proved to be weakly correlated with CF  
196 ( $r=0.13$ , 95%CI: 0.09 to 0.16,  $p<0.0001$ ) whereas both CF and LI drop inversely correlated with  
197 DT ( $r=-0.26$ , 95%CI: -0.29 to -0.22,  $p<0.0001$  for CF;  $r=-0.36$ , 95%CI: -0.39 to -0.33,  $p<0.0001$   
198 for LI). Figure 2 (A to C) shows the resulting mean DT stratified by CF values and LI drop  
199 values. For each 10 grams of CF, LI drops markedly increased from  $22.4\pm 7\Omega$  to  $24.0\pm 8\Omega$  at 5 to  
200 25g CF intervals (5-14 grams of CF vs 15-24 grams of CF,  $p<0.0001$ ), whereas it showed a  
201 smooth transition above 25g ( $24.8\pm 7\Omega$  at  $\geq 25g$  CF intervals,  $p=0.0606$  vs 15-24 g of CF)  
202 (Supplementary figure 1). There was a correlation between shorter DT and larger drop in LI:  
203  $27.2\pm 8\Omega$  at 0-5 seconds of DT interval vs  $22.8\pm 7\Omega$  at 6-10 seconds of DT interval vs  $19.7\pm 6\Omega$  at

204 >10 seconds of DT interval (all comparisons  $p<0.0001$ ). Details of the relationships among the  
205 three parameters are reported in figure 3.

### 206 *Characterization of pulmonary vein location sites*

207 Of the 2895 RF applications, 1544 (53.3%) were sited on the RPVs and 1351 (46.7%) on the  
208 LPVs. Baseline impedance was homogenous across the various location sites ( $158.5\pm 17\Omega$  at  
209 LPVs vs  $157.4\pm 17\Omega$  at RPVs,  $p=0.0822$ ;  $157.2\pm 17\Omega$  at anterior sites vs  $159.4\pm 18\Omega$  at posterior  
210 sites,  $p=0.0643$ ; and  $159.4\pm 17\Omega$  at inferior sites vs  $157.5\pm 15\Omega$  at superior sites,  $p=0.1028$ ). LI  
211 drop was higher at anterior sites ( $23.4\pm 7\Omega$  vs  $22.8\pm 8\Omega$  at posterior sites,  $p=0.029$ ) and at inferior  
212 sites ( $23.5\pm 7\Omega$  vs  $22.5\pm 7\Omega$  at superior sites,  $p=0.0447$ ), whereas it was similar between RPVs  
213 and LPVs ( $22.8\pm 7\Omega$  in RPV pairs vs  $23.2\pm 7\Omega$  at LPVs,  $p=0.0565$ ) (Figure 4A, 4B). RF delivery  
214 time was longer at superior sites ( $8.9\pm 4$  sec vs  $8.4\pm 4$  sec at inferior sites,  $p=0.0437$ ) and in RPV  
215 pairs ( $8.8\pm 4$  sec vs  $8.6\pm 4$  sec in LPV pairs,  $p=0.0334$ ), whereas no differences were found  
216 between posterior and anterior sites ( $8.6\pm 4$  sec at posterior sites vs  $8.7\pm 4$  sec at anterior sites,  
217  $p=0.7824$ ) (Figure 4C). CF values were higher in LPV pairs ( $13.5\pm 8$  g vs  $12.5\pm 7$  g in RPV pairs,  
218  $p=0.0025$ ), whereas no differences were found between superior and inferior sites ( $13.2\pm 8$  g vs  
219  $12.6\pm 8$  g,  $p=0.2358$ ) or between posterior and anterior sites ( $12.7\pm 7$  g vs  $13.2\pm 8$  g,  $p=0.5062$ )  
220 (Figure 4D). Details of the distribution of RF applications, CF values, LI drops and RF delivery  
221 times, according to location sites, are reported in Figure 5 and Supplementary Table 1. Details of  
222 baseline and ablated tissue impedance values are reported in Supplementary Figure 2 and  
223 Supplementary Table 1.

### 224 *First pass isolation and acute outcome*

225 No steam pops or major complications, including atrio-esophageal fistula or tamponade, were  
226 reported during or after the procedures. **In our series a total 169 PVs (94%) were isolated at first**

227 pass ablation, resulting in 40 (89%) patients who had a first pass isolation, whereas 11 residual  
228 gaps in 5 (11%) patients were observed after initial encirclement and required additional RF  
229 applications. LI drop values were larger and CF values were higher where first pass isolation was  
230 achieved (LI drop:  $23.1 \pm 7 \Omega$  at successful sites vs  $16.8 \pm 3 \Omega$  at unsuccessful sites,  $p < 0.0001$ ; CF:  
231  $13 \pm 8 \text{g}$  at successful sites vs  $10.2 \pm 6 \text{g}$  at unsuccessful sites,  $p = 0.0207$ , respectively). At the end of  
232 the procedures, all PVs had been successfully isolated in all study patients. Minor complications  
233 were reported in 3 patients (6.6%) after the procedure: one pericarditis with mild pericardial  
234 effusion, and groin hematomas in 2 patients. Conservative treatment and medical therapy were  
235 effective in all cases, without prolongation of hospital stay.

## 236 **DISCUSSION**

### 237 *Main findings*

238 In this single-arm prospective study, we performed AF catheter ablation by means of a novel  
239 ablation catheter with integrated CF- and LI-sensing capabilities. The ablation strategy, which  
240 was guided by LI information, had a 100% acute procedural effectiveness rate, without causing  
241 any steam pops or major complications. CF significantly impacted on effective lesion formation  
242 during RF PV isolation. The use of higher than 25g contact between the catheter and the tissue  
243 proved to have less impact on LI drop. The inverse correlation of both CF and LI drop with RF  
244 DT indicates that a significant reduction in RF time can be achieved at 45-50 W power in a  
245 point-by-point workflow when LI guidance is combined with CF. These points reflect the value  
246 of LI plus CF in discerning both mechanical contact and electrical coupling, thereby enabling  
247 safe and effective lesions to be created.

### 248 *Ablation guided by local impedance and contact force*

249 The use of highly localized impedance measurements to provide insight into tissue  
250 characteristics and their real-time evaluation seems to be helpful in order to precisely assess the  
251 electrical contact of the catheter and tip stability and to serve as a viable real-time indicator of  
252 tissue characteristics and durability of the lesions created (5-7; 11). Two commercially available  
253 catheters capable of recording LI are currently available. The IntellaNav MiFi OI catheter  
254 (Boston Scientific) generates LI measurements through mini-electrodes on the tip of the ablation  
255 catheter, the maximum value being reported within a three-dimensional mapping environment  
256 (Rhythmia; Boston Scientific). A recently released StablePoint catheter (Boston Scientific)  
257 incorporates CF-sensing capability in addition to LI (8). The ablation strategy for PV isolation  
258 guided by LI technology has proved safe and effective, resulting in a very low rate of AF  
259 recurrence over 1-year follow-up (7). However, as the dedicated ablation catheter (IntellaNAV  
260 Mifi OI, Boston Scientific) used in these studies was not able to collect data on CF sensing, it  
261 was not possible to compare CF and impedance measurements.

262 It is well recognized that, when RF energy is applied, CF is one of the variables, in addition to  
263 catheter stability, power output, temperature, and duration of RF output, that impact on lesion  
264 size and transmuralty (4). CF-guided RF catheter ablation has been associated with a  
265 significantly greater AF/atrial tachycardia-free survival benefit than non-CF-guided ablation in  
266 patients with paroxysmal AF rather than persistent AF. In addition, the CF-guided ablation  
267 strategy also reduced procedure time, fluoroscopy time and RF time, though it had no distinct  
268 effect on the alleviation of procedure-related complications (12). Adding CF sensing to the LI-  
269 sensing technology has the potential to further increase the efficiency of LI-guided catheter  
270 ablation. Indeed, we found that CF significantly impacted on effective lesion formation during  
271 RF PV isolation. However, the benefit of higher than 25g contact between the catheter and the

272 tissue had less impact on the increase in LI drop. Our findings may have relevant implications in  
273 the clinical setting: 1) good catheter-tissue contact improves the drop in LI and shortens the time  
274 needed to achieve it; 2) the lack of benefit of a CF value of above 25 g might avoid excessive  
275 catheter pressure and potential complications. Similar data have already been reported with other  
276 CF-sensing technologies (10, 13); 3) the CF value may help to differentiate the LI value of the  
277 blood pool from that of diseased tissue. Indeed, both the blood pool and diseased tissue display  
278 lower LI values than healthy tissue (5,14). Of note, the magnitude of the mean LI drop observed  
279 in our study ( $23.0 \pm 7 \Omega$ ) was significantly higher than that reported with previous LI technology  
280 (IntellaNAV Mifi OI, Boston Scientific) by other authors: Segreti et al. (5),  $14 \pm 8 \Omega$ ; Das et al.  
281 (6),  $19.8 \pm 11.1 \Omega$ , and Solimene et al. (7),  $13 \pm 8 \Omega$ . To date, only one pilot study, which used the  
282 StablePoint™ ablation catheter (15), showed that a local impedance drop  $> 21.8$  Ohms on the  
283 anterior wall and  $> 18.3$  Ohms on the posterior wall significantly increased the probability of  
284 creating a successful lesion. The CF-LI catheter does not have microelectrodes; instead, its distal  
285 tip serves as the return pole of the LI circuit. The larger electrical field created gives rise to CF-  
286 LI values that are typically 40%–50% greater than those measured by the non-CF-LI catheter  
287 (16). Further studies will therefore be required in order to determine the magnitude of LI drop  
288 that predicts acute PV segment conduction block.

### 289 ***Right power, right duration***

290 Winkle et al. first showed that AF ablations can be performed at 45–50 W for short durations  
291 with very low complication rates. High-power, short-duration ablations have the potential to  
292 shorten procedural and total RF times and to create more localized and durable lesions (17). In  
293 addition, high-power short-duration RF ablation has proved able to significantly shorten  
294 procedure time, fluoroscopy time, left atrial dwell time, and RF ablation time in comparison with

295 a conventional approach, with no difference in safety outcomes between the two groups (18).  
296 When high-power short-duration ablation is performed, several parameters can indicate that a  
297 lesion has been formed and that no further ablation is needed, thereby avoiding ablation for  
298 longer than needed to selectively destroy the target tissue. These parameters include: monitoring  
299 the loss of pacing capture during RF delivery, observing a drop in impedance, and following  
300 such metrics of lesion formation as the Lesion Size Index or the Ablation Index. (19-21). Our  
301 findings showed an inverse correlation of both CF and LI drop with DT, together with a  
302 significant reduction in RF time at 45-50 W power in a point-by-point workflow. This reflects  
303 the value of LI plus CF in discerning both mechanical contact and electrical coupling, thereby  
304 enabling safe and effective lesions to be created.

### 305 *Limitations*

306 This investigation focused on the effect of each single RF application, and no data on medium-  
307 and long-term clinical outcomes were available. Impedance drop was used to assess lesion  
308 formation; however, it is only a surrogate and could be affected by several factors. **The LI values  
309 that we used were empirically chosen. However, they were based on our previous experience in  
310 clinical practice, in which they had allowed us to achieve considerable clinical success. Further  
311 studies are required to identify the best workflow and targeted parameters also for achieving  
312 long-term success.** The effect of using a steerable sheath during ablation may need further  
313 investigation. **Lastly, esophageal temperature monitoring was not performed. However, in our  
314 preliminary experience, applying this procedural workflow, no steam pops or major  
315 complications, including atrio-esophageal fistula or tamponade, occurred during or after the  
316 procedures.**

### 317 **CONCLUSIONS**

318 CF significantly impacts on effective lesion formation during RF PVI. The benefit of higher than  
319 25g contact between the catheter and the tissue appears to have less impact on LI drop.

320

## 321 **DECLARATIONS**

322 **Funding:** No extramural funding was used to support this work.

323 **Conflicts of interest/Competing Interests:** M. Malacrida, M. Cavaiani are employees of Boston  
324 Scientific. The other authors have no conflicts of interest to declare that are relevant to the  
325 content of this article.

326 **Availability of data and material:** The data underlying this article will be shared on reasonable  
327 request to the corresponding author.

328 **Code availability:** Not applicable.

329 **Ethics approval:** This study was performed in line with the principles of the Declaration of  
330 Helsinki. The locally appointed Ethics Committee approved the research protocol.

331 **Consent to participate:** Informed consent was obtained from all individual participants  
332 included in the study.

333 **Consent to publish:** Patients signed informed consent regarding publishing their data.

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441 **Table legends**

442 **Table 1.** Baseline characteristics and procedural parameters

Parameter	n = 45
Age, years	61.6±9
Male Gender, n (%)	28 (62.2)
Indication for ablation:	
• Paroxysmal AF, n (%)	• 26 (57.8)
• Persistent AF, n (%)	• 19 (42.2)
History of atrial flutter/atrial tachycardia, n (%)	6 (13.3)
LVEF, %	55.1±8
Cardiomyopathy, n (%)	15 (33.3)
Hypertension, n (%)	26 (57.8)
Coronary artery disease, n (%)	5 (11.1)
History of heart failure, n (%)	3 (6.7)
COPD, n (%)	2 (4.4)
CKD, n (%)	1 (2.2)
ACE-ARB, n (%)	14 (31.1)
Beta-blockers, n (%)	28 (62.2)
Statin, n (%)	7 (15.6)
Diuretics, n (%)	5 (11.1)
Antiarrhythmics, n (%)	33 (73.3)
Procedure duration, min	107.4±39
Fluoroscopy time, min	11.1±8
RFC applications, n	64±31
RFC duration time, sec	8.7±4
Mean Power, W	47.2±3
Complications during the procedure, n (%)	0 (0)
Minor complications	3 (6.6)
• Mild pericardial effusion	• 1
• Groin hematomas	• 2

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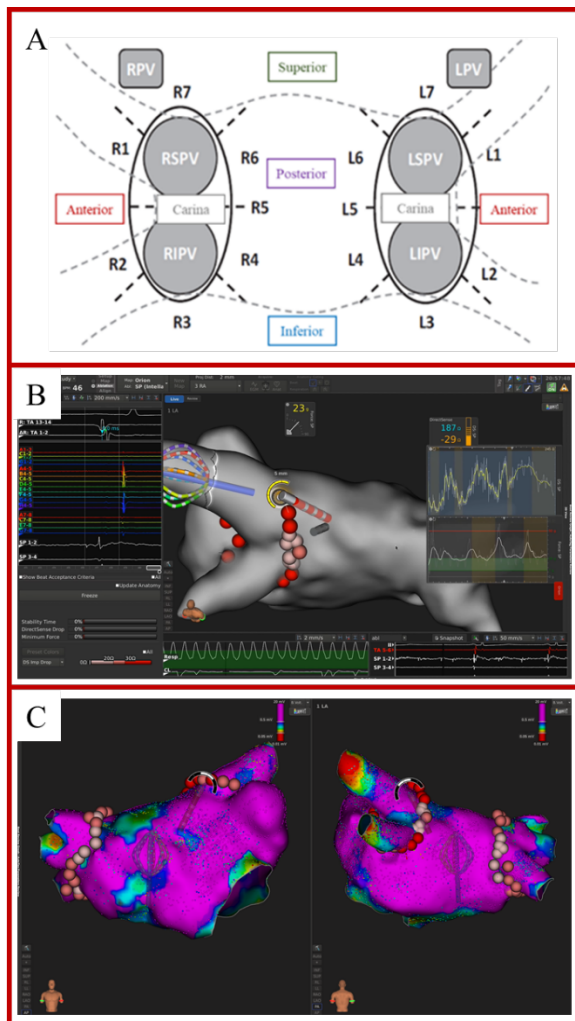
444 AF=atrial fibrillation; PVI=pulmonary vein isolation; LVEF=left ventricular ejection fraction; RFC=radiofrequency

445 catheter; COPD=chronic obstructive pulmonary disease; CKD=chronic kidney disease; ACE=angiotensin-

446 converting enzyme; ARB=angiotensin receptor blocker.

447 **Figure legends**

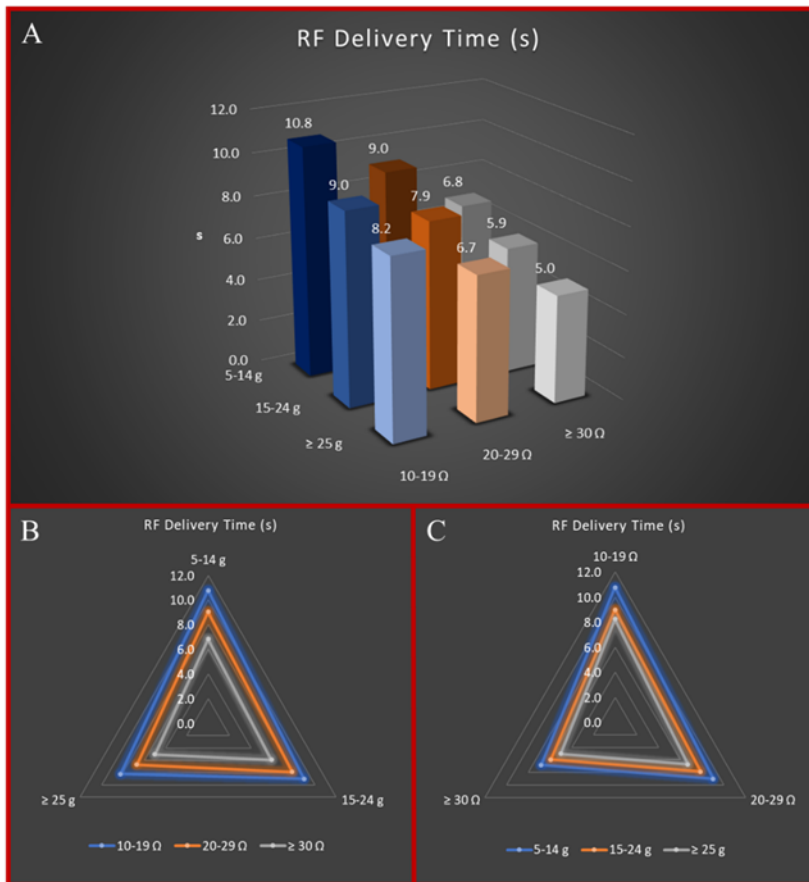
448 **Figure 1. Panel A:** Identification of 7 ablation sites around the right (RPV) and left (LPV) pairs of pulmonary veins.  
449 Anterior superior: R1, L1; Anterior inferior: R2, L2; Inferior: R3, L3; Posterior inferior: R4, L4; Carina: R5, L5;  
450 Posterior superior: R6, L6; Superior: R7, L7. LIPV = left inferior pulmonary vein; LSPV = left superior pulmonary  
451 vein; RIPV = right inferior pulmonary vein; RSPV = right superior pulmonary vein. **Panel B:** Example of  
452 visualization of CF and DirectSense™ tool on the Rhythmia™ mapping system during ablation. **Panel C:** Point-by-  
453 point RF delivery created contiguous ablation spots encircling the PVs. The maximal inter-lesion distance between  
454 two neighboring lesions was set  $\leq 6$  mm and was automatically measured through the Autotag™ software. CF  
455 settings were at the individual operator's discretion, within the range of 5 to 40 g.



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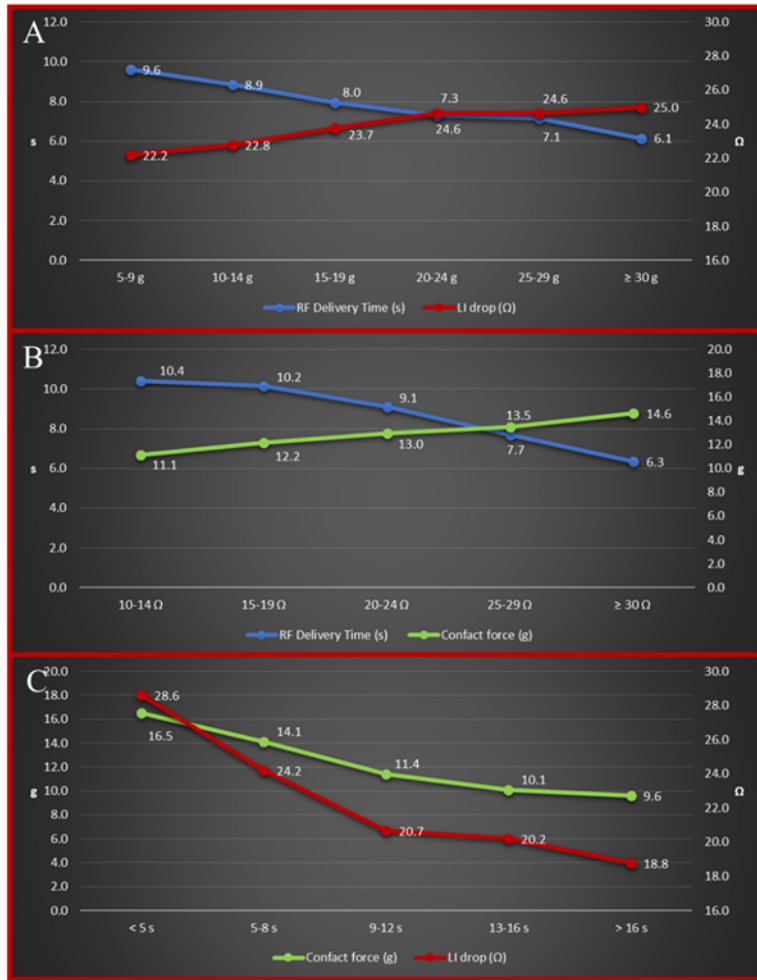
459 **Figure 2. Panel A:** Multidimensional relationship between RF delivery time, CF values and LI drop values. **Panel**  
 460 **B:** Radar plot showing the relationship between RF delivery time and CF values according to different degrees of LI  
 461 drop. This Kiviart chart displays multivariate data (RF delivery time) with values represented on axes starting from  
 462 the same point. The apexes of the Kiviart charts represent different CF intervals (5-14g, 15-24g and  $\geq 25$ g) whereas  
 463 the lines represent different degrees of LI drop (blue line for 10-19 $\Omega$  LI drop values, orange line for 20-29 $\Omega$  LI drop  
 464 values and grey line for LI drop values  $\geq 30\Omega$ ). RF delivery time is represented according to CF and LI drop  
 465 intervals.

466 **Panel C:** Radar plot showing the relationship between RF delivery time and LI drop values according to different  
 467 degrees of CF. This Kiviart chart displays multivariate data (RF delivery time) with values represented on axes  
 468 starting from the same point. The apexes of the Kiviart charts represent different LI drop intervals (10-19 $\Omega$ , 20-29 $\Omega$   
 469 and  $\geq 30\Omega$ ), whereas the lines represent different degrees of CF (blue line for 5-14g CF values, orange line for 15-  
 470 24g CF values and grey line for CF values  $\geq 25$ g). RF delivery time is represented according to CF and LI drop  
 471 intervals.



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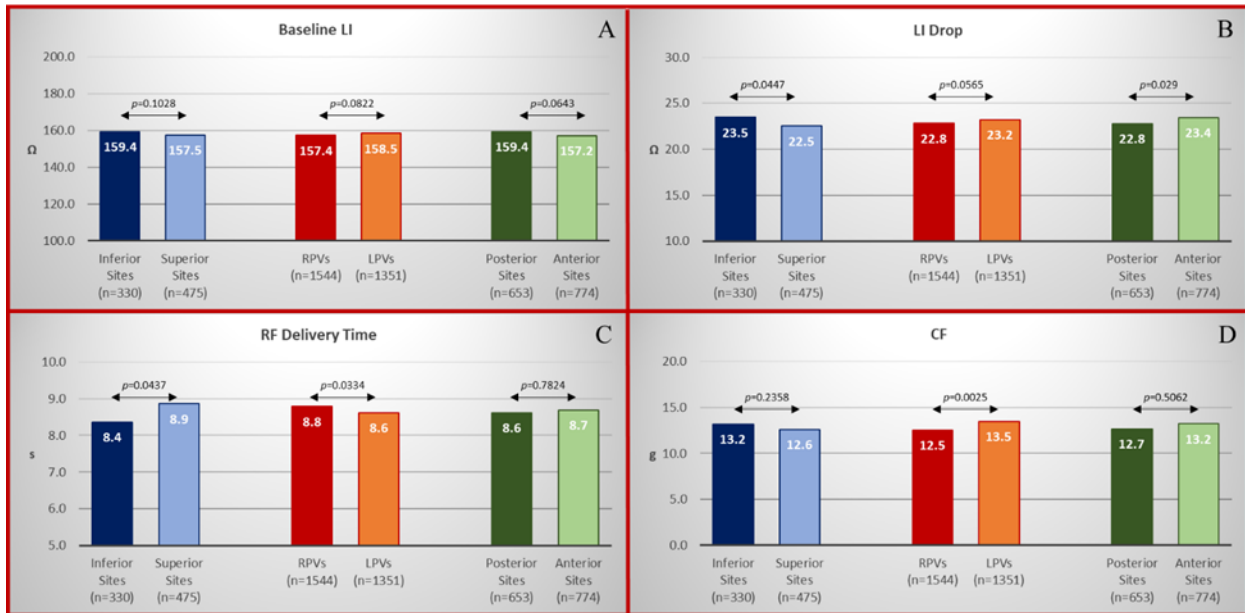
473 **Figure 3.** Details of relationships among the three key parameters: RF delivery time and LI drop according to  
 474 different levels of CF (*panel A*); RF delivery time and CF according to different degrees of LI drop (*panel B*) and  
 475 CF and LI drop according to different values of RF delivery time (*panel C*).  
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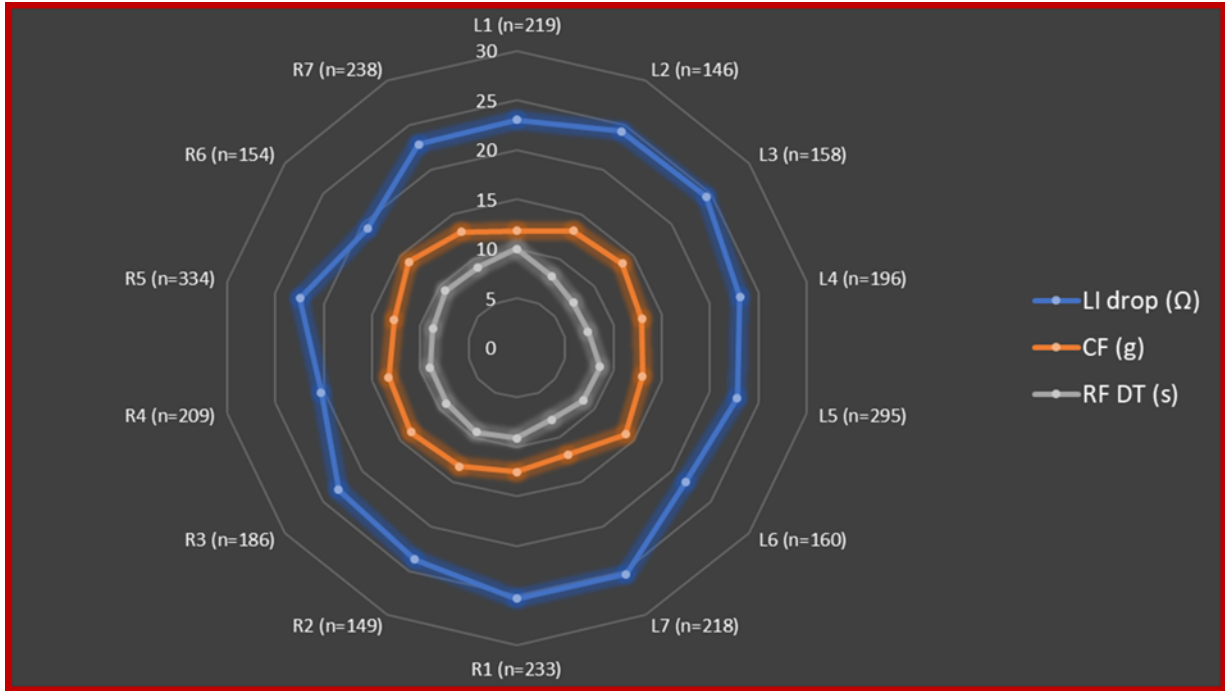
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484 **Figure 4.** Details of the distribution of baseline LI (*Panel A*), ablated tissue impedance (*Panel B*), RF application  
 485 time (*Panel C*) and CF values (*Panel D*) according to location sites: anterior sites vs posterior sites, LPV sites vs  
 486 RPV sites and inferior vs superior sites.



487 **Figure 5.** Details of the distribution of RF applications, CF, ablated tissue impedance values and RF delivery times  
 488 according to location sites. This Kiviart chart displays multivariate data with values represented on axes starting from  
 489 the same point. Each apex of the Kiviart charts represents a location site according with seven distinct sections of  
 490 right (R) and left (L) pairs of PVs (Anterior superior: R1, L1; Anterior inferior: R2, L2; Inferior: R3, L3; Posterior  
 491 inferior: R4, L4; Carina: R5, L5; Posterior superior: R6, L6; Superior: R7, L7). Blue, orange and grey dots represent  
 492 the mean LI drop values, CF values and RF delivery time values according to location sites.  
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511 **Supplementary material**

512 **Supplementary Table 1.** Details of the distribution of RF applications, CF, baseline and ablated tissue impedance

513 values according to location sites

Location Site			n (%)	Baseline LI, $\Omega$	LI drop, $\Omega$	CF, g	RF application time, sec
RPV	Anterior superior	R1	233 (8.0)	161.1±18	25.4±7	12.5±7	9.2±4
	Anterior inferior	R2	149 (5.1)	153.9±18	23.7±8	13.3±8	9.5±4
	Inferior	R3	186 (6.4)	159.0±21	23.0±7	13.7±9	9.1±4
	Posterior inferior	R4	209 (7.2)	157.0±16	20.3±7	13.3±8	9.0±4
	Carina	R5	334 (11.5)	156.1±18	22.4±8	12.7±7	8.7±4
	Posterior superior	R6	154 (5.3)	155.3±13	19.3±6	13.9±10	9.2±4
	Superior	R7	238 (8.2)	159.0±16	22.8±7	13.0±7	9.0±4
LPV	Anterior superior	L1	219 (7.6)	154.3±16	23.0±8	11.8±7	10.0±5
	Anterior inferior	L2	146 (5.0)	157.7±16	24.3±7	13.1±8	8.1±3
	Inferior	L3	158 (5.5)	157.9±15	24.5±8	13.6±10	7.3±4
	Posterior inferior	L4	196 (6.8)	158.3±15	23.1±6	13.0±7	7.4±4
	Carina	L5	295 (10.2)	156.8±17	22.8±8	13.0±8	8.6±4
	Posterior superior	L6	160 (5.5)	158.7±16	21.8±7	14.1±8	8.6±4
	Superior	L7	218 (7.5)	164.9±18	25.4±7	12.0±6	8.1±4

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515 CF=contact force; LI=local impedance; RF=radiofrequency; RPV=right pulmonary vein; LPV=left pulmonary vein.

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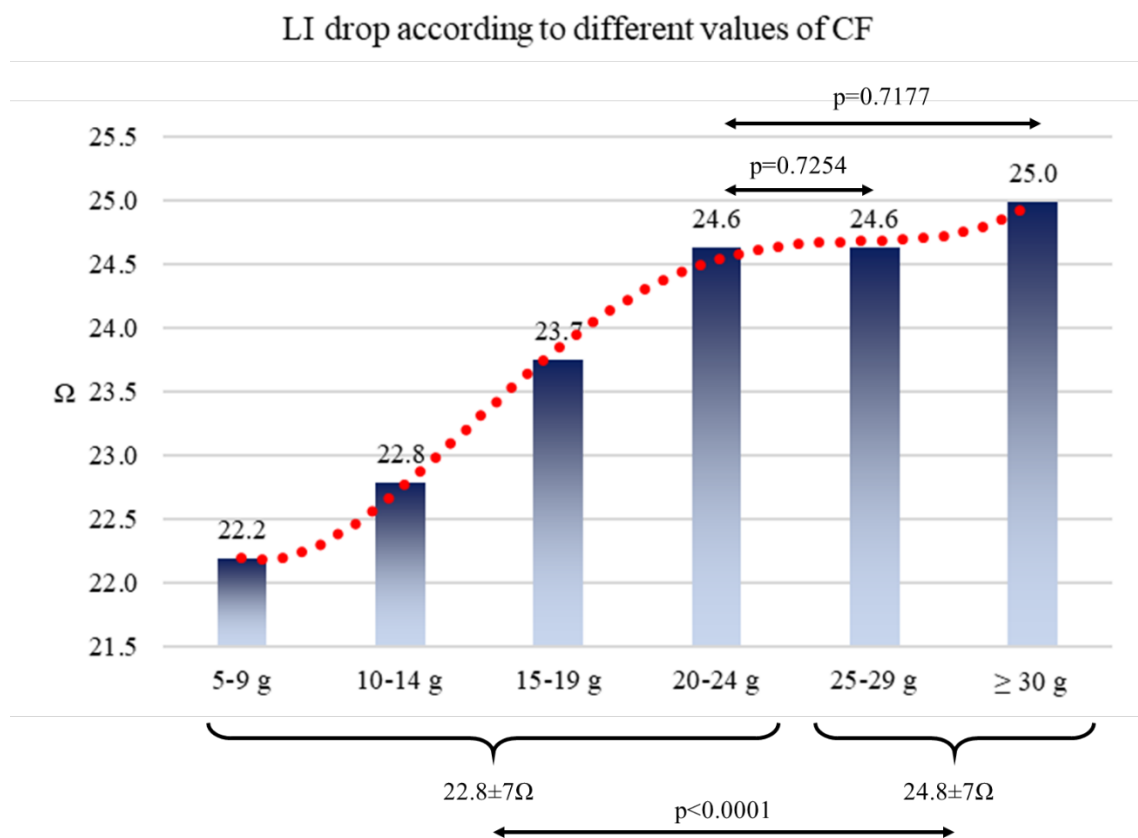
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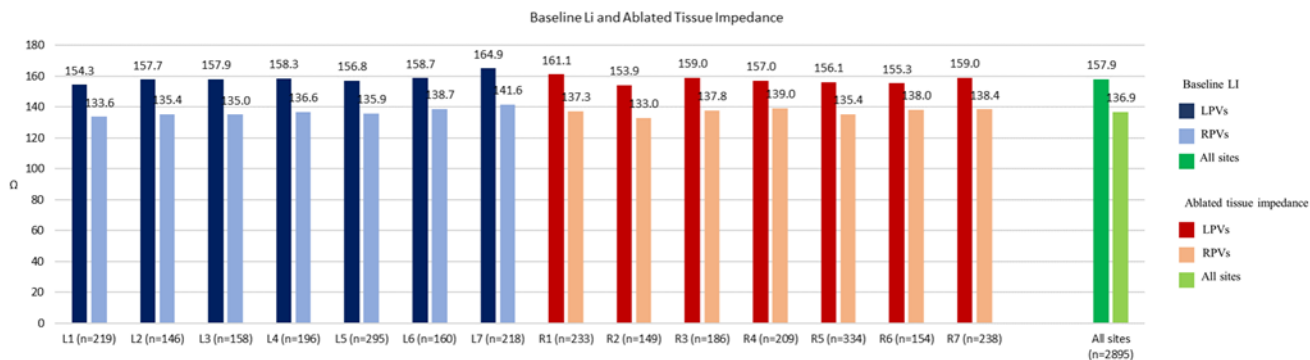
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522 **Supplementary figure 1.** Details of LI drop values according to different values of CF. Although there is a  
 523 significant difference dichotomizing CF values below and above 25 grams ( $22.8 \pm 7\Omega$  at  $< 25\text{g}$  CF intervals vs  
 524  $24.8 \pm 7\Omega$  at  $\geq 25\text{g}$  CF intervals,  $p < 0.0001$ ), LI drops markedly increased from 5 to 25g CF intervals, whereas it  
 525 showed a smooth transition above 25g. No differences were found comparing 20-24g CF interval to both 25-29g and  
 526  $\geq 25\text{g}$  CF intervals. Polynomial trendline is displayed in red.



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536 **Supplementary figure 2.** Details of the distribution of baseline and ablated tissue impedance values according to  
 537 location sites with seven distinct sections of right (R) and left (L) pairs of PVs. Anterior superior: R1, L1; Anterior  
 538 inferior: R2, L2; Inferior: R3, L3; Posterior inferior: R4, L4; Carina: R5, L5; Posterior superior: R6, L6; Superior:  
 539 R7, L7. The mean baseline LI and ablated tissue impedance values at sites of left pair of PVs are reported in red  
 540 tones, at sites of right pair of PVs are reported in blue tones. In green tones are reported the mean values at all sites.  
 541 Dark color denotes baseline LI, light color denotes ablated tissue impedance.



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