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Minimal configuration of body surface potential mapping for discrimination of left versus right dominant frequencies during atrial fibrillation

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Abstract

Background: Ablation of drivers maintaining atrial fibrillation (AF) has been demonstrated as an effective therapy. Drivers in the form of rapidly activated atrial regions can be noninvasively localized to either left or right atria (LA, RA) with body surface potential mapping (BSPM) systems. This study quantifies the accuracy of dominant frequency (DF) measurements from reduced-leads BSPM systems and assesses the minimal configuration required for ablation guidance.

Methods: Nine uniformly distributed lead sets of eight to 66 electrodes were evaluated. BSPM signals were registered simultaneously with intracardiac electrocardiograms (EGMs) in 16 AF patients. DF activity was analyzed on the surface potentials for the nine leads configurations, and the noninvasive measures were compared with the EGM recordings.

Results: Surface DF measurements presented similar values than panoramic invasive EGM recordings, showing the highest DF regions in corresponding locations. The noninvasive DFs measures had a high correlation with the invasive discrete recordings; they presented a deviation of <0.5 Hz for the highest DF and a correlation coefficient of >0.8 for leads configurations with 12 or more electrodes.

Conclusions: Reduced-leads BSPM systems enable noninvasive discrimination between LA versus RA DFs with similar results as higher-resolution 66-leads system. Our findings demonstrate the possible incorporation of simplified BSPM systems into clinical planning procedures for AF ablation.

KEYWORDS

atrial fibrillation, body surface potential mapping, dominant frequency, electrocardiography, lead distribution

1 | INTRODUCTION

Several studies have demonstrated the maintenance of atrial fibrillation (AF) by discrete atrial drivers whose ablation is associated with a favorable outcome, 1-3 making the identification and localization of these drivers the goal for AF mapping.^{2,4} Emerging technologies have used body surface potential mapping (BSPM) systems to noninvasively identify those atrial drivers, either by locating reentrant patterns^{3,5,6} or high dominant frequency (DF) sites.⁷ Although these preliminary results seem promising, the high burden and cost of deploying the BSPM systems impedes their wide spread utilization.

The aim of this study is to determine the minimal configuration of body surface leads necessary to noninvasively identify the atria presumably driving AF to facilitate the incorporation of the noninvasive BSPM approach into the clinical practice. Previous studies have already investigated how the selection of different leads' sets affects the electrocardiogram (ECG) waveform, 8-10 and have concluded that for AF signals, systems with more than 32 electrodes do not increase the signal reconstruction accuracy. 11 However, the ability of these lead sets to characterize the nature and position of possible AF drivers has not been studied. The present study demonstrates that a BSPM system with an approximate uniform distribution of 12 or more electrodes over the torso surface is capable

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of maximal DF identification and localization in the left versus the right atrium.

2 | METHODS

2.1 | Patients

We studied a total of 16 patients admitted for ablation of drug-refractory paroxysmal (N = 11) and persistent (N = 5) AF and various numbers of ablation procedures (0-2).⁶ All patients gave informed consent and the protocol was approved by the Institutional Ethics Committee.

2.2 | Electrophysiological study and electrogram (EGM) recordings

The electrophysiological study was performed under general anesthesia and periodic heparin bolus administration. Intracardiac EGMs were obtained in 14 patients from both atria during the procedure by using the following catheters introduced via the right femoral vein: (1) a standard tetrapolar catheter in the right atrial (RA) appendage; (2) a deflectable 4-mm mapping catheter (Marinr; Medtronic Inc., Minneapolis, MN, USA) in the distal coronary sinus; (3) a decapolar circular mapping Lasso catheter (Biosense Webster, Diamond Bar, CA, USA) used to map the pulmonary veins - left atrial (LA) junctions; and an ablation catheter (Navistar cooled-tip catheter, Biosense Webster). In those 14 patients a sequential mapping of both atria (106.4 \pm 65.8 points mapped, 25 minimum and 231 maximum points) was performed to determine the highest DF site and the preservation of the DF after a second acquisition was required. Then, the ablation catheter was positioned at the highest DF site.⁷ In two additional patients, body surface recordings were obtained simultaneously with a 64-pole basket catheter (Constellation, Boston Scientific, Natick, MA, USA) located sequentially on the right and left atria. The geometry from the left and right atria was reconstructed through the navigation software using the location of the catheters obtained by impedance measurements (CARTO XP, version 7.7; Biosense Webster). In patients arriving in sinus rhythm, AF was induced by burst pacing and AF episode durations longer than 5 minutes were required for inclusion in the study.

2.3 | Body surface potential recordings

Surface ECGs were simultaneously recorded with the EGM recordings by using a vest grid of 66 electrodes distributed as follows: 30 electrodes on the anterior, 34 on the posterior, and two on each lateral side of the torso (Figure 1A). The surface electrodes were located in vertical stripes with an interelectrode vertical distance of 3 cm. An additional left leg electrode was used in order to obtain a Wilson Central Terminal. The signals were recorded using a commercial system (Active Two, Biosemi, Amsterdam, The Netherlands) at a sampling frequency of 2048 Hz and were stored for offline analysis. Ventricular activation was removed by administration of a central venous bolus of adenosine (12–18 mg) and 4-second segments of surface ECGs surrounding the

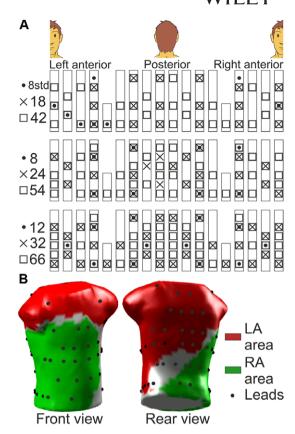


FIGURE 1 Lead and area distribution on the torso. (A) Distribution of the lead sets. (B) 3D torso with the 66 leads (black dots) and the LA and RA dominant influence areas (red and green, respectively). LA, left atrial; RA, right atrial [Color figure can be viewed at wileyonlinelibrary.com]

longest RR interval were used for the analysis. In cases with ventricular activation pauses shorter than 4 seconds, the QRSTs were cancelled. 12

2.4 Lead distribution on torso and 3D model

To analyze the effect of surface lead configuration on AF mapping of DF, different combination sets of the BSPM electrodes were defined. Those sets included uniform distributions of 8, 12, 18, 24, 32, 42, and 54 electrodes as well as a nonuniform distribution of eight leads matching the standard precordial and upper limb ECG electrodes distribution (see Figure 1A). The surface electrical activity was then projected on the surface of a typical adult human 3D torso model with 771 nodes and 1,538 faces (see Figure 1B), in which the surface potentials were reconstructed using the ECG signals provided by the different lead sets. The electrical potentials in areas between the nodes selected as electrodes for a particular lead configuration were interpolated as previously described.⁶

Two areas on the surface of the 3D torso model were assigned as being dominantly influenced by either LA or RA electrical activity (Figure 1B): an LA area (red) covering the left, upper back, and top side; and an RA area (green) covering the right, lower back, and front side.^{6,7} The number of electrodes in the surface LA area was 2, 3, 4, 5, 7, 8, 10, 14, and 19 electrodes for the eight standard, 8, 12, 18, 24, 32, 42, 54, and 66 electrodes, BSPM distributions, respectively, whereas the

surface RA area was covered by 3, 4, 7, 10, 14, 20, 25, 31, and 38 electrodes, respectively. The number of electrodes was then 97.7 \pm 40.3 % greater in the RA region than in the LA along the different BSPM sets, due to the larger RA region covered by the BSPM vest (38 electrodes) than the LA region (19 electrodes).

2.5 | Signal processing and DF

Baseline EGM and ECG signals were processed by decimation to 51.2 Hz and filtering with a Butterworth 10th-order low-pass filter with a cut-off frequency of 2 Hz. These signals were interpolated to 2,048 Hz and subtracted from the original signals to remove low-frequency drift. Resulted potential signals were then low-pass filtered with a 10th-order Butterworth filter with a cut-off frequency of 30 Hz. Power spectral density of 4-second segments of EGMs and ECGs was computed by using Welch periodogram (2-second long Hamming window with an 65,536 point Fast Fourier transform per window and 50% overlap) to determine the DFs.

3 | RESULTS

3.1 | Correspondence of panoramic invasive and full-surface DF measurements

Figure 2 shows an example of the surface DF distribution and its relationship with the panoramic intracardiac DF map of two representative cases of AF with left-to-right gradient (i.e., LA-fastest patient) and right-to-left gradient (i.e., RA-fastest patient). For better anatomical orientation, the DF values measured at the 64-poles catheter are displayed on the atrial shell by the DF measurement from the nearest catheter pole. The invasive panoramic DF map of patient #1 (Figure 2A) shows a highest DF (HDF) region located in the left pulmonary veins where the HDF was around 6 Hz, while the rest of the atrium maintained DF values lower than 5 Hz with a sample of 4.8-Hz measurement site illustrated at the RA appendage. The surface DF map obtained with the full leads set (66 electrodes, Figure 2B) is shown to display two well-demarcated domains: a large part of

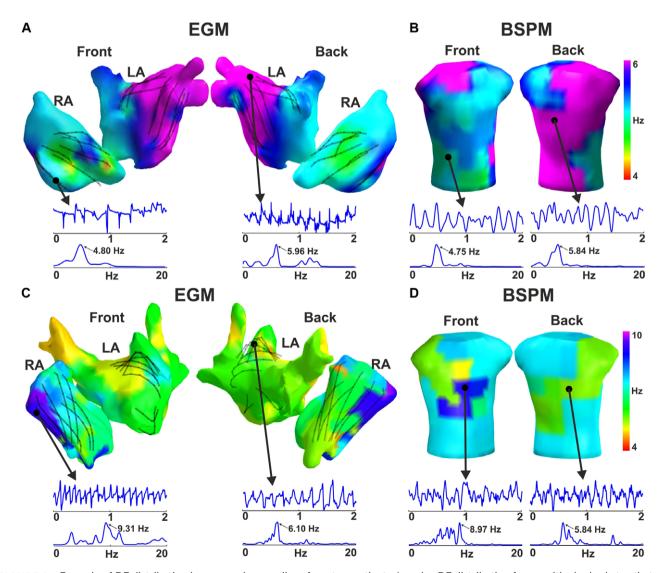


FIGURE 2 Example of DF distribution in panoramic recordings from two patients. Invasive DF distribution from multipolar basket catheters recordings (A and C) and simultaneous noninvasive DFs from BSPM (B and D) and the corresponding voltage signals and their PSD. BSPM, body surface potential mapping; DF, dominant frequency; PSD, power spectral density [Color figure can be viewed at wileyonlinelibrary.com]

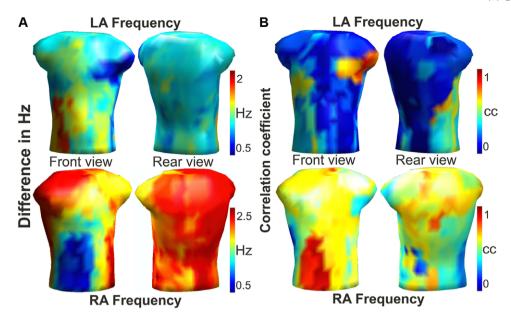


FIGURE 3 Relationship between invasive and noninvasive DF measurements. (A) Surface distribution of the average difference (in Hz) between the LA (top) and RA (bottom) DFs and the BSPM DF for the 14 patients mapped sequentially. (B) Surface distribution of the correlation between the LA (top) and RA (bottom) DFs and the BSPM DF for the same 14 patients. The areas with lowest errors in A (blue) are the areas with the highest cross-correlations in B (red). Abbreviations as in previous figures [Color figure can be viewed at wileyonlinelibrary.com]

the torso surface presents a DF value of 5.84 Hz corresponding to the HDF in the LA and roughly spread over the LA-influenced region shown in Figure 1B, whereas the rest of the torso depicts lower DF values around 4.75 Hz, similar to the DFs invasively measured in the RA. The panoramic invasive DF map from patient #2 (Figure 2C) shows a similar frequency distribution with the HDF region located in the RA appendage (9.31 Hz) while the rest of the atrium was activated between 5 and 7 Hz (6.10 Hz at the posterior LA wall). In this case, the surface DF map (Figure 2D) depicts a region in the right front side of the torso with a DF of 8.97 Hz, whereas the rest of the surface presented lower values, being 5.84 Hz for the DF in the back.

3.2 | Surface-DFs for the full BSPM configuration

The distribution and match between the intraatrial and surface DFs was studied for the 14 patients with discrete EGM recordings. The average frequency differences between the intracardiac maximal DF from the LA and RA and the body surface maximal DFs are depicted in Figure 3A. For each atrium, there is a torso area in which the difference between the intracavitary and noninvasive DF is minimal at <0.5 Hz (blue): the left surface side, including the left axillae, for the LA frequency; and the right front surface side of the torso for the RA frequency. The same surface regions can be identified in Figure 3B as having the highest correlation coefficient (red) between the intracavitary DF measurements from the LA and RA and the surface DF values from the BSPM. The correspondence between the maximal DFs in each atrium and the surface DFs establishes the body surface areas that are predominantly influenced by LA and RA activity and were used to define the LA and RA areas on the torso in Figure 1B.

3.3 | Surface distribution of DFs for reduced-leads BSPM

The errors in the DFs identification using the different lead sets of BSPM were quantified by the difference between their DFs and the intracavitary DFs. As shown in Figure 4, the HDF presented its maximal error for the standard eight-lead BSPM configuration (1.3 \pm 1.8 Hz), and this error decreased for arrangements with more leads with uniform distributions and stabilized at around 0.35 Hz for 12 or more electrodes. The coincidence between DFs obtained from intracavitary and surface recordings was also evaluated by calculating the correlation coefficients for the highest, LA, RA, and LA-RA (gradient) DFs measures for all the patient data set (Figure 5). It is shown that the

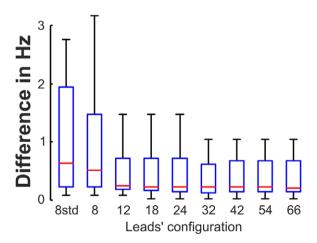


FIGURE 4 Difference (in Hz) between the highest dominant frequency obtained from invasive (electrogram) and noninvasive (body surface potential mapping) recordings as a function of the leads' configuration [Color figure can be viewed at wileyonlinelibrary.com]

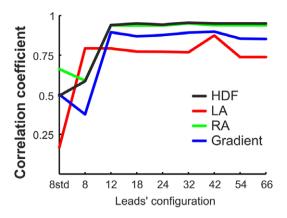


FIGURE 5 Correlation of the dominant frequencies values between the electrogram and body surface potential mapping signals as a function of the leads' configuration for the 14 patients. HDF, highest dominant frequency; LA, left atrium; RA, right atrium [Color figure can be viewed at wileyonlinelibrary.com]

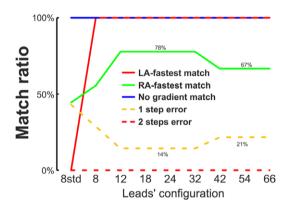


FIGURE 6 Patient classification with noninvasive dominant frequencies measurements. LA, left atrium; RA, right atrium [Color figure can be viewed at wileyonlinelibrary.com]

correlation between invasive and noninvasive DF measurements had low values for low number of surface leads. However, for 12 or more uniformly distributed leads the correlation coefficient reached higher and fixed values for all the lead sets: 0.75 for LA DF, 0.85 for the DF gradient, 0.94 for RA DF, and 0.95 for the HDF.

3.4 | Patient classification from surface DF measurements for reduced-leads BSPM

Noninvasive DF measures were used to classify the AF patients in three groups, according to its LA-to-RA DF gradient: (group I) LA-fastest patients, when the LA-to-RA DF was >0.75 Hz (N = 1); (group II) with no LA-to-RA DF gradient when -0.75 > LA-to-RA DF > 0.75 (N = 4); and (group III) RA-fastest, when the LA-to-RA DF was <-0.75 Hz (N = 9). The classification obtained with the EGM measures was used as gold standard and was compared with the classification carried out with the noninvasive measures (see Figure 6). The outcomes of the noninvasive classification are related as follows: LA-fastest match, when the EGM and noninvasive classification matched for LA-fastest patients (group I); no gradient match, when the EGM and noninvasive classification matched for no gradient patients (group II); RA-fastest match, when the EGM and noninvasive classification matched for

RA-fastest patients (group III); 1-step error, when a LA-fastest or RA-fastest patient was wrongly classified as no gradient patient and vice versa (group II to I/III or vice versa); and 2-step error when a LA-fastest patient was wrongly classified as RA-fastest patient or vice versa (group I to III or vice versa). Figure 6 shows that the DF's pattern is useful for the patient identification for lead configurations with 12 or more leads. For these lead configurations, the LA-fastest match ratio and no gradient match ratios were 100%, the RA-fastest match ratio was 78%, the 1 error ratio was 14%, and the 2 errors ratio was 0%. However, there is a small decrease in the outcome ratios for 42, 54, and 66 lead where the RA-fastest match ratio decreases from 77% to 66% and the 1 error ratio increases from 14% to 21%.

4 | DISCUSSION

4.1 | Major findings

Body surface DF maps obtained with uniformly distributed 66 leads present a good correlation with the invasively obtained DF measures in both atria. The present study demonstrates that fastest atrial localization based on surface HDF distribution assessment with 12 or more electrodes uniformly distributed on the body surface presents a high correlation with the higher-resolution BSPM system of 66 leads. Therefore, reduced-leads BSPM systems enable noninvasive discrimination between LA or RA atrial highest frequency sites with similar results as the higher-resolution 66-leads systems.

4.2 | Mechanisms of AF maintenance and their noninvasive characterization

The mechanisms underlying AF in patients are not always clear. Recent studies utilizing new mapping methods have supported the existence of a hierarchical AF organization of activity whereby identifiable sources of fast activity^{1,4} in the form of ectopic or rotor activation patterns maintain the arrhythmia, 2,3 but controversies still exist. 13,14 Arguably, better and easier identification of localized AF sources would render better targets for ablation, and thus efforts to improve technologies for the localization of AF drivers are ongoing.⁵⁻⁷ Although intracardiac recordings are considered the most reliable method for AF characterization,² they are not free of difficulties¹⁵ and the need for panoramic mapping led to the development of noninvasive AF mapping methods. Noninvasive mapping methods not only would allow for relatively simpler deployment than the usage of multiple intracardiac electrodes, but will also enable studying patients outside of the clinical Electrophysiology (EP) laboratory and preplanning individual ablation procedures. Some of the limitations of noninvasive technologies are their reduced sensitivity to the atrial activity during fibrillation. We have recently studied the effect of the body torso on body surface mapping and found that a significant amount of atrial information is retained on the body surface recordings.^{6,7,16} These studies showed that utilizing the appropriate frequency domain processing, noninvasive analysis of the AF activity on the torso surface enables atrial driver localization at a dominant atrium to facilitate ablation procedures. 1,4

4.3 Lead distribution in body surface electrocardiography

The number of body surface leads and their position has been broadly acknowledged to affect arrhythmia diagnostic accuracy, but in clinical practice the standard ECG leads configuration largely prevails. Indeed, the standard ECG lead set does not accurately respond to all diagnostic cardiology needs, and for this reason, BSPM systems with a broad distribution of leads and higher spatial resolution were developed. However, today there is no standard evaluation or recommendation for the surface electrode configuration for the different cardiac diseases and AF in particular.

The number of electrodes necessary for BSPM utilization has been mostly evaluated for ventricular activity based on the quality of the signals. 8-11,17 Only few studies investigated the role of the number and distribution of BSPM leads in studying atrial electrophysiology. Although the morphology and variability of the standard ECG signal in AF has been widely studied, only a few authors have considered that the standard ECG system is sensing mostly the ventricles and have proposed the use of a different arrangement of the nine standard leads in order to increase the effectiveness in AF characterization. 18 The broader electrode settings of BSPM have been used to study atrial arrhythmias, 5,19,20 but without an evaluation of accuracy for AF mechanisms characterization. In a recent study, Guillem et al. studied the number of leads necessary to properly reconstruct the torso potentials during in AF signals. 11 This study showed that during atrial fibrillatory rhythms a set of 34 electrodes was able to reconstruct the surface potential with the same accuracy than systems with more leads, since the reconstruction error reaches the noise level at the recording electrodes.

In this work, we propose the use of a reduced number of uniformly distributed leads for AF HDF source in interatrial localization, since the surface DF measurements did not increase their diagnostic information beyond a certain number of leads, in this case 12 leads. Although previous studies have concluded that at least 30 leads are necessary to properly reconstruct the surface electrical potential, 8,9,11 the present study makes use of the DF as a main signal characteristic which has been seen common in large surface regions although the individual ECG time series within each region present considerable variations. Thus, although 30 or more leads are necessary to properly reconstruct the ECG signal shape, the DF value is common to most of these ECG signals and then 12 leads would be enough to distinguish between the LA or RA dominance. This observation is also consistent with the natural smoothening filtering effect of the torso volume conductor that projects on the torso surface only the strongest characteristics of the atrial waves, with reduced sensitivity for the small electrical waves, no matter how high the number of leads is used for mapping.6

In this study, we have considered just uniform lead distributions, since in previous studies from our group we found that, in general, uniform lead distributions offered better AF signal reconstruction for the whole torso than nonuniform lead distributions. ¹¹ For limited subsets of leads, election of one left and one right surface electrode (selected as those with best correlations in Figure 3B), offers a similar accuracy

than the standard ECG with errors of 0.92 \pm 1.27 Hz and correlations of 0.18 for HDF, 0.9 for LA, 0.5 for RA, and 0.8 for gradient.

4.4 | Limitations

The study population includes a small number of 16 patients; however, we submit that the use of heterogeneous patients cohort, with paroxysmal and persistent AF as well as varying number of ablation procedures, 6,7,16 enhances the credence of our conclusions on AF interatrial characterization although it may not be representative of the general characteristics of a first detected paroxysmal AF, more prevalent in other AF ablation studies.

The number of leads in the LA surface region was significantly reduced compared to the number of leads in the RA surface region, due to the major covering of the full-resolution BSPM distribution of the RA surface region. This low coverage of the LA region could have provoked the underestimation of the surface LA DFs, as can be observed in the lower correlation value between invasive and noninvasive LA measurements (Figure 5) compared to the other correlation values. Further studious should solve this undercoverage by placing more ECG electrodes in the LA surface regions, concretely in the high anterior and posterior torso.

DF measurements recorded in the EGM catheters can vary inside each of the region considered, so there are atrial locations whose correlation with the surface DF may be lower or higher than reported. Moreover, it has been reported that the DF distribution across the atria can present temporal variations across time. ²¹ Nevertheless, previous studies have also reported the spectral stability of the highest DF region ²² and in our work the highest DF region was identified by performing a detailed DF map of both atria in all patients, where the HDF region was identified at least twice. Therefore, concordance of the highest DF with the simultaneous surface measurements should hold.

This study analyzes AF under the effect of the adenosine. Although previous studies have reported that adenosine infusions accelerate the activation rate during AF by up to 50% in some regions, they also reported that the general DF distribution across the atria remained stable over time. ^{4,23} Thus, although adenosine could have affected the local DF values in our study, their hierarchical distribution across the atria and on the body surface should have been preserved. Based on the error analysis in Figure 3, we expect to be able to detect DF gradients with a reduced body surface electrode configuration in cases of DF gradients larger than about 0.5 Hz.

4.5 | Conclusions

For coarse AF analysis, BSPM systems with about 12 uniformly distributed leads can achieve the same results as systems with up to 66 leads. Low-resolution BSPM systems retain the relevant clinical information that enables noninvasive discrimination between RA- or LA-fastest patients, a highly relevant information for increased ablation effectiveness. The results of this work can be the basis for the BSPM system simplification and wider clinical adoption for better

noninvasive AF preablation screening and planning as well as real-time EP mapping auxiliary guidance.

CONFLICT OF INTEREST

FA served on the advisory board of Medtronic and Sorin. OB received research support from Medtronic and St. Jude Medical. He is a cofounder and Scientific Officer of Rhythm Solutions, Inc., Research and Development Director for S.A.S. Volta Medical and consultant to Acutus Medical. The other authors have no conflict of interest. None of the companies disclosed here financed the research described in this manuscript.

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