Reduced P-wave Voltage in Lead I is Associated with Development of Atrial Fibrillation in Patients with Coronary Artery Disease

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Abstract

Background: Reduced P-wave voltage in lead 1 (PVL1) has been associated with atrial fibrillation (AF) recurrence. This study sought to determine the association between reduced PVL1 and AF in the NSTEMI population and the correlation between reduced PVL1 and interatrial block (IAB)/coronary artery disease (CAD).

Methods: Data were recorded for clinical, echocardiographic, angiographic, electrocardiographic and outcome variables. Patients were followed for a minimum of one year. Chi-square tests, independent samples t-tests and one-way ANOVA were used for the analysis, which was done using IBM SPSS.

Results: A total of 322 consecutive patients were included in the analysis. Patients with new-onset AF had a significantly lower PVL1 (0.085 ± 0.030 mV vs. 0.103 ± 0.037 mV; p = 0.007). There was a significant difference in mean PVL1 between those with no IAB, partial IAB and advanced IAB (p = <0.001). Those with any type of IAB had a significantly lower mean PVL1 than those without (0.094 ± 0.032 mV vs. 0.106 ± 0.038 mV; p = 0.005). Patients who developed AF had a significantly longer P-wave duration (126 ± 20 ms vs. 119 ± 17 ms; p = 0.022). Patients with IAB were more likely to develop new-onset AF (15.4% versus 7.5%, p = 0.025). There were significant co-linear associations between reduced PVL1 and IAB (p = 0.005); reduced PVL1 and diffuse CAD (p = 0.031) and IAB and diffuse CAD (p = 0.022).

Conclusions: Reduced PVL1 and IAB are associated with new-onset AF in patients with NSTEMI. Reduced PVL1 and IAB are correlated with each other indicating a possible common underlying mechanism. Both parameters are associated with CAD.

Introduction

Reduced P-wave amplitude in lead I (PVL1) has recently been shown to be associated with recurrence of atrial fibrillation (AF). In this study, conduction was shown to be displaced in the Bachmann region in patients with lower P-wave voltages using left atrial voltage and activation maps. A possible mechanism for the higher rates of AF recurrence in patients with reduced PVL1 was proposed to be abnormal interatrial conduction along the Bachmann region, the same mechanism as believed to underlie interatrial block (IAB). Interatrial block has previously been shown to be associated with atrial fibrillation in multiple cardiac populations. The P-wave represents atrial depolarization and as such is an indirect measure of atrial conduction. With normal anatomy, in sinus rhythm, the P-wave initiates at the sino-atrial node and travels inferiorly through the right atrium via the intra-atrial conduction pathways and most commonly crosses the interatrial septum superiorly via the Bachmann region, a broad muscular set of fibers. Partial interatrial block (IAB) results from a delay of conduction on this interatrial pathway at the Bachmann region. When this pathway is completely blocked, the right atrium is activated cranio-caudally; however, the left atrium is depolarized from the level of the coronary sinus to the posterior and superior region (retrograde activation) producing the classic biphasic P-wave of advanced IAB. IAB is clinically important due to its correlation with the development or recurrence of AF in various cardiac populations. While the exact pathology underlying the conduction abnormalities seen in IAB have not yet fully been determined it has been hypothesized that electrical remodeling and fibrotic atrial remodeling due to reduction of the blood supply to the Bachmann region may play a key role. In support of this, IAB has been shown to be associated with diffuse coronary artery disease (CAD). This study sought to determine the association of reduced PVL1 with development of AF in a population of patients with NSTEMI and its correlation with IAB and diffuse CAD.

Material and Methods

Patient Selection
Electronic records of a consecutive cohort of patients at Kingston...
variability.\textsuperscript{[1],[24]} P-wave onset was defined as the first upward or downward deflection from the isoelectric baseline and the P-wave offset as the return of the waveform to the baseline (Figure 1b). P-wave duration measurement has been previously described and validated.\textsuperscript{[25]} Partial IAB was defined as a P-wave ≥120ms while advanced IAB was defined as a P-wave ≥120ms with biphasic \textpm{} morphology in the inferior leads (II, III and aVF) according to the most recent consensus definition.\textsuperscript{[19]} Reduced PVL1 was defined as a P-wave voltage in lead I less than 0.10 mV. Echocardiographic and angiographic data were taken from clinical reports. Atrial fibrillation was evaluated through review of medical records, EGG's and holter monitors. AF ≥ 6 minutes' duration was considered as outcome.\textsuperscript{[26]}

**Statistical Methods**

Data were collected in Excel and imported into IBM SPSS (version 24.0 for Windows) for statistical analysis. Data were initially described using means and standard deviations for continuous data, and frequencies and percentages for categorical data. This was followed by a univariate analysis to assess the association of the collected data with the outcome, using one-way ANOVA and independent sample t-tests for the continuous data and chi-square tests (Pearson or Fisher’s Exact as appropriate) for the categorical data.

**Results**

**Population Demographics**

A total of 322 consecutive patients were included in the analysis. The population was 72.3\% male, the mean age was 65.4 ± 11.9 years, the mean ejection fraction was 55.2 ± 12.7\% and the mean left atrial diameter was 38.7 ± 6.0mm. Population characteristics are presented in [Table 1]. The prevalence of PVL1 less than 0.10 mV (reduced PVL1) was 50.3\%, PVL1 between 0.10 and 0.20 mV was 48.8\% and PVL1 > 0.20 mV was 0.9\%. The prevalence of partial IAB was 31.7\% and the prevalence of advanced IAB was 6.5\%. The incidence of new onset AF within one year was 10.6\%. The population was normally distributed in terms of P-wave voltage and duration.

**Associations with Atrial Fibrillation**

Participants who developed new-onset AF within one year had a significantly lower PVL1 (0.085 ± 0.030mV vs. 0.103 ± 0.037mV;
There was a significant difference of mean P-wave duration between PVL1 categories (<0.10mV, 0.10-0.20mV and >0.20mV) (p = 0.009) (Table 3). This difference favored increased P-wave duration with decreased PVL1 category. There was also a significant difference in the presence of advanced IAB between the PVL1 categories (p = 0.009) (Table 3). This difference favored decreased PVL1 with increased severity of IAB category. Patients who had advanced IAB had a significantly lower mean PVL1 than those without advanced IAB (0.074 ± 0.029 mV vs. 0.103 ± 0.362 mV; p<0.001). Patients who had any type of IAB had a significantly lower mean PVL1 than those without IAB (0.094 ± 0.032 mV vs. 0.106 ± 0.038 mV; p=0.005) (Table 2).

**Correlation of IAB Category with P-Wave Voltage**

There was a significant difference of mean P-wave duration between PVL1 categories (<0.10mV, 0.10-0.20mV and >0.20mV) (p = 0.009) (Table 3). This difference favored increased P-wave duration with decreased PVL1 category. There was also a significant difference in the presence of advanced IAB between the PVL1 categories (p = 0.014) and in the prevalence of any IAB (p = 0.035) (Table 4).

**Correlation of IAB Category with P-Wave Voltage**

There was a significant difference of mean P-wave duration between PVL1 categories (<0.10mV, 0.10-0.20mV and >0.20mV) (p = 0.009) (Table 3). This difference favored increased P-wave duration with decreased PVL1 category. There was also a significant difference in the presence of advanced IAB between the PVL1 categories (p = 0.014) and in the prevalence of any IAB (p = 0.035) (Table 4).

### Table 2: Difference in IAB Categories between P-Wave Voltage Categories

<table>
<thead>
<tr>
<th>IAB Category</th>
<th>P-Wave Voltage (mV)</th>
<th>P-Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Absent (n = 199)</td>
<td>0.106 ± 0.038</td>
<td>0.005</td>
</tr>
<tr>
<td>Present (n = 123)</td>
<td>0.094 ± 0.032</td>
<td></td>
</tr>
<tr>
<td>Advanced IAB (n = 301)</td>
<td>0.103 ± 0.362</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Partial IAB (n = 21)</td>
<td>0.074 ± 0.029</td>
<td></td>
</tr>
</tbody>
</table>

### Table 3: Difference in P-Wave Voltage and Duration by IAB and Voltages Categories

<table>
<thead>
<tr>
<th>IAB Category</th>
<th>Mean P-Wave Voltage (mV)</th>
<th>P-Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>No interatrial block</td>
<td>0.106 ± 0.038</td>
<td></td>
</tr>
<tr>
<td>Partial interatrial block</td>
<td>0.098 ± 0.031</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Advanced interatrial block</td>
<td>0.074 ± 0.029</td>
<td></td>
</tr>
</tbody>
</table>

### Table 4: Method for the Measurement of P-Wave Duration and Amplitude

<table>
<thead>
<tr>
<th>IAB Category</th>
<th>P-Wave Voltage</th>
<th>Absent (n = 199)</th>
<th>Present (n = 123)</th>
<th>P-Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>&lt;0.10 mV</td>
<td>90 (45.2%)</td>
<td>72 (58.5%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>0.10 - 0.20 mV</td>
<td>106 (53.3%)</td>
<td>51 (41.5%)</td>
<td>0.035</td>
<td></td>
</tr>
<tr>
<td>&gt;0.20 mV</td>
<td>3 (1.5%)</td>
<td>0 (0.0%)</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Advanced IAB</th>
<th>Absent (n = 301)</th>
<th>Present (n = 21)</th>
<th>P-Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>&lt;0.10 mV</td>
<td>145 (48.2%)</td>
<td>17 (81.0%)</td>
<td></td>
</tr>
<tr>
<td>0.10 - 0.20 mV</td>
<td>153 (50.8%)</td>
<td>4 (19.0%)</td>
<td>0.014</td>
</tr>
<tr>
<td>&gt;0.20 mV</td>
<td>3 (1.0%)</td>
<td>0 (0.0%)</td>
<td></td>
</tr>
</tbody>
</table>

#### Discussion

Reduced PVL1 was found to be significantly associated with the development of new-onset AF in this population. In addition, reduced PVL1 and IAB were found to be significantly correlated with each other. It is plausible that reduced PVL1 and IAB may be associated with the same pathological process leading to increased P-wave duration and reduced voltage, namely atrial fibrosis. Park et al. have recently demonstrated a significant correlation between reduced PVL1 and displaced conduction in the Bachmann region using left atrial voltage and activation maps.[1] Atrial fibrosis delays cardiac electrical conduction and reduces voltage, phenomena which have been well described previously.[27-31] Since the P-wave voltage depends on the direction of electrical propagation relative to the axis of the lead being measured and the myocardial mass and intervening substrates; it has been proposed that reduced P-wave voltage may be a result of an altered atrial conduction pattern and decreased myocardial mass due to atrial fibrotic scarring and increased degree of electro-anatomical remodeling.[1] It has recently been shown that diffuse CAD is associated with IAB and development of AF in the NSTEMI population.[11] In this current study, both reduced PVL1 and IAB are also significantly correlated with diffuse CAD. Therefore it is possible that the mechanism underlying both decreased PVL1 and IAB is fibrosis of the atria, particularly in the Bachmann region.[32,33].

#### Limitations

This study was retrospective in nature and as such may present inherent bias. AF was determined by clinical examination, ECG and Holter monitor reports; thus silent AF episodes may not have been recorded.

#### Conclusions

Reduced PVL1 is associated with new-onset AF in the NSTEMI population. In addition, PVL1 and IAB are significantly correlated with each other and with diffuse CAD. While the exact mechanism responsible for each have yet to be worked out, it is possible that the underlying cause could stem from fibrosis of the atria.

#### References

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