

## Original Article

# Importance of the relationship between sinus cycle length and junctional rhythm cycle length (occured during radiofrequency ablation) in predicting the successful modification of the slow pathway in Atrioventricular Nodal Re-entrant Tachycardias

Javier Jimenez-Candil, MD, PhD, Jose Luis Morinigo, MD, PhD, Claudio Ledesma, MD, Víctor Leon, MD, Candido Martín-Luengo MD, PhD.

Department of Cardiology, University Hospital, Salamanca, Spain

Address for correspondence: Javier Jimenez-Candil, MD, PhD, Department of Cardiology. University Hospital. Paseo de San Vicente, 58-182, 37007 Salamanca. Spain. E-mail: jimenezcandil/at/secardiologia.es

**Potential conflicts of interest:** None to disclose.

### Abstract

**Background:** In atrioventricular nodal re-entrant tachycardias (AVNRT), the achievement of Junctional Rhythms (JR) during Radiofrequency Ablation (RF) is a sensitive but non-specific marker of success. Our aim is to analyze prospectively the predictors of non-inducibility of AVNRT, focusing on the characteristics of the JR.

**Methods:** We included 75 patients with reproducibly inducible AVNRT. Ablation was performed following an electro-anatomical approach. After each application, the induction protocol was repeated.

**Results:** A total of 341 applications were performed. Although the achievement of  $\geq 1$  JR was necessary to obtain the non-inducibility, and the cumulative number of junctional beats (CJB) was higher in effective applications, no CJB cut-off was associated with a success rate higher than 75%. After the observation of a significant correlation between the sinus cycle length (CL) pre-RF and the CL of the JR (JR-CL) ( $c=0.52$ ;  $p<0.001$ ), the sinus CL pre-RF/JR-CL ratio (CL-ratio) adequately differentiated the successful vs. unsuccessful applications:  $1.41\pm 0.23$  vs.  $1.17\pm 0.2$  ( $p<0.001$ ). In a multivariate analysis, a CJB  $\geq 11$  ( $p<0.001$ ) and a CL-ratio  $\geq 1.25$  ( $p<0.001$ ) were found to be the only independent predictors of success. The combination of  $\geq 11$  of CJB with a CL ratio  $\geq 1.25$  achieved non-inducibility in 97% of our patients.

**Conclusions:** 1) The specificity of the occurrence of JR as a marker of the successful ablation of AVNRT is increased by the CL-ratio. 2) The achievement of  $\geq 11$  of CJB with a CL ratio  $\geq 1.25$  predicts non-inducibility in almost all patients.

**Key Words:** Atrioventricular nodal re-entry tachycardia; Ablation; Junctional ectopy

## Introduction

Atrioventricular nodal re-entrant tachycardia (AVNRT) is the most common cause of paroxysmal supraventricular tachycardias<sup>1</sup>. Selective radiofrequency catheter ablation (RF) of the slow AV nodal pathway has become the first-line curative treatment mode in patients with AVNRT<sup>2-5</sup> and relies on the non inducibility of the tachycardia as a criterion of success<sup>6</sup>. However, in 5 to 15 % of cases, AVNRT is not inducible or not reproducibly inducible during electrophysiological studies<sup>7,8</sup>. Although ablation of the slow AV nodal pathway is an accepted treatment for such patients<sup>4,8</sup>, the optimal end-point of the treatment has not been well established because evidence of a junctional rhythm (JR) during RF is a non-specific marker of success<sup>7,9</sup>.

Additionally, previous studies have not been able to pinpoint the clinical usefulness of the cycle length (CL) of JR in predicting the achievement of successful slow pathway ablation because the values observed during effective and ineffective energy applications overlap considerably<sup>9</sup>. We speculated that the CL of the JR (JR-CL) obtained during Radiofrequency Ablation would be related to the basal sinus CL; if this were the case, the ratio between sinus CL before application of RF and the mean CL of the JR (CL-ratio) could be a marker of successful applications. The purpose of the present study was hence to analyse prospectively the predictors of non-inducibility of AVNRT after applications of RF, focusing on the characteristics of the JR, the JR-CL, and the relationship of this latter with the sinus CL prior to RF ablation.

## Methods

### *Characteristics of the study population*

From July 2004 to March 2006, a total of 82 consecutive patients with clinically documented supraventricular tachycardia underwent an electrophysiological study and RF catheter ablation due to typical AVNRT at our institution. The study was approved by the Institutional Review Board of our institution. In 7 patients AVNRT was not reproducibly inducible; thus, our study population finally comprised 75 patients (mean age: 48 ± 9; females: 84%; hypertension: 24%). No patient had major structural heart disease.

### *Electrophysiological Study*

After obtaining written informed consent, baseline electrophysiological study was performed with patients in the fasting unsedated state. All antiarrhythmic drugs were discontinued at least five elimination half-lives prior to study. Three multipolar electrode catheters were introduced percutaneously via femoral veins and positioned under fluoroscopic guidance in the right atrium, the His bundle region, and the right ventricular apex. In some patients, a decapolar electrode catheter was introduced into the left antecubital vein and placed in the coronary sinus. Intracardiac electrograms were filtered at 30 to 500 Hz and simultaneously displayed with surface ECG leads I, II, V1 and V6 on a multichannel oscilloscope (Cardiolab 6.0, GE Medical Systems, Milwaukee, USA). A programmable stimulator (Model 5328, Medtronic, Inc) was used to deliver an electric stimulus with 2-ms duration at twice the diastolic threshold. Incremental pacing and programmed stimulation were performed in the right atrium and right ventricle to define anterograde and retrograde AV nodal conduction and to confirm that AVNRT was inducible. In 21 patients the tachycardia was not inducible in the baseline state; in these cases, isoproterenol (at graded doses from 1 to 4 µg/min IV) was infused to

facilitate induction. AVRNT was diagnosed using previously described criteria; intra-atrial re-entrant tachycardia and tachycardia incorporating a midseptal or paraseptal accessory pathway were excluded<sup>10-13</sup>.

Junctional beats were identified based on the QRS configuration and duration identical to that of sinus beats and the absence of AV conduction from a preceding P wave. All episodes of JR during the applications of RF were analysed as regards the ablation site, the total number of junctional ectopic beats, the duration of JR, and the CL of the JR.

### ***Mapping and Ablation***

Mapping and ablation were performed using a multipolar catheter (Marinr MC, Medtronic Inc., USA) with a distal electrode size of 4mm, following an anatomically guided approach<sup>3</sup>. Briefly, the triangle of Koch, extending from the coronary sinus ostium up to the His bundle region, is divided into three regions designated posterior, mid and anterior. The ablation catheter was placed along the tricuspid septal annulus down to the posterior aspect of the interatrial septum adjacent to the coronary sinus ostium (posterior zone), obtaining an AV electrogram ratio of 0.1 to 0.3. At the end of each RF delivery, the inducibility of AVNRT was tested and, if still inducible, a second RF pulse is delivered to an adjacent site with a higher AV ratio. In the case of further unsuccessful pulses, the catheter is moved towards a more mid and anterior position if necessary.

The RF energy was delivered while temperature was monitored at the catheter tip, which was limited to a maximum of 55 °C, with a preset maximal power of < 45 W. The duration of the RF pulses was 30 seconds; in the presence of an accelerated JR (CL<450 ms), a JR without ventriculo-atrial conduction, a PR prolongation, a rise in impedance or a displacement of the catheter, the application was discontinued immediately. After each application, regardless of whether or not JR was observed, the induction protocol was repeated, including the infusion of isoproterenol in the cases in which it had been necessary previously.

### ***Location of the ablation site***

Each ablation site was determined using three different fluoroscopic projections: right anterior oblique 20°, front view, and left anterior oblique 60°. Three areas of equal height between two principal reference points (inferior edge of the coronary sinus ostium and the detection site of distal His-bundle deflection) were determined: posterior, mid-septal and anterior.

### ***Follow-up***

After the procedure, the patients were monitored for 24 hours prior to discharge. No antiarrhythmic drugs were prescribed. Follow-up information was obtained at the time of the clinical revisits, which took place 6 and 12 months after discharge. Whenever the patient had symptoms suggestive of tachycardia, they were advised to seek consultation from our arrhythmia clinic or contact their physicians to verify possible AVNRT recurrence.

### ***Definitions***

- Reproducibly inducible tachycardia: Tachycardia was induced three or more consecutive times with the same protocol.

- Non-inducible tachycardia: Post-ablation, AVNRT was no longer inducible with no more than a single atrial echo.
- Successful application: An application that achieved non-inducibility.
- Sinus Cycle Length: Mean cycle length of the 10 consecutive beats registered immediately before or after each RF application.
- Junctional Rhythm: The occurrence of  $\geq 2$  consecutive junctional beats.
- Junctional Rhythm Cycle Length: Average of the cycle length of all junctional rhythms or beats occurred during an application. In the case of isolated junctional beats, it was determined the coupling interval with the precedent sinus beat.
- Cumulative Junctional Beats: Number of junctional beats achieved from the first application to the present application.
- CL-ratio: Ratio of Sinus Cycle Length before RF/Junctional Rhythm Cycle Length.

### **Statistical**

### **analysis**

The statistical analysis was performed using the SPSS 11.5 for Windows (SPSS Inc., Chicago, Illinois). Normal and continuous variables were described using the mean and standard deviation, whereas categorical variables were summarized by the number of patients and percentage. In order to establish cut-off points of continuous variables with the best sensitivity and specificity, we determined the Receiver-Operating Characteristic Curves. Comparison of categorical variables was performed with the Chi-square test (or Fisher's exact test if  $n < 5$ ). Comparison of two normal (determined by the Kolgomorov-Smirnov test) and continuous variables was accomplished with Student's t test. Comparison of  $> 2$  continuous variables was performed using the ANOVA-test. Multivariate analysis was performed using the stepwise logistic regression test, including the variables with statistical significance in the univariate analysis. A P value  $< 0.05$  was considered to represent a significant difference.

### **Results**

A total of 341 RF applications were performed (mean applications per patient:  $4.6 \pm 2.4$ ; median: 4; range: 1-13). In 190 applications (56 %) we achieved JR. The median of beats/application was 8. Non-inducibility was achieved in all cases, and no major complication occurred, in particular neither transient nor persistent second or third degree AV block. In all successful applications at least one JR was achieved. **Table 1.** No recurrences of AVNRT were recorded during a follow-up period of 1 year.

### **Determinants of the JR-CL**

The mean JR-CL was  $574 \pm 114$  ms (median: 570 ms; interquartile range: 150 ms). The mean of the sinus CL before RF was  $704 \pm 104$  ms (median: 700). There was a positive and significant correlation between the sinus CL before RF and the mean CL of the subsequent JR achieved during the application (Pearson coefficient: 0.56;  $p < 0.001$ ). **Figure 1.** Applications preceded by a sinus CL of  $\geq 700$  ms were associated with a significantly higher mean JR-CL:  $624 \pm 111$  versus  $512 \pm 84$  (95 % CI of the difference: 83; 141;  $p < 0.001$ ).

The JR-CL was shorter in applications with a duration of the atrial electrogram  $\geq 50$  ms and in those performed after isoproterenol infusion. Other variables analysed were not related to any significant difference in the JR-CL. **Table 2.** In a multivariate analysis for accelerated JR (JRCL  $\leq 550$  ms), a sinus CL of  $< 700$  ms (OR: 4.3; 95% CI: 2.3-8.2;  $p < 0.001$ ) was the only

independent predictor. The use of isoproterenol (OR: 1.47; 95% CI: 0.67-3.2; p=0.3) or a duration of the atrial electrogram of  $\geq 50$  ms (OR: 1.9; 95% CI: 0.98-3.7; p=0.07) ceased to be statistically significant.

**Table 1.** Characteristics of the applications of RF.

Variable	Overall	Successful applications	Unsuccessful applications	P value
Temperature, °C	55±2	55±3	55±1	0.4
Duration, s	27±6	26±5	27±6	0.7
Sinus rhythm CL before RF, ms	704±104	708±104	705±105	0.9
Sinus rhythm CL after RF, ms	700±105	702±106	698±107	0.8
Drugs used for induction	28%	28%	27%	0.9
Site of ablation mid-septal	54%	69%	54%	0.004
Stability	81%	86%	80%	0.2
Duration of atrial electrogram, ms	48±13	59±11	45±12	<0.001
A/V voltage ratio	0.2±0.9	0.23±0.06	0.2±0.09	0.02
Voltage of atrial electrogram, mV	2.1±0.8	2.2±0.6	2±0.9	0.6
Complex atrial electrogram	24%	23%	22%	0.4
Junctional Rhythm	55%	100%	42%	<0.001
Nº of Junctional Beats	5±3	12±5	3±5	<0.001
Cumulative Junctional Beats	10±7	23±14	6±10	<0.001
Duration of JR, seconds	2.9±3.7	6.2±3.3	2.2±3.6	0.007
Time to first JR, seconds	11±7	9±8	11±6	0.3
JR-CL	573±120	511±84	615±113	<0.001
CL-ratio	1.28±0.26	1.41±0.23	1.17±0.2	<0.001
JR without VA conduction	23 %	48%	11%	<0.001

RF: Radiofrequency Ablation. JR: Junctional Rhythm. JR-CL: Junctional Rhythm Cycle Length.

### *Predictors of non-inducibility*

Successful applications showed a more prolonged duration of the atrial electrogram, a higher A/V voltage ratio, more junctional beats, more CJB, a longer duration of the JR, a shorter JR-CL, a higher CL-ratio, and a higher frequency of the absence of VA conduction during the JR, and of mid-septal ablation sites. **Table 1.**



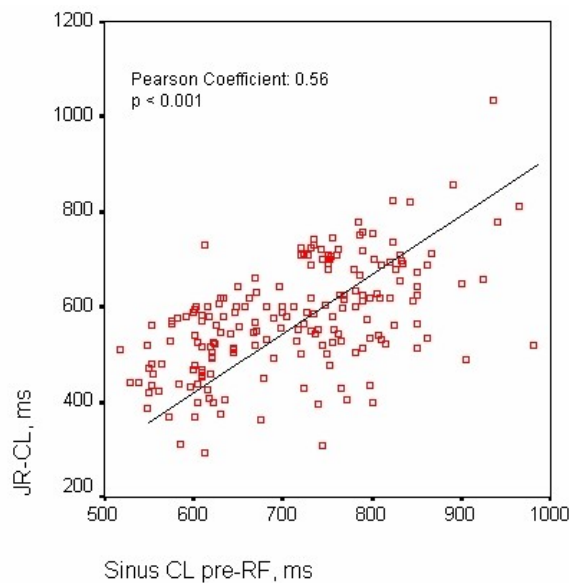


Figure 1: Diagram showing the correlation between the sinus CL and the JRCL.

Table 2. Comparison of the Junctional Rhythm Cycle Length in relation to several variables. Univariate analysis.

Variable	JR-CL*, ms	Statistical analysis
Age>55	572±112 versus 562±109	95% CI of the difference: -12 to 34; p=0.2
A/V ratio ≥0.2	565±113 versus 573±111	95% CI of the difference: -34 to 23; p=0.1
Duration of the atrial electrogram ≥50 ms	559± 103 versus 599±128	95% CI of the difference: -73 to -6; p=0.021
Complex atrial electrogram at the ablation sites	564±138 versus 576±112	95% CI of the difference: -27 to 51; p=0.5
Use of isoprenaline	517±104 versus 590±112	95% CI of the difference: -34 to -111; p=0.004
Sinus Cycle Length ≥ 700 ms	625±111 versus 512±84	95% CI of the difference: 83 to 141; p<0.001
Site of application mid-septal	582±135 versus 586±111	95% CI of the difference: -16 to 58; p=0.2

Two continuous variables showed an adequate correlation with non-inducibility (defined as an area under the ROC curve ≥ 0.8): CJB and the CL ratio. Table 3. The best cut-off point of the continuous variables with statistical significance were a CJB of ≥11, a CL ratio of ≥1.25, a duration of JR of ≥3.5 s, a JR-CL of ≤550 ms, a duration of atrial electrogram of ≤50 ms, and an A/V voltage ratio of ≥0.2. Their predictive values are shown in Table 3.

**Table 3.** Statistical values for the probability of success of several variables.

Variable	Area under ROC curve	Best cut-off point	S	E	PPV	NPV
CL-ratio	0.86	1.25	95 %	85 %	77 %	94 %
JR-CL, ms	0.77	550	73 %	72 %	65 %	80 %
CJB	0.80	11	95 %	62 %	58 %	99 %
Duration of atrial electrogram, ms	0.73	50	89 %	65 %	45 %	95 %
A/V voltage ratio	0.59	0.2	80 %	60 %	26 %	87 %
Duration of JR, s	0.68	3.5	82 %	77 %	54 %	94 %
Mid-septal ablation site	-	-	70 %	47 %	27 %	85 %

ROC curve: Receiver-Operating Characteristic Curve. S: sensitivity. E: Specificity. PPV: Positive predictive value. NPV: Negative Predictive Value. JR-CL: Junctional Rhythm Cycle Length. CJB: Cumulative number of Junctional beats. JR: Junctional Rhythm.

All of these variables, together with a mid-septal site of ablation and the absence of VA conduction during the JR, were associated with a significantly higher frequency of success in the univariate analysis. However, in the multivariate analysis, only two persisted as independent markers of non-inducibility: a CJB of  $\geq 11$  and a CL ratio of  $\geq 1.25$ . **Table 4.**

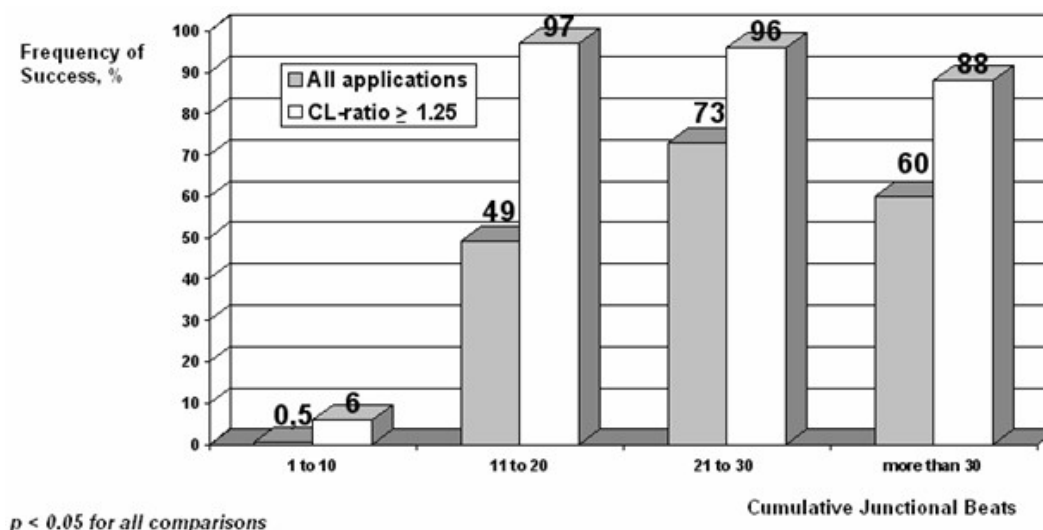
**Table 4.** Statistical analysis of predictors of success.

Variable	Frequency of success*	Univariate analysis	Multivariate analysis
CJB $\geq 11$	57% versus 0.5%	OR: 173 (95% CI: 27-215); p<0.001	OR: 81 (95% CI : 15-413) ; p<0.001
Duration of the atrial electrogram $\geq 50$ ms	45% versus 4.4 %	OR: 17 (95% CI: 8-38); p<0.001	OR: 4 (95% CI : 0.85-17); p=0.086
CL-ratio $\geq 1.25$	78% versus 5%	OR: 24 (95% CI: 10-57); p<0.001	OR : 8 (95% CI: 15-41); p<0.001
JR-CL $\leq 550$ ms	20% versus 65%	OR: 0.44 (95% CI : 0.32-0.6) ; p=0.001	OR : 0.6 (95% CI : 0,1-3.4) ; p=0.6
A/V voltage ratio $\geq 0.2$	26% versus 13%	OR: 1.2 (95% CI : 1.1-1.3); p=0.005	OR : 1.4 (95% CI : 0.3-6.7; p=0.6
Mid-septal ablation site	27% versus 15%	OR: 2.1 (95% CI: 1.2-3.7); p=0.008	OR : 0.7 (95% CI : 0,1-3.7) ; p=0.7
Duration of JR $\geq 3.5$ s	54% versus 6%	OR: 15 (95% CI: 7-29) ; p<0.001	OR: 0.8 (95% CI: 0.1-3.5); p=0.8
JR without VA conduction	68% versus 23%	OR: 7.1 (95% CI: 3.5-14); p<0.001	OR: 3.3 (95% CI: 0.6-13); p=0.3

\* Present versus absent. OD: Odds Ratio. CI: Confidence Interval. CJB: Cumulative number of Junctional beats. JR-CL: Junctional Rhythm Cycle Length. JR: Junctional Rhythm.

Although the achievement of at least of one JR was necessary to demonstrate non-inducibility, no CJB was associated with a success frequency greater than 75 %. Nevertheless, a CL ratio of  $\geq 1.25$  significantly increased the positive predictive value for non-inducibility across the whole range of CJB values. **Figure 2.**

A total of 74 applications (performed in 70 patients) displayed a CJB of  $\geq 11$  and a CL ratio of  $\geq 1.25$ ; of these, 70 (95 %) were successful. The success frequency was significantly higher in attempts with a CJB of  $\geq 11$  and a CL ratio of  $\geq 1.25$ : 95% vs. 2% (OR: 13; 95% CI: 6-43;  $p < 0.001$ ). Applications with a CJB of  $\geq 11$  and a CL ratio of  $\geq 1.25$  had a sensitivity, specificity, negative predictive value and positive predictive value for non-inducibility of 93 %, 98 %, 98 % and 95 %, respectively. The four unsuccessful applications with a CJB of  $\geq 11$  and a CL ratio of  $\geq 1.25$  occurred in 2 patients (two in each). Thus, in 68/70 patients (97 %) the achievement of a CJB of  $\geq 11$  and a CL ratio of  $\geq 1.25$  was predictor of success. Focusing on the remaining five patients, four became non-inducible with a CJB of  $\geq 11$  and a CL-ratio of  $< 1.25$ , and in one the number of CJB was 6 and the CL-ratio was 1.54.



**Figure 2.** Bar diagram showing the frequency of success of the applications in relation to the number of CJB and the CL-ratio.

In 33 patients (44 %) the successful application abolished the dual AV nodal pathway physiology. CL-ratio correlated with the probability of the dual AV nodal pathway physiology elimination (C-coefficient: 0.75;  $p < 0.001$ ). Clasifying the last applications into three groups according to the tertiles of CL-ratio ( $\square < 1.28$ ;  $\square 1.29-1.42$ ;  $\square > 1.43$ ), we found that the applications with higher CL-ratio were associated with a more elevated frequency of the abolition of the slow pathway function: 20% vs. 44% vs. 68% ( $p < 0.001$  for the trend). Other variables analysed did not show significant differences. **Table 5.**

## Discussion

The major finding of our study was that despite a positive and significant correlation between the sinus CL before RF and the mean of the JR-CL, the CL ratio increased the specificity and the positive predictive value of the amount of JR in predicting the acute success of slow pathway RF ablation.



**Table 5.** Relationship between the abolition of the dual AV nodal pathway physiology and different variables.

Variable	Values*	Statistical analysis
CJB	25±18 vs. 22±9	95% Confidence Interval of the Difference: -3; 9; p=0.3
CL-ratio	1.49±0.24 vs. 1.34±0.18	95% Confidence Interval of the Difference: 0.04; 0.25; p=0.005
JR-CL, ms	496±90 vs. 522±77	95% Confidence Interval of the Difference: -66; 12; p=0.2
Mid-septal ablation site, %	68 vs. 71	95% Confidence Interval: 0.4-3.3; p=0.6
Atrial electrogram duration, ms	58±10 vs. 59±10	95% Confidence Interval of the Difference: -5; 4; p=0.8
Complex atrial electrograms; %	27 vs. 26	95% Confidence Interval: 0.3-2.6; p=0.9

\* Present versus absent. OD: Odds Ratio. CI: Confidence Interval. CJB: Cumulative number of Junctional beats. JR-CL: Junctional Rhythm Cycle Length. JR: Junctional Rhythm.

### ***Junctional rhythm during slow pathway RF ablation: mechanism and implications***

The precise mechanism responsible for the JR is not well established. Different hypotheses have been proposed, such as: a) the postganglionic release of noradrenaline from the sympathetic nerve endings, produced by the RF current, which may increase junctional automaticity<sup>14</sup>; b) an enhanced automaticity of heat-sensitive cells located close to the AV node in response to thermal effects of RF<sup>15</sup>; and, c) the thermal current conducted through specialized atrionodal fibers to the AV node<sup>16</sup>.

Although in some rare cases successful slow-pathway ablation is possible in the absence of junctional ectopy<sup>17</sup>, the achievement of a JR is a sensitive marker of success<sup>9,18</sup>. In the present study, the occurrence of a JR was universal in the effective applications; this also has been reported by others<sup>9</sup>. However, the specificity of this finding is limited; it appears in 25-65 % of unsuccessful attempts (9, 19). On the other hand, JR with a CL of < 350 ms and absence of ventriculoatrial conduction during JR have been identified as markers of AV block<sup>20,21</sup>.

### ***Predictors of Junctional Rhythm Cycle Length***

In previous reports, the relationship between the JR-CL and several variables related to RF applications has been analysed: neither mid-septal ablation sites<sup>19</sup> nor the duration of the local atrial electrogram<sup>22</sup> modified the JR-CL. We found that the JR-CL was related to the sinus cycle length, with a positive and significant correlation between both. Although in our study a duration of the atrial electrogram of ≥50 ms and the use of isoproterenol were also associated with shorter JR-CL, in the multivariate analysis the only independent predictor of JR-CL ≤550 ms was the sinus CL pre-RF. It is clear that the emergence of the JR always occurs with a CL shorter than the sinus rhythm, because JR with a CL longer than that of sinus rhythm would be suppressed by sinus node activity. Moreover, several works have shown that high levels of catecholamines enhance the automaticity of the nodal tissue<sup>23</sup> and facilitate the emergence of the

JR during RF<sup>24</sup>; it is possible that the sinus CL, as an indicator of the level of activation of the sympathetic nervous system<sup>25</sup>, might be associated with the JR-CL during RF applications through this mechanism.

### ***Specificity of the Junctional Rhythm in predicting successful ablation***

Because the sensitivity of JR for successful slow pathway ablation is very high, it is useful to use the occurrence of the JR as marker of effective RF applications. In general, the bursts of JR are significantly longer at successful sites<sup>9,19,26</sup>, but the number of junctional beats objectified during effective and ineffective applications is considerably overlapped. Thus, the quantity of junctional ectopy during attempted RF treatment is not useful in predicting whether slow pathway ablation has been achieved<sup>9</sup>. Other authors have proposed that the total amount of JR is related to the total abolition of slow pathway conduction and may serve as a marker of success; unfortunately the specificity of this variable is very low<sup>26</sup>. In our study, the cumulative number of junctional beats (CJB) correlated with the non-inducibility; even though the correlation was adequate, no CJB presented a sufficiently high positive predictive value to be used as a predictor of success.

Other variables have been found to be independent predictors of successful RF attempts; thus the following may increase the specificity of the JR as marker of success: the ablation site (mid-septal rather than posterior)<sup>19</sup>, the duration of the atrial electrogram (including slow pathway potentials when present)<sup>19</sup> and a higher ablation temperature<sup>27</sup>.

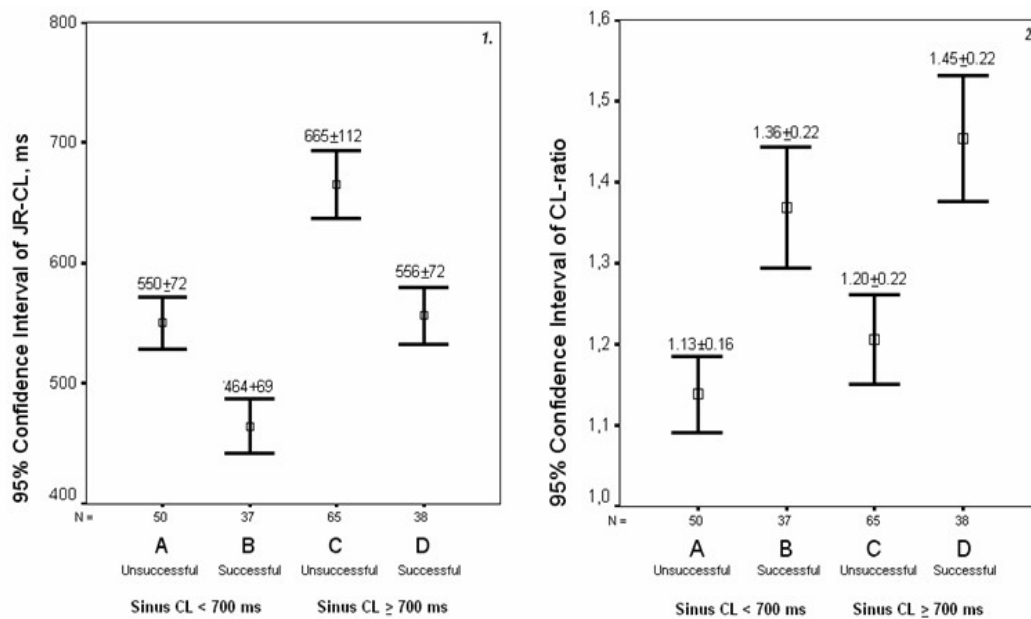
During the application of RF, the JR is thought to be a marker of thermal injury, a shorter JR-CL has been associated with a high degree of lesion of the AV node<sup>21,28</sup>. This may have two practical consequences. Firstly, it has been demonstrated that a JR with a CL under 350 ms is a predictor of conduction block<sup>21</sup>. Secondly, the JR-CL could increase the specificity of the occurrence of JR for predicting the successful modification of the slow pathway with RF, as exemplified by either tachycardia non-inducibility or abolition of the dual AV nodal pathway physiology. This point has not yet been clarified. In several studies the effective applications did not differ in JR-CL compared with the junctional ectopy that occurred during failed attempts<sup>9,19</sup>; in other, the JR-CL was significantly shorter in successful applications (the authors only reported a univariate analysis), although with limited clinical value<sup>29</sup>. In the present study the JR-CL of the successful attempts was also shorter but, after adjusting for other significant variables, particularly the CL-ratio, the JR-CL was not an independent marker of success. The explanation of this finding could lie in the significant correlation found between the sinus CL and the JR-CL, which produces produced an important overlap between the JR-CL of effective applications preceded by a larger sinus CL and ineffective applications preceded by a shorter sinus CL. **Figure 3.**

In our series, the CL-ratio had the best correlation with the probability of success, avoiding the overlap of JR-CL between effective and ineffective applications. Its best cut-off point (a CL ratio of  $\geq 1.25$ ) increases significantly the specificity and positive predictive value for non-inducibility of the different ranges of CBJ. The combination of a CJB of  $\geq 11$  and a CL-ratio of  $\geq 1.25$  maintained a higher sensitivity, had a positive predictive value for success of 95 %, and achieved non-inducibility in 97% of our patients.

Finally, up to 40 % of patients have residual slow pathway function after successful ablation of AVNRT<sup>2,30</sup>; our data show that, among successful applications, the higher the CL-ratio the more probable the elimination of the dual AV nodal pathway physiology.

*Clinical implications*

The major clinical implication of the present study is that, in an electro-anatomical approach, the combination of a CJB of  $\geq 11$  and a CL ratio of  $\geq 1.25$  is useful to assess when the successful slow pathway modification has been achieved. This could allow to test the inducibility more efficiently, avoiding unnecessary applications and shortening the duration of the procedures. In addition, since in 5-15 % of patients with documented paroxysmal supraventricular tachycardia the tachycardia is non-inducible (or non-reproducibly inducible)<sup>7,8</sup>, and since for these patients the slow pathway ablation is accepted as treatment<sup>4,8</sup>, the achievement of a CJB of  $\geq 11$  and a CL-ratio of  $\geq 1.25$  could be used as the end-point of the RF.



**Figure 3.** Scatter plot showing the JR-CL and the CL ratio of the applications when they were classified by the Sinus Cycle Length pre-RF and the result of the application. **Panel 1:** A vs. D: 95 % Confidence Interval of the Difference: -34; 42; p=0.8. B vs. C: 95 % Confidence Interval of the Difference: 85; 135; p<0.001. **Panel 2:** A vs. D: 95 % Confidence Interval of the Difference: 1.08; 1.34; p<0.001. B vs. C: 95 % Confidence Interval of the Difference: 0.12; 0.48; p=0.008.

**Study Limitations**

- The lower number of patients could decrease the statistical power of our findings.
- Since the location of the ablation sites was determined exclusively with fluoroscopic and electrical references, its accuracy might not be optimal.
- The number of cumulative junctional beats (CJB) included all junctional beats occurred from the first to the present application. This could decrease the value of the parameters assessed in the present application, since there is an influence from previous applications. This is a difficulty common to all studies including several RF applications in a single patient.
- The analysis was performed taking in consideration isolated junctional beats and junctional rhythms together. This did not allow to determine differences between both.

## **Conclusions**

The specificity of the occurrence and amount of the JR as marker of successful slow pathway ablation is insufficient. Effective applications achieved JR with significantly shorter JR-CL, but since the JR-CL correlates with the sinus CL, the JR-CL values in effective and non-effective applications overlap, depending on the sinus CL. The CL-ratio appears as an independent predictor of successful attempts, increasing the specificity of the JR and becoming an useful tool for identifying the effective applications. New studies involving larger numbers of patients are needed to confirm the value of our findings.

## **References**

1. Akhtar M, Jazayeri MR, Sra J, Blanck Z, Deshpande S, Dhala A. Atrioventricular nodal reentry. Clinical, electrophysiological, and therapeutic considerations. *Circulation* 1993;88:282-95.
2. Jackman WM, Beckman KJ, McClelland JH, Wang X, Friday KJ, Roman CA, et al. Treatment of supraventricular tachycardia due to atrioventricular nodal reentry, by radiofrequency catheter ablation of slow-pathway conduction. *N Engl J Med* 1992;327:313-8.
3. Jazayeri MR, Hempe SL, Sra JS, Dhala AA, Blanck Z, Deshpande SS, et al. Selective transcatheter ablation of the fast and slow pathways using radiofrequency energy in patients with atrioventricular nodal reentrant tachycardia. *Circulation* 1992;85:1318-28.
4. Blomstrom-Lundqvist C, Scheinman MM, Aliot EM, Alpert JS, Calkins H, Camm AJ, et al. ACC/AHA/ESC guidelines for the management of patients with supraventricular arrhythmias--executive summary. a report of the American college of cardiology/American heart association task force on practice guidelines and the European society of cardiology committee for practice guidelines (writing committee to develop guidelines for the management of patients with supraventricular arrhythmias) developed in collaboration with NASPE-Heart Rhythm Society. *J Am Coll Cardiol* 2003;42:1493-531.
5. Haissaguerre M, Gaita F, Fischer B, Commenges D, Montserrat P, d'Ivernois C, et al. Elimination of atrioventricular nodal reentrant tachycardia using discrete slow potentials to guide application of radiofrequency energy. *Circulation* 1992;85:2162-75.
6. Kalman J. Catheter Ablation of Atrioventricular Nodal Reentrant Tachycardia. In: Zipes D, Jalife J, editors. *Cadiac Electrophysiology. From Cell to Bedside*. Philadelphia: Saunders; 2004. p. 1069.
7. McGavigan AD, Rae AP, Cobbe SM, Rankin AC. Junctional rhythm - a suitable surrogate endpoint in catheter ablation of atrioventricular nodal reentry tachycardia? *Pacing Clin Electrophysiol* 2005;28:1052-4.
8. Bogun F, Knight B, Weiss R, Bahu M, Goyal R, Harvey M, et al. Slow pathway ablation in patients with documented but noninducible paroxysmal supraventricular tachycardia. *J Am Coll Cardiol* 1996;28:1000-4.

9. Jentzer JH, Goyal R, Williamson BD, Man KC, Niebauer M, Daoud E, et al. Analysis of junctional ectopy during radiofrequency ablation of the slow pathway in patients with atrioventricular nodal reentrant tachycardia. *Circulation* 1994;90:2820-6.
10. Knight BP, Zivin A, Souza J, Flemming M, Pelosi F, Goyal R, et al. A technique for the rapid diagnosis of atrial tachycardia in the electrophysiology laboratory. *J Am Coll Cardiol* 1999;33:775-81.
11. Leitch J, Klein GJ, Yee R, Murdock C. Invasive electrophysiologic evaluation of patients with supraventricular tachycardia. *Cardiol Clin* 1990;8:465-77.
12. Knight BP, Ebinger M, Oral H, Kim MH, Sticherling C, Pelosi F, et al. Diagnostic value of tachycardia features and pacing maneuvers during paroxysmal supraventricular tachycardia. *J Am Coll Cardiol* 2000;36:574-82.
13. Martinez-Alday JD, Almendral J, Arenal A, Ormaetxe JM, Pastor A, Villacastin JP, et al. Identification of concealed posteroseptal Kent pathways by comparison of ventriculoatrial intervals from apical and posterobasal right ventricular sites. *Circulation* 1994;89:1060-7.
14. Chen MC, Guo GB. Junctional tachycardia during radiofrequency ablation of the slow pathway in patients with AV nodal reentrant tachycardia: effects of autonomic blockade. *J Cardiovasc Electrophysiol* 1999;10:56-60.
15. Thibault B, de Bakker JM, Hocini M, Loh P, Wittkamp FH, Janse MJ. Origin of heat-induced accelerated junctional rhythm. *J Cardiovasc Electrophysiol* 1998;9:631-41.
16. Yu JC, Lauer MR, Young C, Liem LB, Hou C, Sung RJ. Localization of the origin of the atrioventricular junctional rhythm induced during selective ablation of slow-pathway conduction in patients with atrioventricular node reentrant tachycardia. *Am Heart J* 1996;131:937-46.
17. Hsieh MH, Chen SA, Tai CT, Yu WC, Chen YJ, Chang MS. Absence of junctional rhythm during successful slow-pathway ablation in patients with atrioventricular nodal reentrant tachycardia. *Circulation* 1998;98:2296-300.
18. Baker JH, Plumb VJ, Epstein AE, Kay GN. Predictors of recurrent atrioventricular nodal reentry after selective slow pathway ablation. *Am J Cardiol* 1994;73:765-9.
19. Poret P, Leclercq C, Gras D, Mansour H, Fauchier L, Daubert C, et al. Junctional rhythm during slow pathway radiofrequency ablation in patients with atrioventricular nodal reentrant tachycardia: beat-to-beat analysis and its prognostic value in relation to electrophysiologic and anatomic parameters. *J Cardiovasc Electrophysiol* 2000;11:405-12.
20. Li HG, Klein GJ, Stites HW, Zardini M, Morillo CA, Thakur RK, et al. Elimination of slow pathway conduction: an accurate indicator of clinical success after radiofrequency atrioventricular node modification. *J Am Coll Cardiol* 1993;22:1849-53.
21. Lipscomb KJ, Zaidi AM, Fitzpatrick AP, Lefroy D. Slow pathway modification for atrioventricular node re-entrant tachycardia: fast junctional tachycardia predicts adverse



prognosis. Heart 2001;85:44-7.

22. Tebbenjohans J, Pfeiffer D, Schumacher B, Manz M, Luderitz B. Impact of the local atrial electrogram in AV nodal reentrant tachycardias: ablation versus modification of the slow pathway. J Cardiovasc Electrophysiol 1995;6:245-51.

23. Lee K, Chun H, Liem LB. Effect of adenosine and verapamil in catecholamine-induced accelerated atrioventricular junctional rhythm: insights into the underlying mechanism. Pacing Clin Electrophysiol 1999;22:866-70.

24. Matsushita T, Chun S, Sung RJ. Influence of isoproterenol on the accelerated junctional rhythm observed during radiofrequency catheter ablation of atrioventricular nodal slow pathway conduction. Am Heart J 2001;142:664-8.

25. Olguin J, Zipes D. Specific Arrhythmias: Diagnosis and Treatment. In: Braunwald E, Zipes D, Libby P, editors. Heart Disease. A Textbook of Cardiovascular Disease. 6 th ed. Philadelphia: W.B Saunders Company; 2001. p. 821.

26. Iakobishvili Z, Kusniec J, Shohat-Zabarsky R, Mazur A, Battler A, Strasberg B. Junctional rhythm quantity and duration during slow pathway radiofrequency ablation in patients with atrioventricular nodal re-entry supraventricular tachycardia. Europace 2006;8:588-91.

27. Wagshal AB, Crystal E, Katz A. Patterns of accelerated junctional rhythm during slow pathway catheter ablation for atrioventricular nodal reentrant tachycardia: temperature dependence, prognostic value, and insights into the nature of the slow pathway. J Cardiovasc Electrophysiol 2000;11:244-54.

28. Thakur RK, Klein GJ, Yee R, Stites HW. Junctional tachycardia: a useful marker during radiofrequency ablation for atrioventricular node reentrant tachycardia. J Am Coll Cardiol 1993;22:1706-10.

29. Hsieh MH, Chen SA, Tai CT, Chiang CE, Chang MS. Electrophysiologic characteristics of different ectopic rhythms during slow pathway ablation in patients with atrioventricular nodal reentrant tachycardia. J Interv Card Electrophysiol 1998;2:203-9.

30. Strickberger SA, Daoud E, Weiss R. A randomized comparison of fixed power and temperature monitoring during slow pathway ablation in patients with atrioventricular nodal reentrant tachycardia. J Cardiovasc Electrophysiol 1997;7:299-303.