

John D. Fisher, MD



### Multisite pacing in CRT – does it help?

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#### **Disclosures**

- Consultant: Medtronic.
- Investigator: many device & drug companies.
- ♥ Fellowship support: Boston Scientific, Medtronic, St. Jude Medical, Biotronik

### **Dual-Site Pacing**

- Atrial
  - For prevention/reduction of AF
  - Remains controversial
- For termination (ATP) of atrial and ventricular arrhythmias (Mehra and others).
- For atrial and ventricular CRT.
  - Some early Bi-V CRT was called "multisite".

# Multisite Pacing (MSP) What are we talking about?

- ♥ RA + LA pacing.
- ♥ RV + LV pacing.
- ♥ RA + RA pacing.
- RV + RV pacing
- ♥ RV + RV + LV pacing.
- **♥**RV +LV + LV pacing.
  - Present area of greatest interest

### Pilot for dual site atrial pacing to prevent atrial fibrillation (DAPPAF)

#### Long-Term Outcome of Patients With Drug-Refractory Atrial Flutter and Fibrillation After Single- and Dual-Site Right Atrial Pacing for Arrhythmia Prevention

PHILIPPE DELFAUT, MD, SANJEEV SAKSENA, MD, FACC, ATUL PRAKASH, MD, MRCP, RYSZARD B. KROL, MD, Pid

Millburn, New Jersey

Objectives. An initial crossover study comparing dual- and single-site right atrial pacing was performed followed by a longterm efficacy and safety evaluation of dual-site right atrial pacing in patients with drug-refractory atrial fibrillation (AF). Also examined was the efficacy of two single-site right atrial pacing modes (high right atrium and coronary sinus ostium) and the long-term need for cardioversion, antithrombotic and antiarrhythmic drug therapies during dual-site atrial pacing.

Methods. Thirty consecutive patients with drug-refractory symptomatic AF and documented primary or drug-induced bradycardia were implanted with a dual chamber rate-responsive pacemaker and two atrial leads. Single-site atrial pacing was performed at the high right atrium or the coronary sinus ostium. Continuous atrial pacing was maintained.

Results. Mean arrhythmia-free intervals increased from  $9 \pm 10$  days in the control period preceding implant to  $143 \pm 110$  days (p < 0.0001) in single-site right atrial pacing and  $195 \pm 96$  days in dual-site right atrial pacing (p < 0.005 versus single-site pacing and p < 0.0001 versus control). Dual-site right atrial pacing significantly increased the proportion of patients free of AF

recurrence (89%) as compared to single-site right atrial pacing (62%, p = 0.02). High right atrial pacing and coronary sinus ostial pacing had similar efficacy for AF prevention. Effective rhythm control was achieved in 86% of patients during dual right atrial pacing. Seventy-eight percent of patients at 1 year and 56% at 3 years remained free of symptomatic AF. The need for cardioversion was reduced after pacemaker implant (p < 0.05) and antithrombotic therapy was reduced (p < 0.06) without any thromboembolic event. Coronary sinus ostial lead dislodgement was not observed after discharge.

Conclusions. Atrial pacing in combination with antiarrhythmic drugs eliminates or markedly reduces recurrent AF. Prevention of AF is enhanced by dual-site right atrial pacing. High right atrial and coronary sinus ostial pacing do not differ in efficacy. Dual-site right atrial pacing is safe, achieves long-term rhythm control in most patients, decreases the need for cardioversion, and antithrombotic therapy can be selectively reduced.

> (J Am Coll Cardiol 1998;32:1900-8) ©1998 by the American College of Cardiology

# Conventional CRT (RA) + RV + LV

#### Multisite Pacing for End-Stage Heart Failure: Early Experience

SERGE CAZEAU, PHILIPPE RITTER, ARNAUD LAZARUS, DANIEL GRAS, HAYSSAM BACKDACH, OLIVIER MUNDLER,\* and JACQUES MUGICA

From the Centre Chirurgical du Val d'Or, Saint-Cloud, and the "Service de Médecine Nuclinire, Hôpital Lariboisière, Paris, France

CAZEAU, S., ET AL.: Multisite Pacing for End-Stage Heart Failure: Early Experience. Our objective was to improve hemodynamics by synchronous right and left site ventricular pacing in patients with severe congestive heart failure (CHF). Previous studies reported a benefit of dual chamber pacing with a short AV delay in patients with severe CHF. Other works, however, show contradictory results. Deleterious effects due to a desynchronization of right (RV) and left ventricular (LV) contractions have been suggested. This study included eight subjects with widened QBS and end-stage heart failure despite maximal medical therapy, who refused, or were not eligible to undergo heart transplantation. Each patient underwent a baseline, invasive hemodynamic evaluation with insertion of three temporary leads to allow different pacing configurations, including RV apex and outflow tract pacing, and biventricular pacing between the RV outflow tract and LV and RV apex and LV. According to the results of this baseline study, the configuration of preexistent pacemakers was modified or new systems were implanted to allow hiventricular pacing, which, in patients with sinus rhythm, was atrial triggered. Biventricular pacing increased the mean cardiac index (Cl) by 25% (from a baseline of 1.83 ± 0.30 L/min per m<sup>2</sup>, P < 0.006), decreased the mean V wave by 26% (from a baseline of 36  $\pm$  12 mmHg, P < 0.004), and decreased pulmonary capillary wedge pressure by 17% (from a baseline of 31  $\pm$  10 mmHg, P < 0.01). Four patients died (1 preoperatively, 1 intraoperatively, 2 within 3 months, and 1 of a noncardiac cause). The four surviving patients have clinically improved from New York Heart Association Functional Class IV to Class II. In these survivors, CI decreased by 15% (P < 0.007) when multisite pacing was turned off during follow-up. In patients with end-stage heart failure, multisite pacing may be associated with a rapid and sustained hemodynamic improvement. (PACE 1996; 19[Pt. II]:1748-1757).

#### Cazeau et al 1996

- "Multisite" here = conventional CRT
  - RVA or RVOT to LV produced favorable results.

# CRT may favor Reversion of AF to SR, (...which may further improve CHF?)

Clin Res Cardiol 98:189-194 (2009) DOI 10.1007/s00392-008-0740-z

ORIGINAL PAPER

Melanie Hauck Alexander Bauer Frederik Voss Hugo A. Katus Ruediger Becker Effect of cardiac resynchronization therapy on conversion of persistent atrial fibrillation to sinus rhythm

### Dual Site RV RV + RV

Some reports that if an LV site cannot be achieved, then dual RV may be better than single site RV.



Dual-site right ventricular pacing.

A rescue alternative in cardiac resynchronisation therapy implantation failure? More efficient stimulation for patients with borderline cardiac resynchronisation therapy indication?

Less harmful ventricular pacing?

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Department of Cardiology, Medical University, Lublin, Poland





#### CASE REPORT

#### Cardiac resynchronisation therapy versus dual site right ventricular pacing in a patient with permanent pacemaker and congestive heart failure

Oruganti Sai Satish \*, Kuan-Hung Yeh b, Ming-Shien Wen b, Chun-Chieh Wang b. \*

Submitted & September 2004, and accepted after revision 30 January 2005

#### KEYWORDS

cardisc resynchronisation therapy; heart failure; complete heart block; pacomaker; atrial fibrillation Abstract: A 46-year-old male patient who had long-term right vertricular (RV) pacing for symptomatic complete heart block, initially by an epicardial, later with an endocardial pacing lead at the RV apex, developed congestive heart failure (CHF) and chronic strial fibrillation 7 years following the pacemaker implantation and was medically treated. During follow-up, his pacemaker was upgraded to a cardiac respectionisation therapy (CRT) device, because of uncontrolled CHF symptoms, New York Heart Association (HTHA) functional class IV, while on drugs. The patient's symptomatic status improved to NYtHA functional class II with CRT. After 17 months of CRT, the battery became depleted, because of the high capture threshold of the left ventricular lead. The patient was then given dual site RV pacing (RV outflow tract + RV apex) in place of CRT, which showed similar efficacy at 52 weeks follow-up.

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### Dual site RV + single-site LV. RV + RV + LV

#### Dual-site right ventricular and left ventricular pacing in a patient with left ventricular systolic dysfunction and atrial fibrillation using a standard CRT-D device



David Chase a., Vipin Kumar\*, Amit Hooda\*

J Saudi Heart Assoc 2013

\*Christian Medical College Hospital, Velices, Tambrado 632000

In patients undergoing cardiac resynchronication therapy with delibrillator (CRT-D) implantation for left ventricular systolic dysfunction (LVSD) accompanied by permanent atrial fibrillation (AF), generally, the unused strial port in plugged at device implantation. We describe an alternative use for the atrial-port in this case report.

A 43 year old gentleman with LVSD due to left ventricular non-compaction (LVNC) and AF of unknown dutation underwent a CRT-D implantation after optimization of cardiac tailure treatment. The atrial-part which would otherwise have been plugged was connected to a high right ventricular septal (RVS) paring-lead and the shock-lead was positioned at the right ventricular ages (RVA). This approach permitted modified cardiac resynchronization in a high RVS to left ventricular (LV) and RVA pacing sequence using the high RVS and LV pacing combined with a shock vector including the RV ages. A standard CRT-D device with a minimum programmable A-V delay of 36 ms (technically RVS to LV delay in the 'DDD' pacing mode) was used. The device was programmed to a 'DDO' pacing mode (sequential multi-site ventricular pacing with some programmability). The mode switch openation was programmed 'OBF' since atrial sensing is unavailable. Device-delivered shacks did not cardiovert the patient back to sixus rhythm suggesting that the AF was permanent (no prior cardioversion attempts were made on the presumption that the chances of maintaining sinus rhythm, given the underlying cardiac condition, were low). Subsequently, the patient required radio-frequency ablation of the atrio-ventricular node for conducted AF. Symptomatic, orbicardiographic and radiological improvement preceded atrio-ventricular node ablation.

Conclusion: Amongst AF patients with permanent AF undergoing CRT-D implantation, those patients who are likely to have the CRT-D device atrial-ports plugged could benefit from having both the options of (i) a RVA shock vector as well as (ii) a high RVS-pacing feasible, by utilizing the atrial-port of a conventional CRTD device for a RVS pacing lead, should a RVA shock-lead position be preferred. New device programming algorithms will be necessary to make putient-customized programming in this lead configuration firelible, more useful clinically and easy.

<sup>\*</sup>India

### Dual site RV, + LV CRT Chase et al 2013

- AF patient.
- A-port connected to RVS lead.
  - Near 0ms AV I.
- RV & LV ports connected normally.
- Good response.

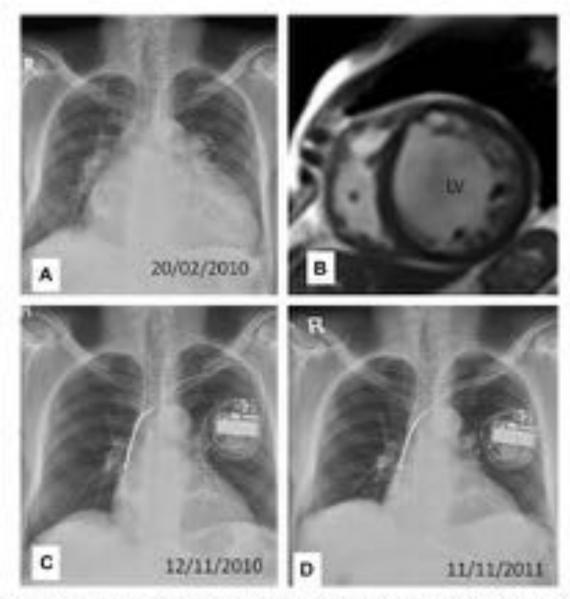


Figure 3: A  $\phi$  8 - Biseline closel X-ray and ME image; C  $\phi$  D - Sorial follow up closel X-ray images.



# Effect of triangle ventricular pacing on haemodynamics and dyssynchrony in patients with advanced heart failure: a comparison study with conventional bi-ventricular pacing therapy

Kentaro Yoshida<sup>†</sup>, Yoshihiro Seo\*<sup>†</sup>, Hiro Yamasaki, Kazuyuki Tanoue, Nobuyuki Murakoshi, Tomoko Ishizu, Yukio Sekiguchi, Satoru Kawano, Sadanori Otsuka, Shigeyuki Watanabe, Iwao Yamaguchi, and Kazutaka Aonuma

- RV + RV + LV
- **♥ 21 patients, over medium-term follow up.**
- Dual RV, + 1LV better than standard CRT.
- ♥ No RCT yet.

### However...

#### Right Ventricular Outflow Versus Apical Pacing in Pacemaker Patients with Congestive Heart Failure and Atrial Fibrillation

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BRUCE VAN NAITA, M.D., † † and BRUCE L. WELKOFF, M.D., ### for the ROVA Inventigation.

From Transmits Hospitals of Christiand, Flast Western Research Christian, Ohn, USA, (Made of College of Vergion, Vergion, Entercoverable Versional, Microsoft, Vergion, USA, Dis. Subs Medical CRMS. Symbol. Symbol. Statistics, USA, (Consoling) of Schools, Orbeita, Noticeala, USA, Western, Ohn, USA, Ohn, USA, Marian Christian, USA, Microsoft Christian, Ohn, USA, Marian Christian, Mina, Astron., USA, (Ohndryberta) Start Group, Afragarque, New Mirania, USA, (Original Massa Research Christ. South, Westington, USA, (Ohndryberta) Start Group, Afragarque, New Mirania, USA, (Original Massa Research Christ. South, Westington, USA, (Ohndryberta) Start Group, Afragarque, New Miranian, Vergion, Marian, Start, Christian, USA, (Ohndryberta), Colleges Cart Group, Marian, Group, USA, (Ohndryberta), USA, (Ohndrybe

Right Ventricular Pacing Site in Bleast Failure, templecine: Prior make suggest that right ventricular optical (EVX) pacing for distriction effects. Whether the right ventricular unifore tract (EVXE) is a more optimal site for permanent pacing in potents with congretive heart failure (CSE) has not been constituted.

Methods and Broads: We constructed a translaminal, cross-over trial to determine whether quality of Bir (QCL) is better after 3 months of BYOT than BYO, parting in 165 passession recipions with CBC, left ventricular (LV) conside dedunction (LV) ejection fraction (LM), and chronic atrial liberitation (LV). An additional aim was in compare dual-site (BYOT ± BYO). How delays with single-site BYO, and BYOT paring, QEO duration was chertor during BYOT (DCT ± BYO), but and dual-site (BST ± DF met Box BYO) paring, (TM ± DF me, P < 0.000), At a months, the BYOT group had higher (P ± 0.01) rein-continual QCE, subscale scores than the BYO, group, At-Connection, there were no significant differences in QCE, some between BYOT and BYO, groups, Comparing BYOT to BYO, paring within the came patient, montal health subscale scores must better (P ± 0.00) during BYOT pacing, After 7 months of biline-up, LVEF was higher (P = 0.00) in those analysed to BYO, nather than BYOT pacing between months a and 2, After 3 months of dual-site BY pacing, physical functioning was works (P = 0.00) than during BYOT pacing, and New York Blood through BYO, pacing, montal health was works (P = 0.02) than during BYOT pacing. There were no other significant differences between BYO, BYOT and dual-site BY pacing in QXX, source, NYEA class, distance walked in 6 minutes, LV species. BYOT and dual-site BY pacing in QXX, source, NYEA class, distance walked in 6 minutes, LV species.

Conclusion: In patients with CRE, LV dynhauction, and chronic AE, RVOT and dual-site RV pacing shorten QES duration but ofter 3 months do not contributely improve QEA, or other clinical entropes computed with RVA pacing, of Continuous Encountering, lot, As you (ARS-200), Newcolor (2003).

### Disappointing Stambler et al, JCE 2003

- In pts with CHF, LV Dysfunction, CAF:
  - RVOT or dual site RV pacing shorten QRS.
    - But after 3 months, no change in QOL or other outcomes compared with RVA pacing.
  - Thus CRT with LV pacing is needed.

### **Multisite LV pacing**

Europace doi:10.1093/europace/euu197

# A review of multisite pacing to achieve cardiac resynchronization therapy

Christopher Aldo Rinaldi<sup>1\*</sup>, Haran Burri<sup>2</sup>, Bernard Thibault<sup>3</sup>, Antonio Curnis<sup>4</sup>, Archana Rao<sup>5</sup>, Daniel Gras<sup>6</sup>, Johannes Sperzel<sup>7</sup>, Jagmeet P. Singh<sup>8</sup>, Mauro Biffi<sup>9</sup>, Pierre Bordachar<sup>10</sup>, and Christophe Leclercq<sup>11</sup>

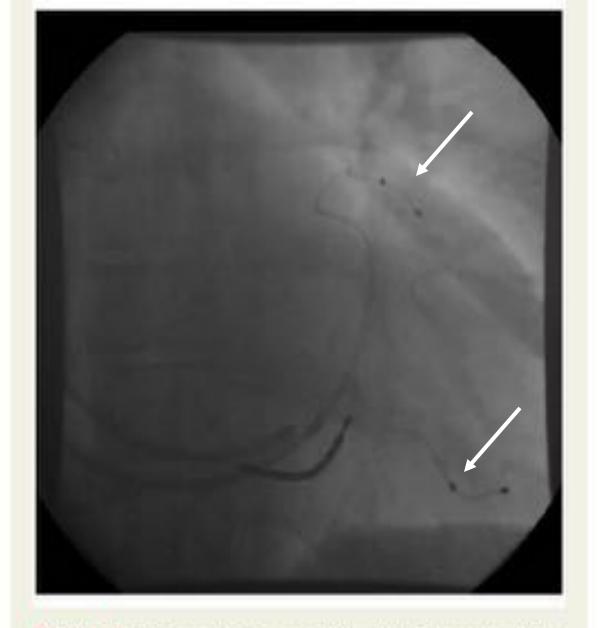


Figure 1 Anteroposterior projection of deployment of two bipolar LV pacing leads in separate branches of the CS (anterolateral and posterolateral) to achieve dual LV lead CRT.

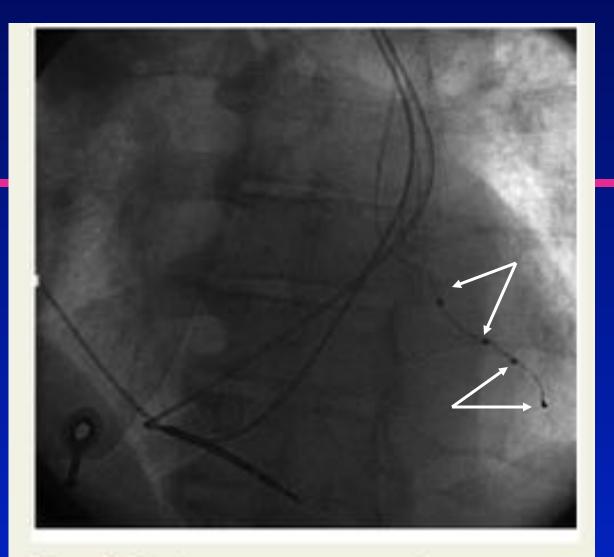


Figure 2 Top: Anteroposterior projection. Coronary sinus angiography performed with a telescopic catheter placed into a lateral cardiac vein. Bottom: LAO 30° projection. Implantation of a quadripolar LV lead using a similar approach compared with a regular bipolar LV lead. The patient has a persistent left-sided superior vena cava.

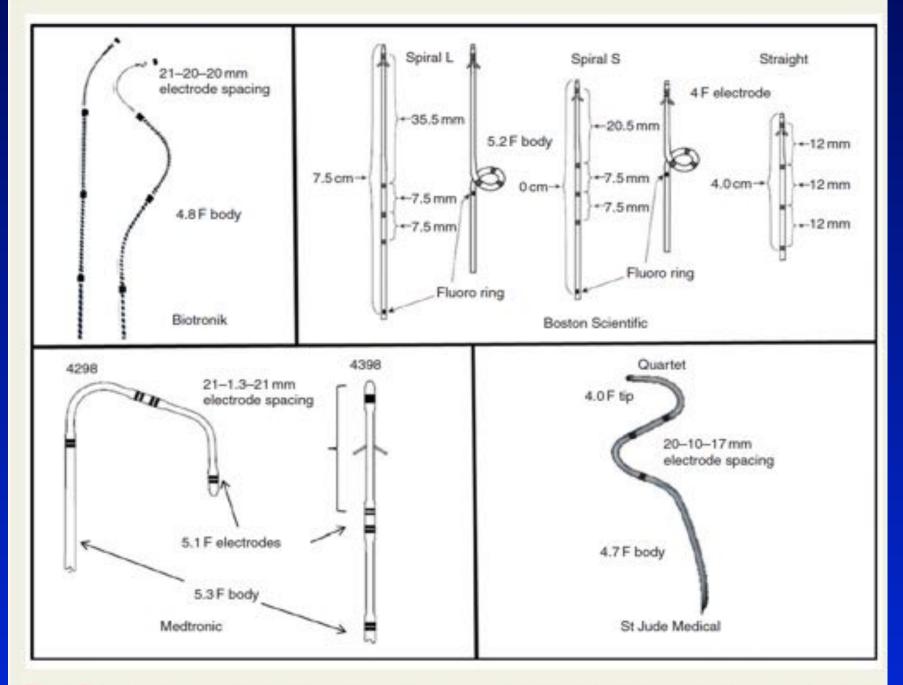


Figure 3 Currently available quadripolar LV leads, showing differences in lead design (reproduced/adapted and with permission from Boston Scientific, St Jude Medical, and Medtronic).

### Features of Quadripolar LV Leads

- ▼ Need only 1 LV lead.
- Choice of poles may avoid phrenic nerve stimulation.
- May or may not provide multisite LV stimulation depending on anatomy.
- May or may not provide CRT equal to 2 separate LV leads.

## Ischemic CM may benefit most from Multisite LV or Endocardial Pacing

#### ♥ Ginks Circ A&E 2012, Europace 2012

Benefits of Endocardial and Multisite Pacing Are Dependent on the Type of Left Ventricular Electric Activation Pattern and Presence of Ischemic Heart Disease Insights from Electroanatomic Mapping

Matthew R. Ginks, MD; Anoop K. Shetty, MBBS; Pier D. Lambiase, PhD; Simon G. Duckett, MD; Julian Bostock, MSc; Janet L. Peacock, PhD; Kawal S. Rhode, PhD; Cliff Bucknall, MD; Jaswinder Gill, MD; Peter Taggart, MD; Christophe Leclercq, MD; Gerald S. Carr-White, PhD; Reza Razavi, MD; C. Aldo Rinaldi, MD



Europace (2012) 14, 373-379 doi:10.1093/europace/eur336

#### CLINICAL RESEARCH

Pacing and Resynchronization Therapy

Multi-site left ventricular pacing as a potential treatment for patients with postero-lateral scar: insights from cardiac magnetic resonance imaging and invasive haemodynamic assessment

Matthew R. Ginks<sup>1,2\*</sup>, Simon G. Duckett<sup>1,2</sup>, Stamatis Kapetanakis<sup>1</sup>, Julian Bostock<sup>1</sup>, Shoaib Hamid<sup>1</sup>, Anoop Shetty<sup>1,2</sup>, Yingliang Ma<sup>2</sup>, Kawal S. Rhode<sup>2</sup>, Gerald S. Carr-White<sup>1</sup>, Reza S. Razavi<sup>1,2</sup>, and C. Aldo Rinaldi<sup>1,2</sup>

#### LV MSP can overcome scar limitations of CRT

#### Biophysical Modeling to Simulate the Response to Multisite Left Ventricular Stimulation Using a Quadripolar Pacing Lead

STEVEN A. NIEDERER, Ph.D.,\* A.K. SHETTY, M.B.B.S.,\*,† G. PLANK, Ph.D.,‡ J. BOSTOCK, M.B.B.S.,† R. RAZAVI, M.D.,\*,† N.P. SMITH, Ph.D.,\*,§ and C.A. RINALDI, M.D.,\*,†

From the \*Imaging Sciences & Biomedical Engineering Division, King's College London, London, United Kingdom; and †Department of Cardiology, St Thomas' Hospital, London, United Kingdom; ‡Institut für Biophysik, Medizinische Universität Graz, Graz, Austria; and §Computing Laboratory, University of Oxford, Oxford, United Kingdom

Background: Response to cardiac resynchronization therapy (CRT) is reduced in patients with posterolateral scar. Multipolar pacing leads offer the ability to select desirable pacing sites and/or stimulate from multiple pacing sites concurrently using a single lead position. Despite this potential, the clinical evaluation and identification of metrics for optimization of multisite CRT (MCRT) has not been performed.

Methods: The efficacy of MCRT via a quadripolar lead with two left ventricular (LV) pacing sites in conjunction with right ventricular pacing was compared with single-site LV pacing using a coupled electromechanical biophysical model of the human heart with no, mild, or severe scar in the LV posterolateral wall.

Result: The maximum dP/dt<sub>max</sub> improvement from baseline was 21%, 23%, and 21% for standard CRT versus 22%, 24%, and 25% for MCRT for no, mild, and severe scar, respectively. In the presence of severe scar, there was an incremental benefit of multisite versus standard CRT (25% vs 21%, 19% relative improvement in response). Minimizing total activation time (analogous to QRS duration) or minimizing the activation time of short-axis slices of the heart did not correlate with CRT response. The peak electrical activation wave area in the LV corresponded with CRT response with an R<sup>2</sup> value between 0.42 and 0.75.

Conclusion: Biophysical modeling predicts that in the presence of posterolateral scar MCRT offers an improved response over conventional CRT. Maximizing the activation wave area in the LV had the most consistent correlation with CRT response, independent of pacing protocol, scar size, or lead location. (PACE 2012; 35:204–214)

#### CLINICAL RESPONSE TO MULTISITE BIVENTRICULAR PACING

Poster Contributions
Poster Sessions, Expo North
Saturday, March 09, 2013, 3:45 p.m.-4:30 p.m.

**ACC 2013** 

Session Title: Heart Failure: Cardiac Resynchronization Therapy

Abstract Category: 17. Heart Failure: Therapy

Presentation Number: 1177-311

Authors: Jose Cuellar-Silva, Dan Dan, Andrew Wickliffe, Thomas Deering, Serge Cazeau, Ioanna Kosmidou, Piedmont Heart Institute, Atlanta, GA, USA, Georgia Health Sciences University, Augusta, GA, USA

Background: Lack of response to cardiac resynchronization therapy (CRT) remains a clinical challenge and is predominantly attributed to suboptimal LV lead positioning. Multisite pacing (MS-CRT) has emerged as an alternative to biventricular pacing, however clinical responses have not been thoroughly investigated.

Methods: 66 patients (45 males) with CHF (NYHA III and ambulatory IIII) underwent initial MS-CRT (ICRT, n=49) or upgrade to MS-CRT from standard CRT after worsening of clinical status or adverse remodeling (UCRT, n=17). Major adverse clinical events (MACE) were death, heart failure (HF) hospitalization and cardiac transplantation or ventricular assist device implant at one year.

Results: Clinical characteristics and baseline LVEF, were similar between groups except for AF (64.7% vs 33.3% for the UCRT and ICRT groups, p=0.025) and history of ventricular arrhythmias (41.3% vs 16.7% for the UCRT and ICRT groups, p=0.039). Mean LVEF change after implant was not different between the ICRT and UCRT groups (8.0±8% vs 4.1±6.3%, p=0.27). At one year, the probability for MACE was similar between the two groups (LR=0.892). Time to first HF hospitalization was not different between groups (222.3±134d vs 267.2±118d for the ICRT and UCRT groups, p=0.818).

Conclusions: Upgrade to MS-CRT in CRT non-responders results in clinical response similar to the observed response with initial MS-CRT implant.

Multisite pacing may be useful in CRT non responders.



# Acute haemodynamic comparison of multisite and biventricular pacing with a quadripolar left ventricular lead

Bernard Thibault<sup>1</sup>\*, Marc Dubuc<sup>1</sup>, Paul Khairy<sup>1</sup>, Peter G. Guerra<sup>1</sup>, Laurent Macle<sup>1</sup>, Lena Rivard<sup>1</sup>, Denis Roy<sup>1</sup>, Mario Talajic<sup>1</sup>, Edward Karst<sup>2</sup>, Kyungmoo Ryu<sup>2</sup>, Patrice Paiement<sup>3</sup>, and Taraneh G. Farazi<sup>2</sup>

<sup>1</sup>Department of Medicine and Research Center, Montreal Heart Institute and Université de Montréal, 5000 Belanger Street, Montreal, Quebec, QC H1T 1C8, Canada,
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Received 27 September 2012; accepted after revision 19 December 2012; online publish-ahead-of-print 27 February 2013

#### Aims

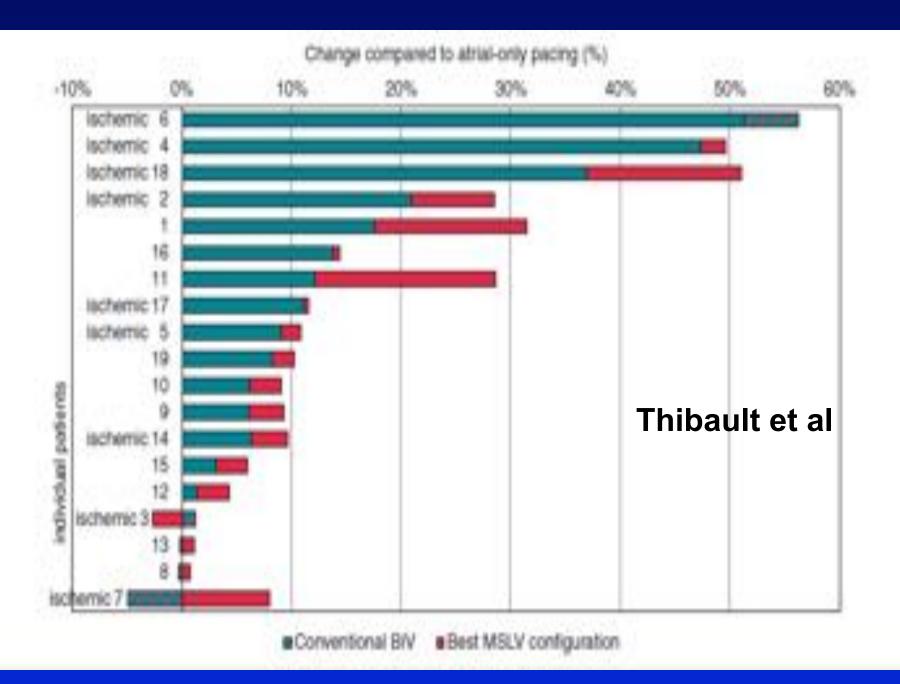
Pacing from multiple sites in the left ventricle (LV) may bring about further resynchronization of the diseased heart compared with biventricular (BiV) pacing. We compared acute haemodynamic response (LV dP/dt<sub>max</sub>) of multisite and BiV pacing using a quadripolar LV lead.

#### Methods and results

In 21 patients receiving cardiac resynchronization therapy, a quadripolar LV lead and conventional right atrial and ventricular leads were connected to an external pacing system. A guidewire pressure sensor was placed in the LV for continuous dP/dt measurement. Four multisite pacing configurations were tested three times each and compared with BiV pacing using the distal LV electrode. Nineteen patients had useable haemodynamic data. Median increase in LV dP/dt<sub>max</sub> with BiV vs. atrial-only pacing was 8.2% (interquartile range 2.3%, 15.7%). With multisite pacing using distal and proximal LV electrodes, median increase in LV dP/dt<sub>max</sub> was 10.2% compared with atrial-only pacing (interquartile range 6.1%, 25.6%). In 16 of 19 patients (84%), two or more of the four multisite pacing configurations increased LV dP/dt<sub>max</sub> compared with BiV pacing. Overall, 72% of all tested configurations of multisite pacing produced greater LV dP/dt<sub>max</sub> than obtained with BiV pacing. Pacing from most distal and proximal electrodes was the most common optimal configuration, superior to BiV pacing in 74% of patients.

#### Conclusion

In the majority of patients, multisite pacing improved acute systolic function further compared with BiV pacing. Pacing with the most distal and proximal electrodes of the quadripolar LV lead most commonly yielded greatest LV dP/dt.....



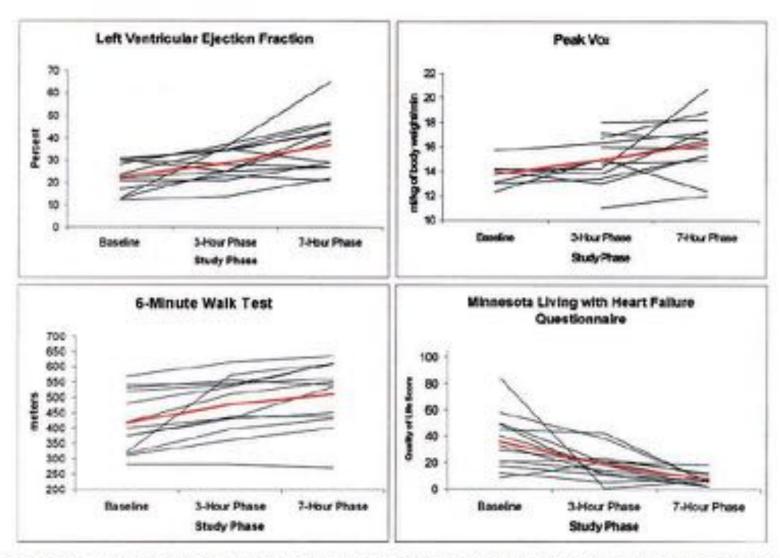


Figure 3. Changes in echocardiographic left ventricular systolic and diastolic function, maximal and submaximal exercise function, and clinical course during follow-up. A: Time course of left ventricular ejection fraction in the study phases compared with baseline in single patients (black lines) and in the overall study population (red line). B. C: Time course of the peak Voy (in mL/kg of body weight/min) and the distance walked in to minutes (in meters) in the study phases compared with baseline in single patients (black lines) and in the overall study population (red line), respectively. D: Patients' perception of the effects of heart failure on their daily lives (access of the Minnesota Living with Beart Failure Questionnaire) in the study phases compared with baseline in single patients (black lines) and in the overall study population (red line). Subjects had mean scores that were substantially above the benchmark of 10 (e.g., reduced quality of life).

Table I	Studies of MPS delivered by a quadripolar	
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Author, year	Number of patients	Study type	Findings
Thibault et al. (2013)	19 (21)	Acute comparative study  Measurement invasive haemodynamic evaluation (dP/dt)	72% of patients, MPP improved acute systolic function vs. conventional CRT. Pacing most distal and proximal electrodes most commonly yielded greatest LVdP/dt <sub>max</sub>
Rinaldi et al. (2013)	41(52)	Comparative study after implant Measurement echocardiographic dyssynchrony (TDI)	64% of patients MPP resulted in significant reduction in dyssynchrony vs. conventional CRT
Pappone et al. (2013)	44	Randomized comparative study at the time of implant Measurement: invasive haemodynamic evaluation (pressure—volume loops)	Main finding: CRT with MPP can significantly improve acute LV haemodynamic parameters assessed with PV loop measurements as compared with conventional CRT

But remember RV + RV + LV Also poduced favorable early results That lessened over time.

# How many sites are needed to optimize MSP?

Studied in canines with chronic LBBB

#### Acute electrical and hemodynamic effects of multisite left ventricular pacing for cardiac resynchronization therapy in the dyssynchronous canine heart

Sylvain Ploux, MD, \*\*\* Marc Strik, MD, \*\* Arne van Hunnik, BSc, \*\* Lars van Middendorp, MD, \*\*
Marion Kuiper, BSc, \*\* Frits W. Prinzen, PhD \*\*

From the "Department of Physiology, Cardiovascular Research Institute Maastricht, Maastricht University, Maastricht, The Netherlands, <sup>†</sup>Höpital de Haut-Lévêque, CHU de Bordeaux, Pessac, France, and <sup>‡</sup>L'Institut de Rythmologie et modélisation Cardiaque, Université de Bordeaux Segalen, Bordeaux, France.

BACKGROUND Multisite left ventricular (multi-LV) epicardial pacing has been proposed as an alternative to conventional single-site LV (single-LV) pacing to increase the efficacy of cardiac resynchronization therapy.

OBJECTIVE To compare the effects of multi-LV versus single-LV pacing in dogs with left bundle branch block (LBBB).

METHODS Studies were performed in 9 anaesthetized dogs with chronic LBBB using 7 LV epicardial electrodes. Each electrode was tested alone and in combination with 1, 2, 3, and 6 other electrodes, the sequence of which was chosen on the basis of practical real-time electrical mapping to determine the site of the latest activation. LV total activation time (LVTAT) and dispersion of repolarization (DRep) were measured by using approximately 100 electrodes around the ventricles. LV contractifity was assessed as the maximum derivative of left ventricular pressure (LVdP/dt<sub>max</sub>).

RESULTS Single-LV pacing provided, on average, a  $-4.0\% \pm 9.3\%$  change in LVTAT and  $0.2\% \pm 13.7\%$  change in DRep. Multi-LV pacing markedly decreased both LVTAT and DRep in a stepwise fashion to reach  $-41.3\% \pm 5\%$  (P < .001 for overall comparison) and  $-14.2\% \pm 19.5\%$  (P < .02 for overall comparison) in the septuple-LV pacing configuration, respectively. Single-LV pacing provided a mean increase of  $10.7\% \pm 7.7\%$  in LVdP/dtmax

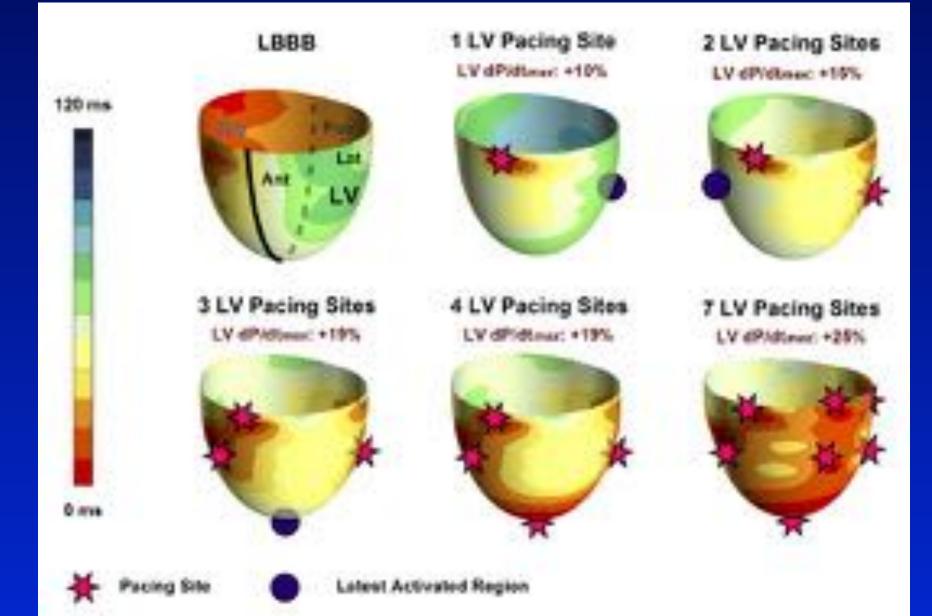
 ${\rm LVdP/dt_{max}}$  incrementally increased by the addition of pacing electrodes to 16.4%  $\pm$  8.7% (P<.001 for overall comparison). High response to single-LV pacing could not be improved further during multi-LV pacing.

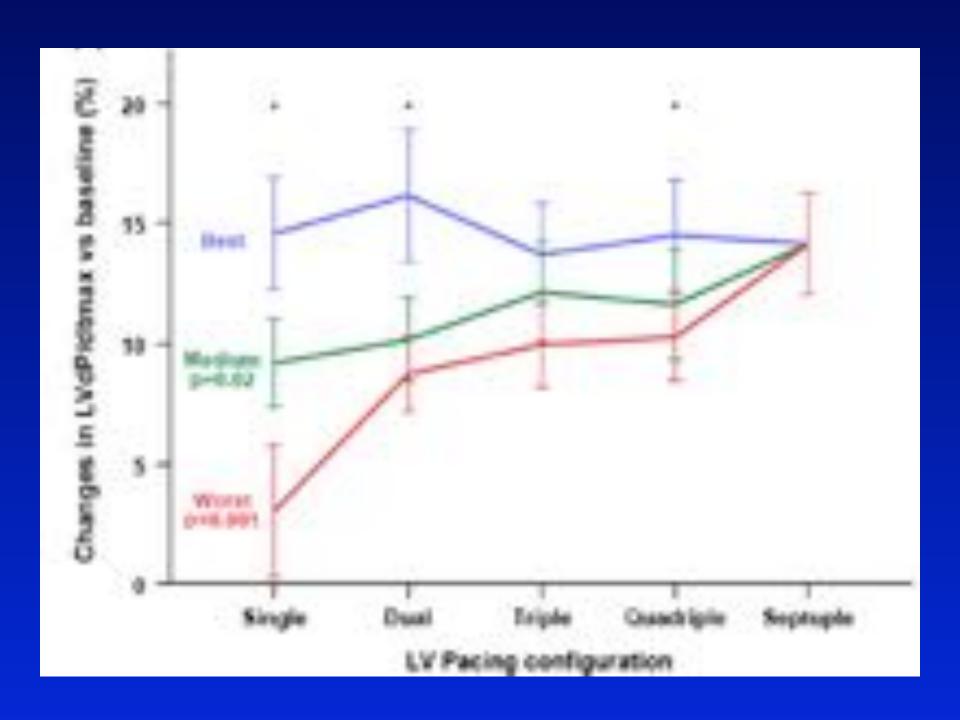
CONCLUSIONS Compared with single-LV pacing, multi-LV pacing can considerably reduce both LVTAT and DRep in dogs with LBBB, but the improvement in contractility is limited to conditions where single-LV pacing provides suboptimal improvement. Further studies are warranted to determine whether these acute effects translate in antiarrhythmic properties and better long-term outcomes.

KEYWORDS Cardiac resynchronization therapy; Multisite left ventricular pacing: Heart failure; Left bundle branch block; Cardiac mapping; Biventricular pacing

ABBREVIATIONS CRT = cardiac resynchronization therapy; DRep = dispersion of repolarization; LBBB = left bundle branch block; LV = left ventricular; LVdP/dt<sub>mex</sub> = maximum derivative of left ventricular pressure; LVTAT = left ventricular total activation time; multi-LV = multisite left ventricular; RV = right ventricular; single-LV = single-site left ventricular

(Heart Rhythm 2014;11:119–125) © 2014 Heart Rhythm Society. All. rights reserved.





### **MultiPoint Pacing Trial**

ClinTrials.gov NCT02066467

- BSC-Guidant European trial.
- **2014-2016**
- **V** Randomize:
  - Conventional CRT.
  - CRT-MSP with Quad LV lead.

### **MORE-CRT**

### ClinTrials.gov NCT02006069

- SJM European trial.
- **2014-2017.**
- All get quad LV lead but programmed to conventional CRT.
- **After 6 months:** 
  - responders continued as CRT.
  - Non-responders randomized to conventional or MSP-CRT.

### **Hierarchy?**

- **♥** Sometimes a good LV position cannot be achieved through the coronary sinus.
- **▼** A "hierarchy" has been suggested:
  - RV + LV
    - Maybe RV + LV + LV.
    - Or RV + RV +LV
  - Dual site RV (RV + RV).
  - RVOT.
  - RVA. Known to be bad with poor LV function.

### "Thinking out of the Box"

# Impulse Dynamics™ Multisite RV Cardiac Contractility Modulation (CCM) delivered during RP at multiple RV sites

First Human Chronic Experience with Cardiac Contractility
Modulation by Nonexcitatory Electrical Currents for Treating
Systolic Heart Failure: Mid-Term Safety and Efficacy Results
from a Multicenter Study

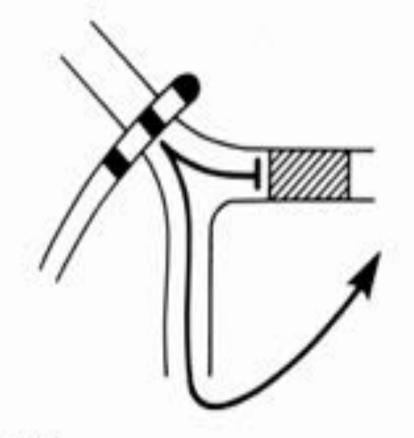
CARLO PAPPONE, M.D., Ph.D., GIUSEPPE AUGELLO, M.D.,
SALVATORE ROSANIO, M.D., Ph.D., GABRIELE VICEDOMINI, M.D.,
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GUENTER STIX, M.D., and HERWIG SCHMIDINGER, M.D.

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Conclusion: CCM therapy appears to be safe and feasible. Proarrhythmic effects of this novel therapy seem unlikely. Preliminary data indicate that CCM gradually and significantly improves systolic performance, symptoms, and functional status. CCM therapy for 7 hours per day is associated with greater dispersion near the mean, emphasizing the need to individually tailor CCM delivery duration. The technique appears to be attractive as an additive treatment for severe HF. Controlled randomized studies are needed to validate this novel concept. (J Cardiovase Electrophysiol, Vol. 15, pp. 418-427, April 2004)

### **His Bundle Pacing for CRT**

- ♥ Early studies (Rosen, Narula) indicated that some LBBB originated proximally in fibers in the HB that were destined to become the LBB.
- Early attempts at chronic HB pacing (Furman, Karpawich) foiled by development of fibrosis.
- Small modern steroid-eluting screws have re-opened interest in HB pacing.





QRS:

NSR: LBBB

H-Paced: LBBB

LBBB

Normal



Fig. 6.

Diagrammatic sketches of the left-sided conduction system as observed in 49 hearts (from Demoulin, J. C., Thesis, in preparation).

Updated from: Demoulin JC and Kulbertus HE. Br Heart J 1972;34:807-14

### Cardiac Resynchronization Through Selective His Bundle Pacing in a Patient with the So-Called InfraHis Atrioventricular Block

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MORINA-VAZQUEZ, P., ET AL.: Cardiac Resynchronization Through Selective His Bundle Pacing in a Patient with the So-Called InfraHis Atrioventricular Block. We present a case of infraHis AV block in which selective His bundle pacing with His-ventricular conduction through the conduction system was accomplished. While further investigations are developed, this approach may be an alternative for cardiac resynchronization in cases of difficult coronary sinus access. (PACE 2005; 28:726–729)

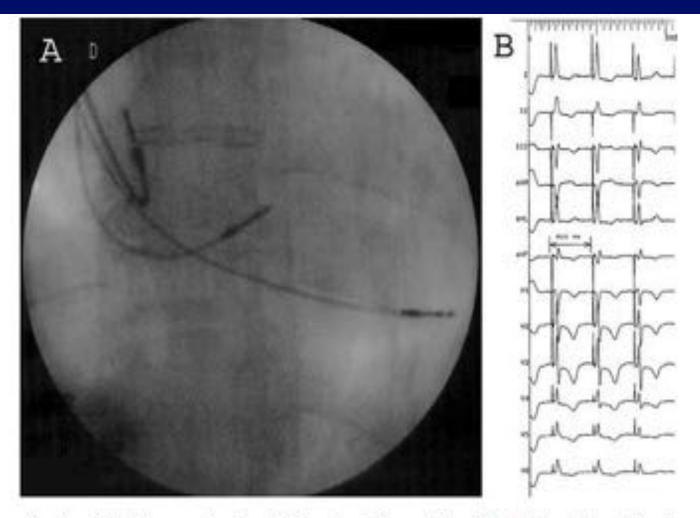


Figure 2. Panel A: PA x-ray showing the final position of the right atrial, right ventricular, and His bundle leads. Panel B: Continuous pacing at the final position of the His bundle lead catheter beside the reference temporary catheter, the third complex is a pure His bundle capture, while the other two QRS complex are minimally fused.

PACE, Vol. 28 July 2005

## Permanent His-bundle pacing is feasible, safe, and superior to right ventricular pacing in routine clinical practice © © ©



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**BACKGROUND** Right ventricular pacing (RVP) has been associated with heart failure and increased mortality. His-bundle pacing (HBP) is more physiological but requires a mapping catheter or a backup right ventricular lead and is technically challenging.

**OBJECTIVE** We sought to assess the feasibility, safety, and clinical outcomes of permanent HBP in an unselected population as compared to RVP.

METHODS All patients requiring pacemaker implantation routinely underwent attempt at permanent HBP using the Select Secure (model 3830) pacing lead in the year 2011 delivered through a fixed-shaped catheter (C315 HIS) at one hospital and RVP at the second hospital. Patients were followed from implantation, 2 weeks, 2 months, 1 year, and 2 years. Fluoroscopy time (FT), pacing threshold (PTh), complications, heart failure hospitalization, and mortality were compared.

RESULTS HBP was attempted in 94 consecutive patients, while 98 patients underwent RVP. HBP was successful in 75 patients (80%). FT was similar (12.7  $\pm$  8 minutes vs 10  $\pm$  14 minutes; median 9.1 vs 6.4 minutes; P = .14) and PTh was higher in the HBP group than in the RVP group (1.35  $\pm$  0.9 V vs 0.6  $\pm$  0.5 V at 0.5 ms; P < .001) and remained stable over a 2-year follow-up period. In patients with >40% ventricular pacing (>60% of patients), heart failure

hospitalization was significantly reduced in the HBP group than in the RVP group (2% vs 15%; P = .02). There was no difference in mortality between the 2 groups (13% in the HBP group vs 18% in the RVP group; P = .45).

CONCLUSION Permanent HBP without a mapping catheter or a backup right ventricular lead was successfully achieved in 80% of patients. PTh was higher and FT was comparable to those of the RVP group. Clinical outcomes were better in the HBP group than in the RVP group.

KEYWORDS His-bundle pacing: Para-Hislan pacing: Right ventricular pacing; Heart failure; Clinical outcomes; Safety; Feasibility

ABBREVIATIONS AF = atrial fibrillation; AV = atrioventricular; DHBP = direct His-bundle pacing; ECG = electrocardiogram/electrocardiographic; FT = fluoroscopy time; HB = His-bundle; HBP = His-bundle pacing; HF = heart failure; HFH = heart failure hospitalization; LV = left ventricular; LVEF = left ventricular ejection fraction; PHP = para-Hisian pacing; PTh = pacing threshold; RBBB = right bundle branch block; RVP = right ventricular pacing

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### HB pacing may give results similar to conventional CRT

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Heart Rhythm, Vol 12, No 2, February 2015

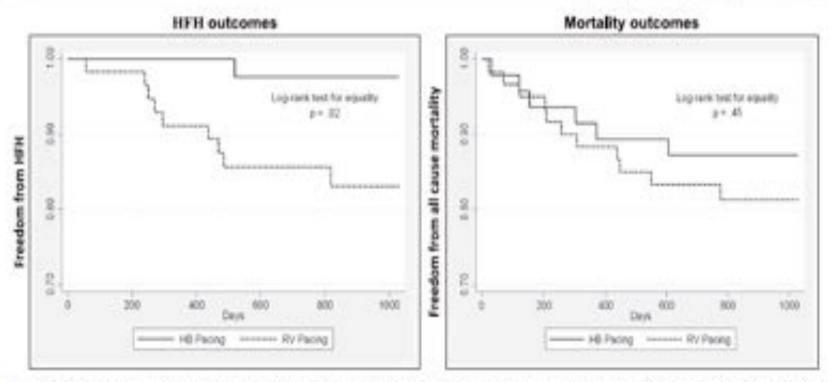


Figure 5 Kaplan-Meier survival curves depicting heart failure hospitalization (HFH) and all-cause mortality outcome differences in patients with >40% ventricular pacing, comparing His-bundle (HB) pacing with right ventricular (RV) pacing.

### **Conclusions**

- CRT is important for heart failure patients especially with LBBB.
- Conventional CRT (RV + LV) is sufficient for most.
- For others, quadripolar LV leads give option of LV MSP.
- Other entities also should be considered:
  - Endocardial and surgically placed LV leads, unconventional stimulation, His Bundle pacing.

### All Done!



### AF + CHF: AVJ ablation + CRT, or PVI?

Journal of the American College of Cardiology © 2006 by the American College of Cardiology Foundation Published by Elsevier Inc. Vol. 48, No. 4, 2006 888N 0735-1097/06/832.00 doi:10.1016/j.incc.2006.03.056

Four-Year Efficacy of Cardiac Resynchronization Therapy on Exercise Tolerance and Disease Progression

The Importance of Performing Atrioventricular Junction Ablation in Patients With Atrial Fibrillation

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Milan and Pavia, Italy, Magdeburg, Germany, and Lugano, Switzerland

### AF + CHF: AVJ ablation + CRT, or PVI?

#### Pulmonary-Vein Isolation for Atrial Fibrillation in Patients with Heart Failure

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Gemma Pelargonio, M.D., Johannes Brachmann, M.D.,
Volker Schibgilla, M.D., Aldo Bonso, M.D., Michela Casella, M.D.,
Antonio Raviele, M.D., Michel Haissaguerre, M.D., and Andrea Natale, M.D.,
for the PABA-CHF Investigators\*

#### CONCLUSIONS

Pulmonary-vein isolation was superior to atrioventricular-node ablation with biventricular pacing in patients with heart failure who had drug-refractory atrial fibrillation. (ClinicalTrials.gov number, NCT00599976.)

### AV Node Ablation in AF + CRT: Choose patients carefully: There's no turning back!

