

How to identify optimal ECG patterns for CRT

인하대 병원

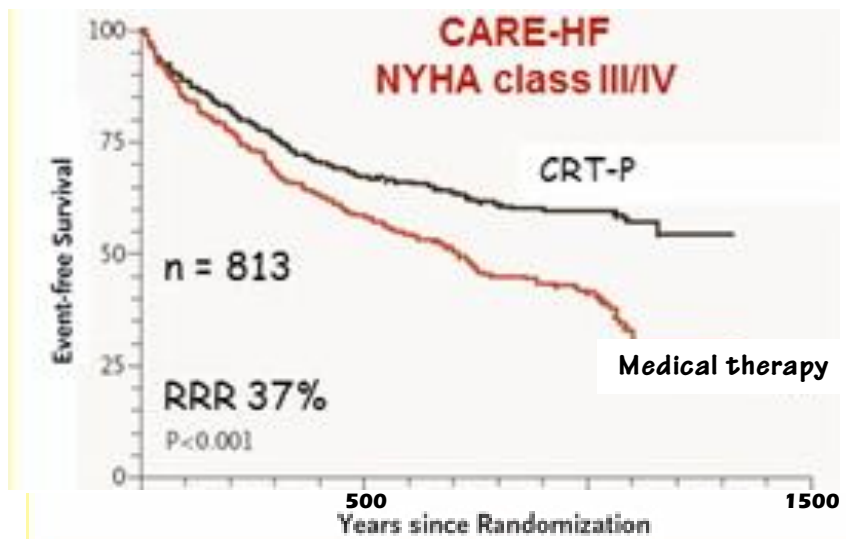
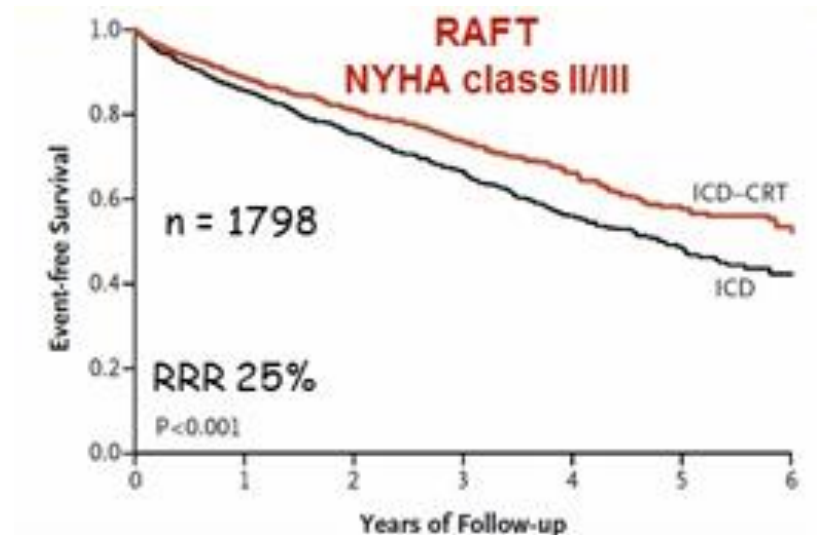
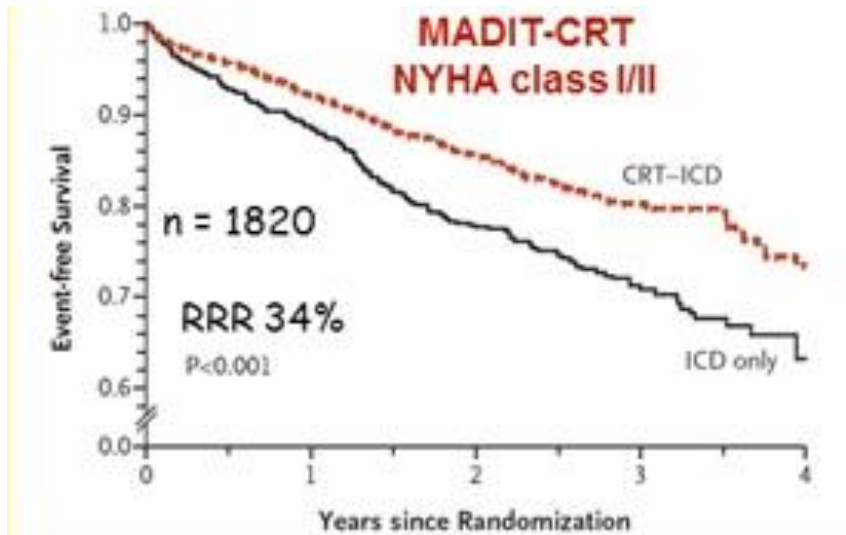
심장내과

김대혁

CRT

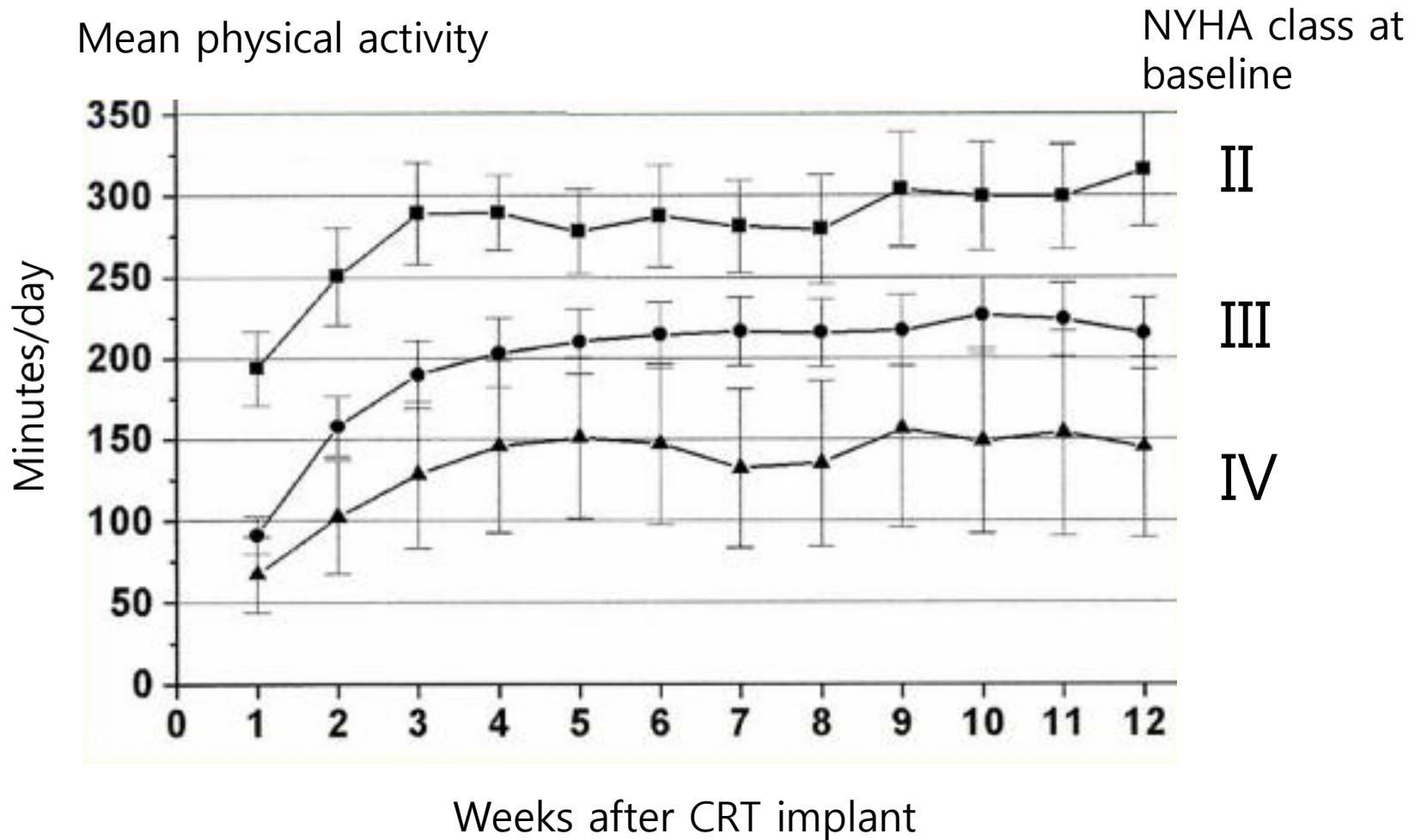
- **Interventricular dyssynchrony** : delayed activation of one ventricle with respect to the other
- **Intraventricular dyssynchrony** : late activation of the lateral region of the LV chamber vs activation of the septum.
- **CRT** : attempt to normalize the timing of activation of the left and right ventricle or lateral wall and septum----> improve the consequence of this impaired activation.

Effect of CRT in major clinical trials



N Engl J Med 2009; 361:1329-38
N Engl J Med 2010;363:2385-95
N Engl J Med 2005;352 :1539-49

Effect of CRT are evident soon after implant



Cardiac resynchronization therapy (CRT): Clinical trials, guidelines, and target populations

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Over the last 10 years, several large, well-designed clinical trials have firmly established the role of cardiac resynchronization therapy (CRT) as a recommended treatment strategy for moderate-to-severe heart failure (HF). A review of the relevant results from the MUSTIC, MIRACLE, CONAK-CD, and MIRACLE ICD trials reveals that in patients with New York Heart Association (NYHA) class III–IV HF, CRT produces consistent improvements in quality of life, functional status, and exercise capacity while also providing strong evidence for reverse remodeling and diminished functional mitral regurgitation, resulting in reductions in both HF hospitalizations and all-cause morbidity and mortality. In patients with earlier NYHA class I–II HF, the benefit of CRT has been more controversial. The principal ongoing challenges addressed in this article include the substantial 30% of patients who receive a CRT device but fail to respond, the wide variations in how to define “response” vs “nonresponse,” and how to identify patients who will benefit from CRT, especially narrow QRS (<120 ms), those with right bundle branch block, and those with mild-to-moderate

(NYHA class I–II) HF. An important result of this uncertainty is the lack of a good sense of the optimal rate of CRT implantation, making consideration of the data reviewed in this article crucial for identifying important gaps of knowledge and mechanisms of action that need to be studied in the near future.

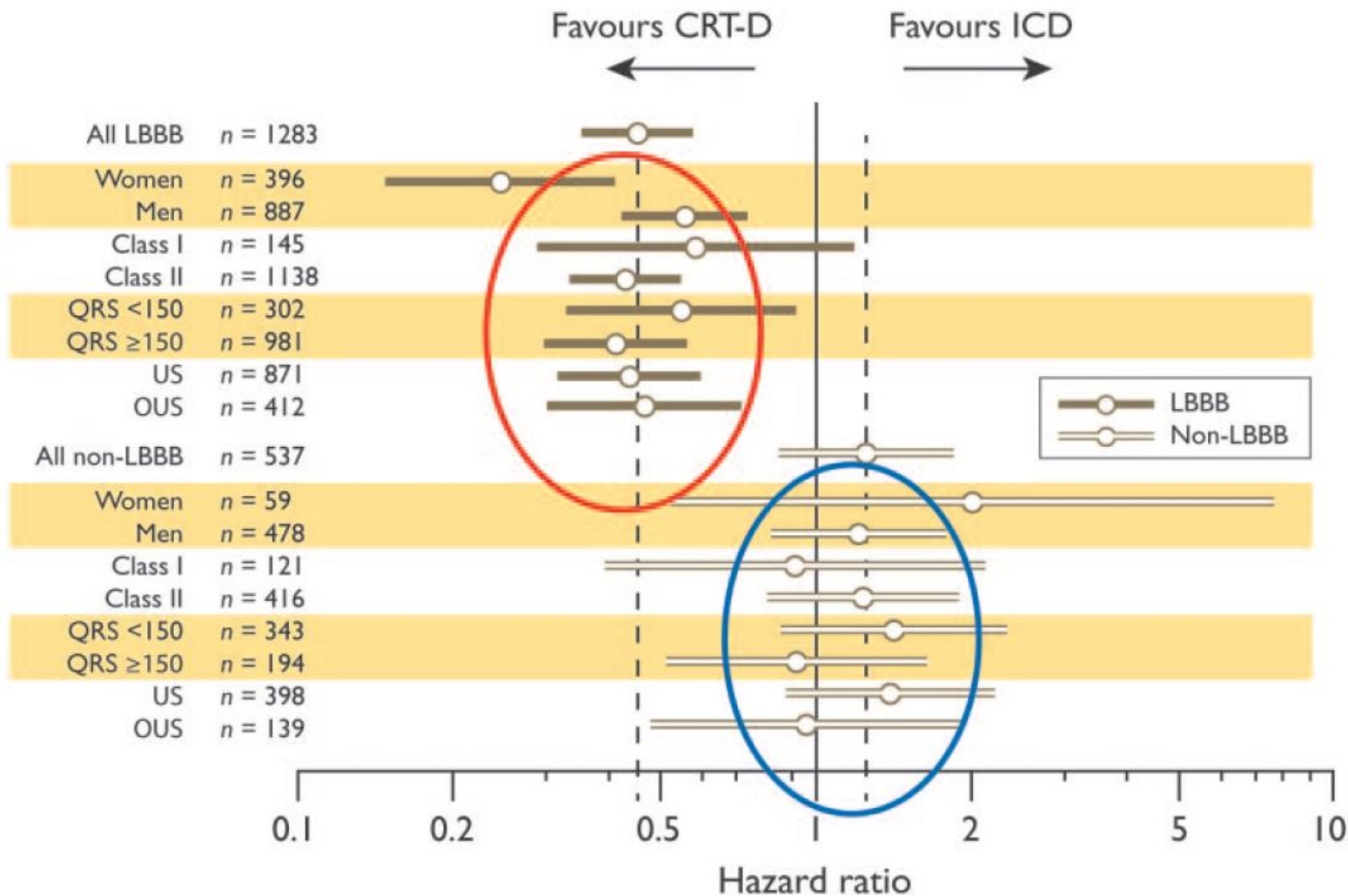
KEYWORDS Arrhythmia; Biventricular pacing; Cardiac resynchronization therapy (CRT); Dyssynchrony; Heart failure; Narrow QRS; Nonresponder

ABBREVIATIONS ACE = angiotensin-converting enzyme; BiV = biventricular; CRT = cardiac resynchronization therapy; EF = ejection fraction; ESC = European Society of Cardiology; HF = heart failure; ICD = implantable cardioverter-defibrillator; LBBB = left bundle branch block; LV = left ventricular; NYHA = New York Heart Association; RBBB = right bundle branch block

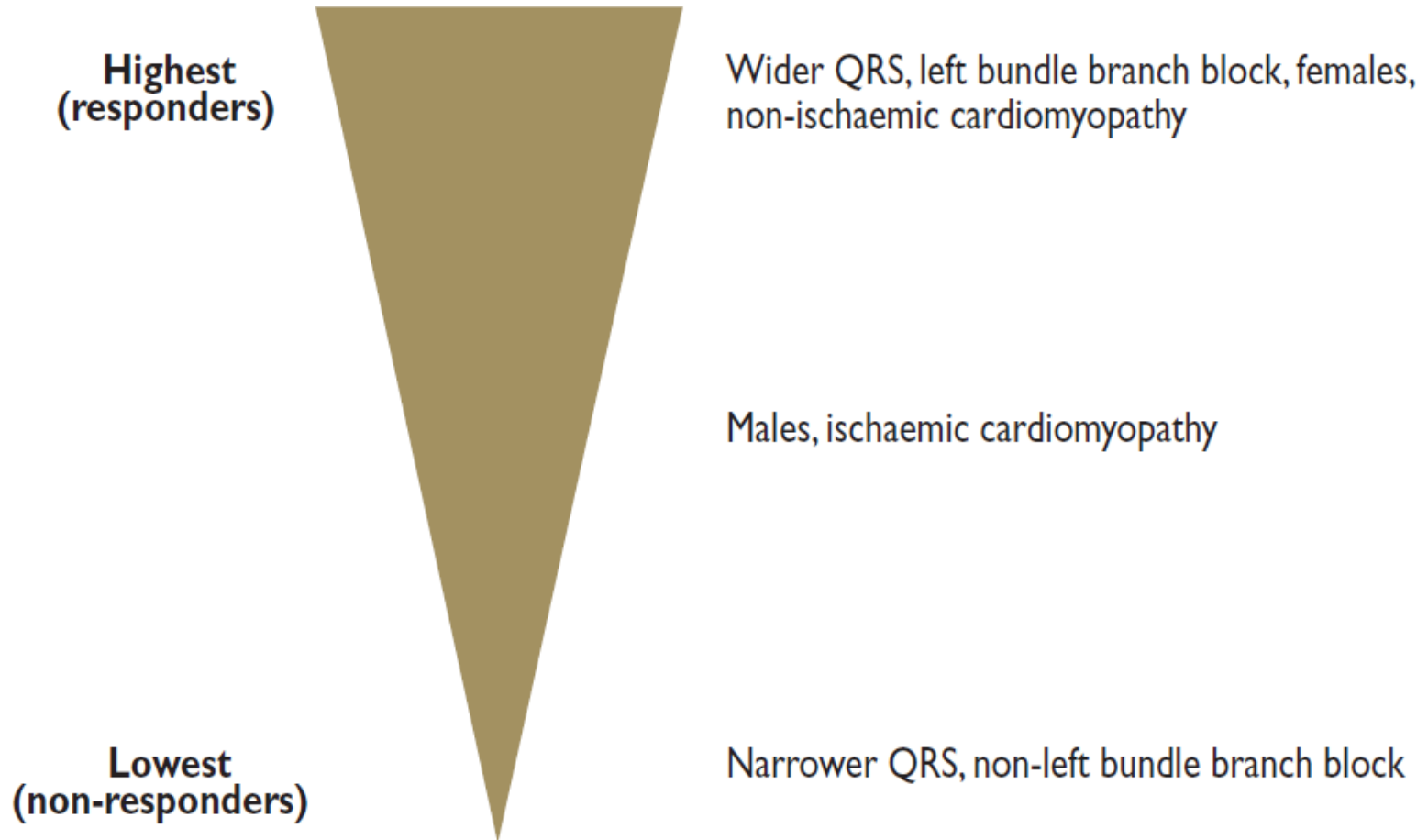
(Heart Rhythm 2012;9:S3–S13) © 2012 Heart Rhythm Society. Published by Elsevier Inc. All rights reserved.

**include the substantial 30% of patients who receive a CRT device
But failed to respond,**

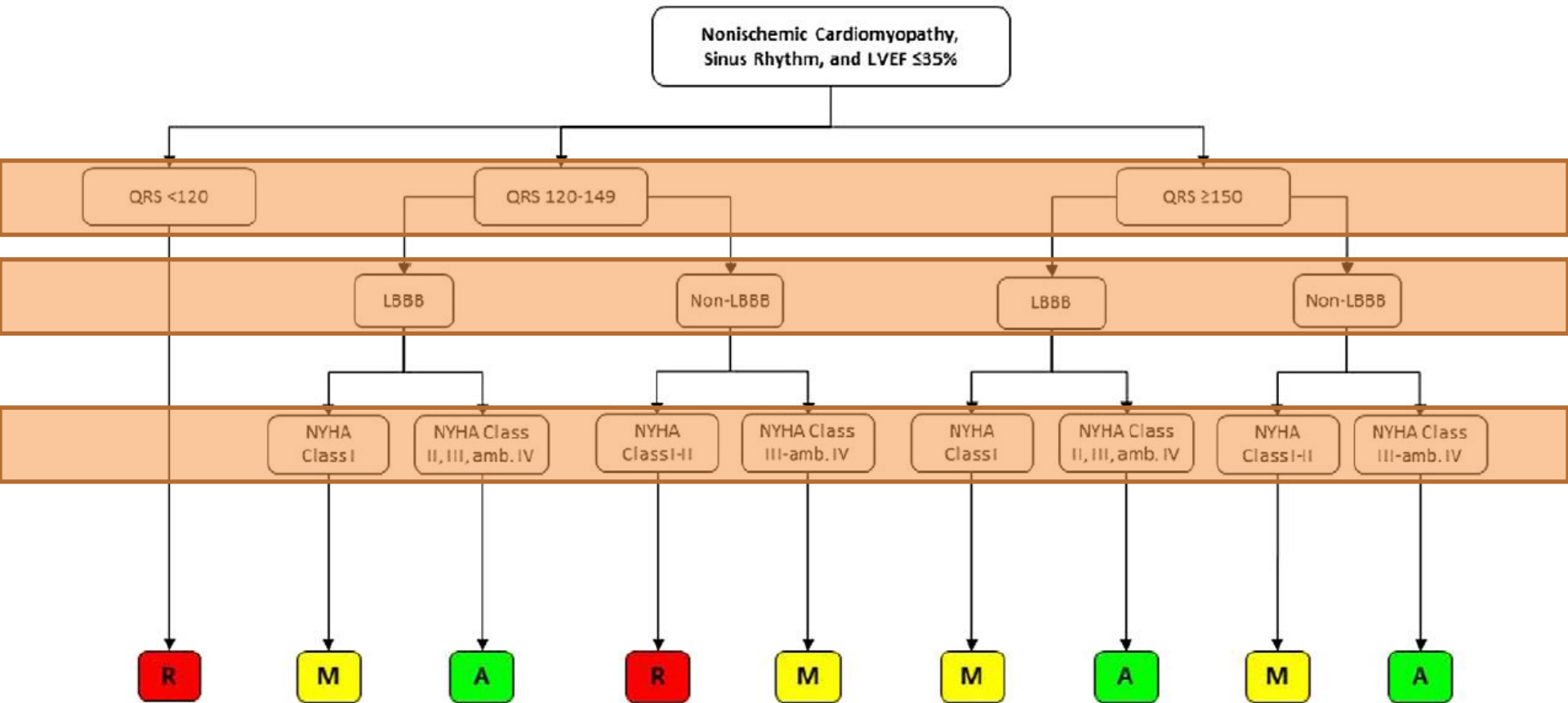
Clinical factors influencing the likelihood to response to CRT



Magnitude of benefit from CRT

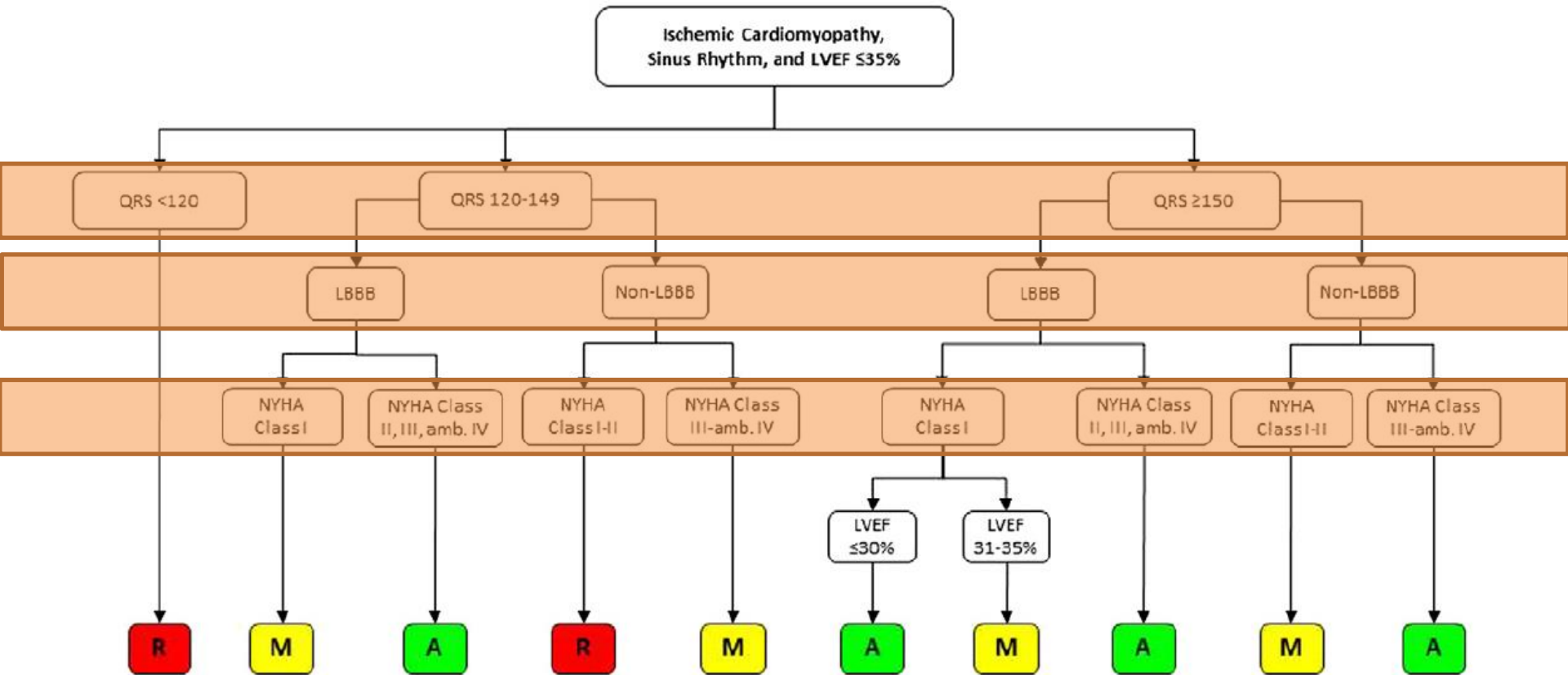


Appropriate use criteria for CRT



A = Appropriate; amb = ambulatory; CRT = cardiac resynchronization therapy; LBBB = left bundle branch block; LVEF = left ventricular ejection fraction; M = May Be Appropriate; NYHA = New York Heart Association; R = Rarely Appropriate.

Appropriate use criteria for CRT



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Cause that contribute to the degree of response to CRT

- Type of underlying heart ds(ischemic vs non-ischemic)
- Type of ventricular conduction disturbance(LBBB vs non-LBBB)
- Suboptimal pharmacological medical Tx.
- Presence of (supra)ventricular arrhythmias
- Position of LV lead
- Timing of RV and LV pacing/Atrioventricular interval

Optimization of CRT

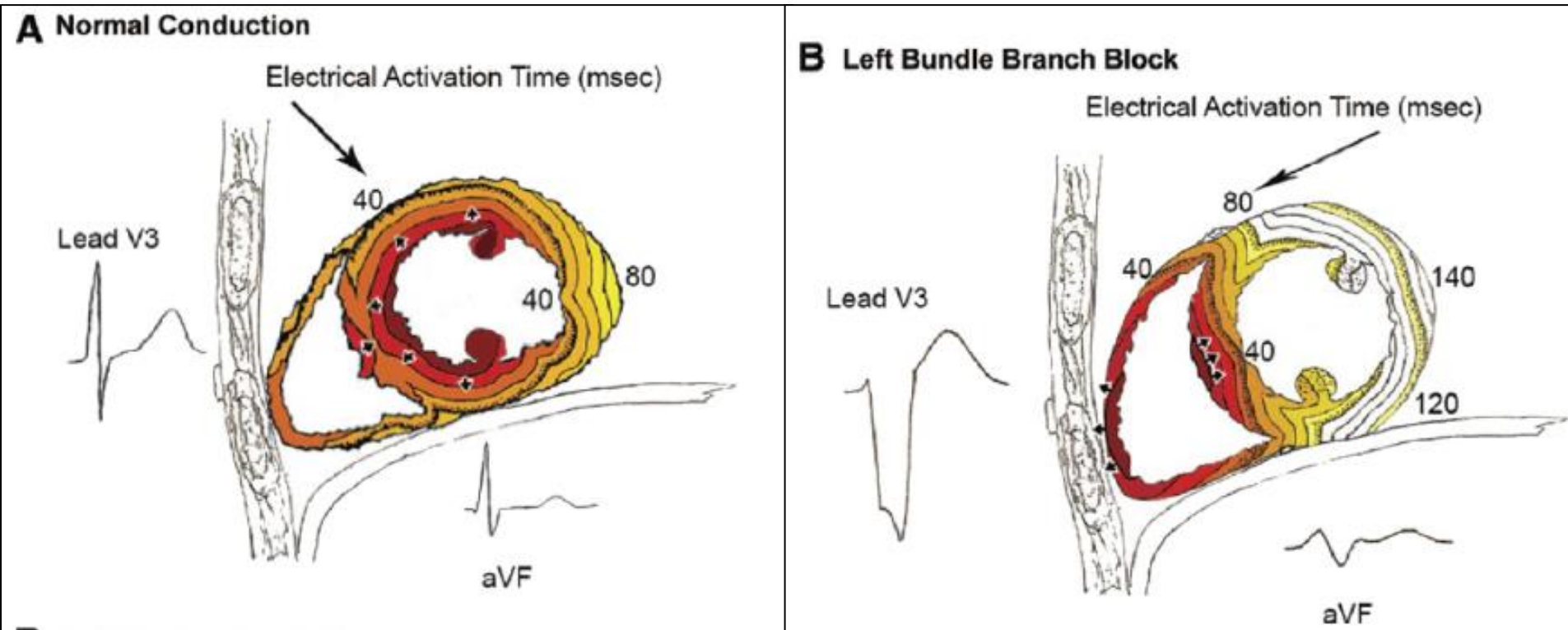
- How to achieve biventricular pacing as close to 100% as possible
- How to select the best LV lead position
- How to program the AV interval in order to achieve the max. contribution of LA contraction to LV filling
- How to eliminate the residual LV dyssynchrony through device VV interval optimization.

Strategy for using the standard 12-lead ECG for evaluation and improving application of CRT

- To assess the electrical substrate of a patient: presence of a **typical LBBB pattern** and **absence of scar**
- To assess the **QRS pattern** during LV pacing in order to judge its **proper position**
- For optimization of the programmed **AV delay** and **VV interval**

True LBBB

Normal Conduction vs LBBB



- During LBBB development
 - 1) QRS duration prolongs by more than 40ms
 - 2) Initial electrical force(QRS morphology) must change because of the different activation of septum

Criteria of LBBB

ECG parameters for complete LBBB according to guidelines of European Society of Cardiology (ESC) [11], American Heart Association (AHA)/American College of Cardiology Foundation (ACCF)/Heart Rhythm Society (HRS) [12], Strauss et al. [13], MADIT-CRT [14] and REVERSE [15] clinical CRT trials.

ECG parameter for complete LBBB	ESC	AHA	Strauss	MADIT	REVERSE
QRS duration (ms) \geq	120	120	♀130 ♂140	130	120
QS or rS in V ₁	Yes	Yes	Yes	Yes	Yes
Positive T in V ₁	Yes	No	No	No	No
Normal ID R in V ₁ -V ₃	No	Yes	No	No	No
ID R in V ₅ \geq 60 ms	No	Yes	No	No	No
ID R in V ₆ \geq 60 ms	Yes	Yes	No	No	No
ID R in I \geq 60 ms	Yes	No	No	No	No
Notch-/slurred R in I, aVL and V ₅ -V ₆	No	Yes	No	No	No
Mid-QRS notch/slurring in \geq 2 leads of V ₁ -V ₂ , V ₅ -V ₆ , I, aVL	No	No	Yes	No	No
RS pattern allowed in V ₅ -V ₆	No	Yes	Yes	Yes	Yes
Absent q in V ₅ -V ₆	No	Yes	No	Yes	Yes
Absent q in I	No	Yes	No	No	No
QS with positive T in aVR	Yes	No	No	No	No
Usually discordant T	Yes	Yes	No	No	No

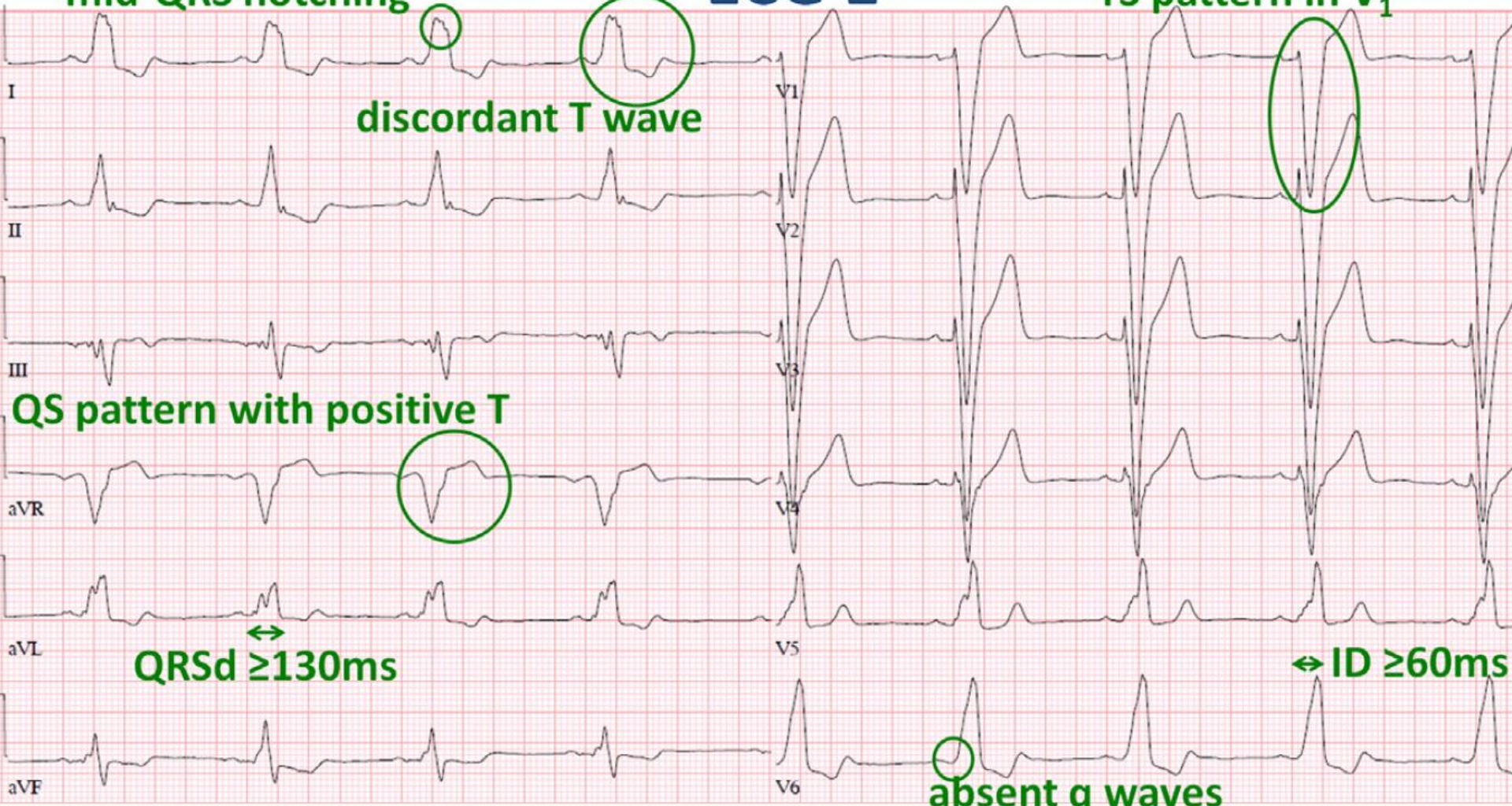
ID : intrinsicoid deflection defined as time from start QRS to R peak

ECG 1

mid-QRS notching

discordant T wave

rS pattern in V₁



QRSd ≥ 130 ms

ID ≥ 60 ms

absent q waves

ECG 2

mid-QRS slurring

rS pattern in V₁

no discordant T wave

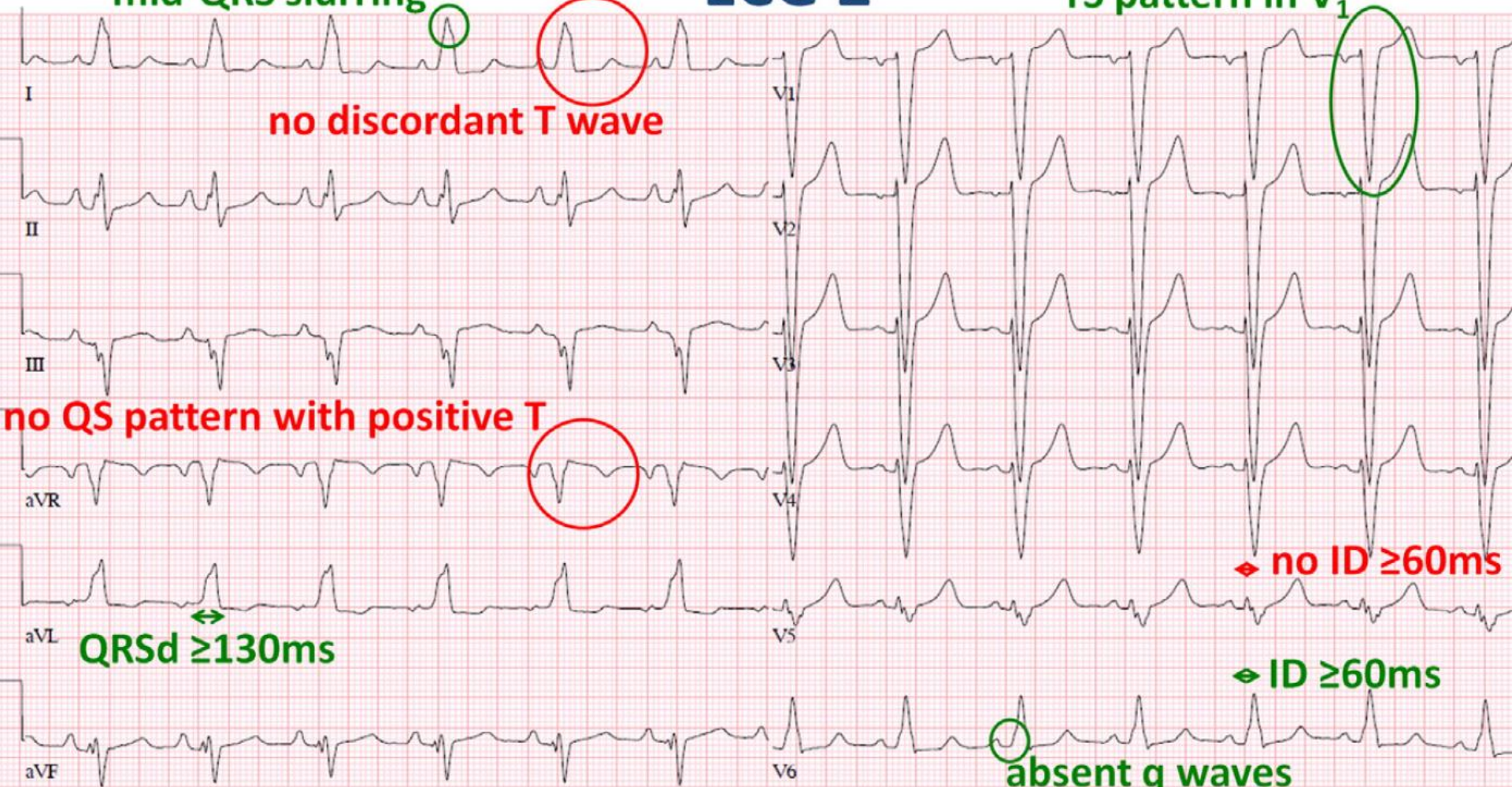
no QS pattern with positive T

QRSd ≥ 130 ms

no ID ≥ 60 ms

ID ≥ 60 ms

absent q waves



ECG 3

no mid-QRS notching

no discordant T wave

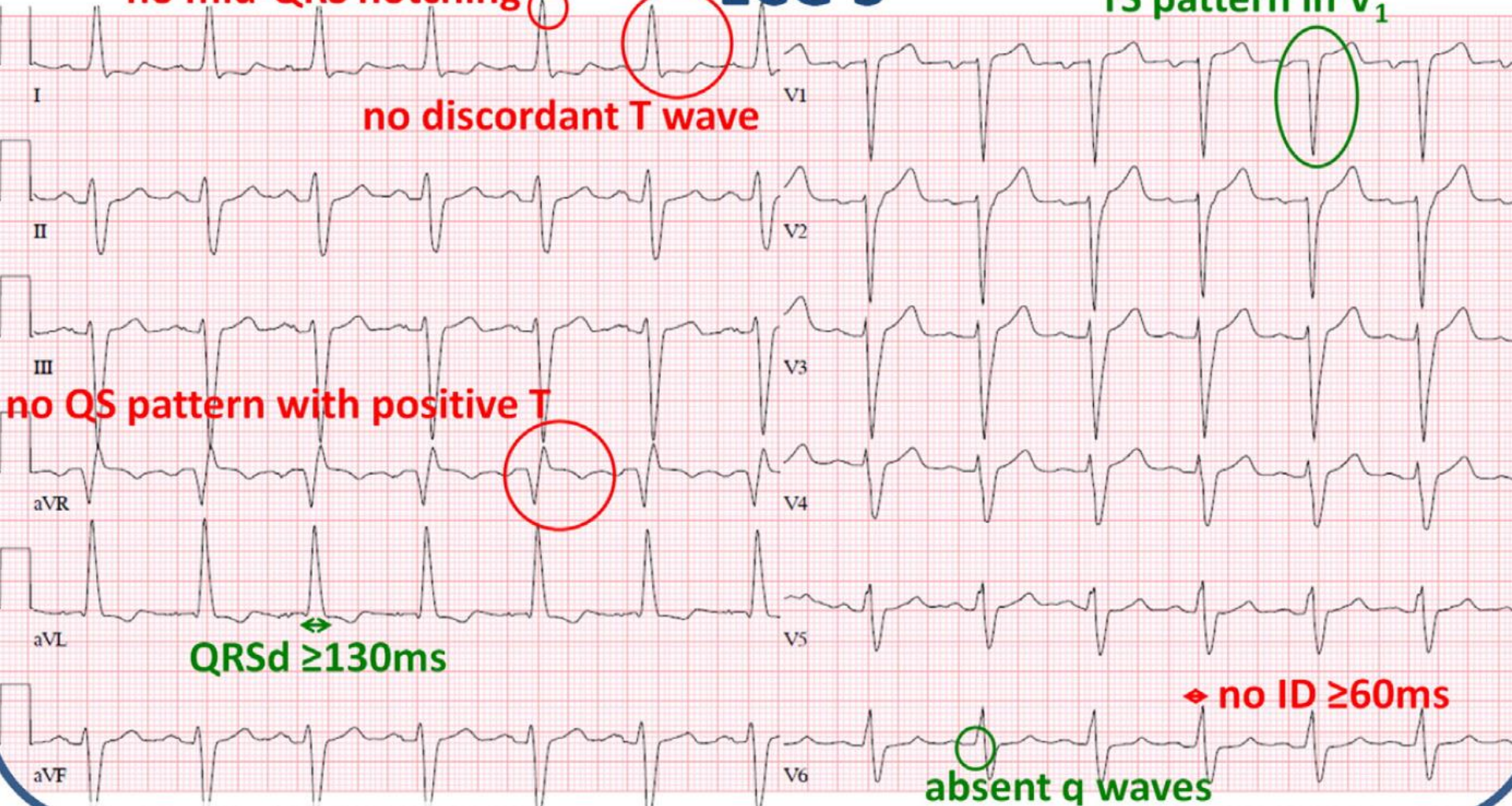
rS pattern in V₁

no QS pattern with positive T

QRSd ≥ 130 ms

no ID ≥ 60 ms

absent q waves



3D contact and Noncontact mapping of patients with LBBB by conventional ECG criteria

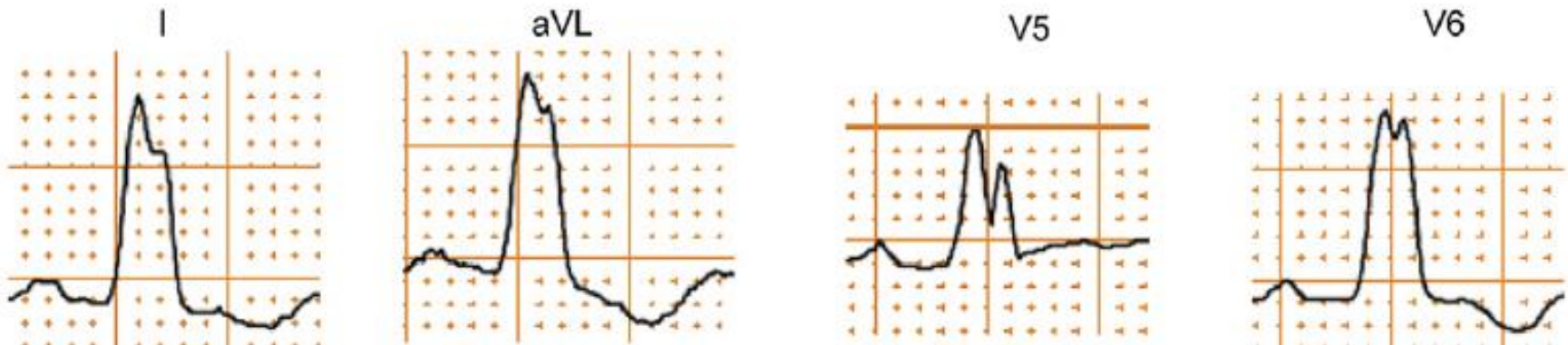
Location of LV Breakthrough Site	Site of Line of Block				Pathogenesis: DCM/CAD	QRS Duration		Time to LV Breakthrough		Time of Total LV Activation		Distance From LV Breakthrough Site to Line of Block NCM, mm
	None	Ant	Lat	Inf		Automatic, ms	Maximum, ms	NCM, ms	CM, ms	NCM, ms	CM, ms	
Anterior (n=2)			2		2/0	100±9	136±11	11±3	5±0	86±20	94±32	98±6
Septal (n=22)												
Basal (n=4)		1	2	1	2/2	149±20	168±24	16±25	13±18†	107±30	114±28	92±35†
Middle (n=4)	1	2		1	3/1	149±35	167±34	38±25	25±36	82±20	99±16	63±15
Apical (n=14)		9	4	1	10/4	168±17*	195±29*	59±25*	62±22*	101±20	106±17	63±10*
<i>P</i>					0.681	0.03	0.035	0.010	0.001	0.346	0.648	0.009

Ant indicates anterior; Lat, lateral; Inf, inferior; CAD, coronary artery disease; DCM, idiopathic dilated cardiomyopathy; CM, contact mapping; and NCM, noncontact mapping.

*Statistical difference between anterior and septal-apical at the level of significance $P < 0.05$.

†Statistical difference between septal-basal and septal-apical at the level of significance $P < 0.05$.

- **1/3rd of patients** did not have significant delay between the start of activation of the RV endocardium and the start of activation of the LV endocardium



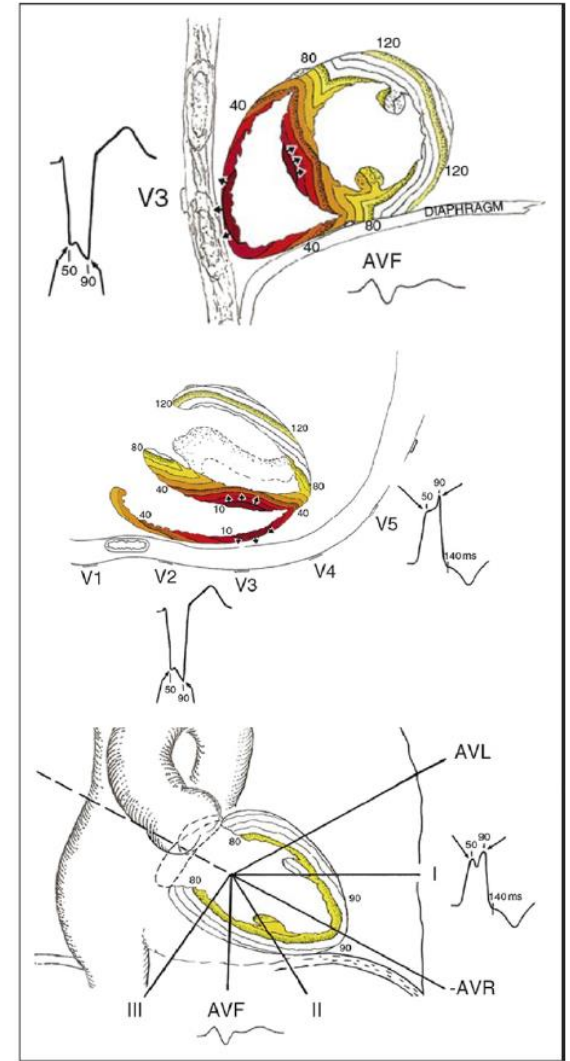
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Usually discordant T	Yes	Yes	No	No	No

ID : intrinsicoid deflection defined as time from start QRS to R peak

Notch at QRS

- **First notch** : time when the electrical depolarization wave front reaches the endocardium of the LV(after proceeding through the septum)
- **Second notch** : occur when the depolarization wave front begins to reach the epicardium of the posterolateral wall.
- These notches are best seen in lead **I, aVL, V₁, V₂, V₅ and V₆**

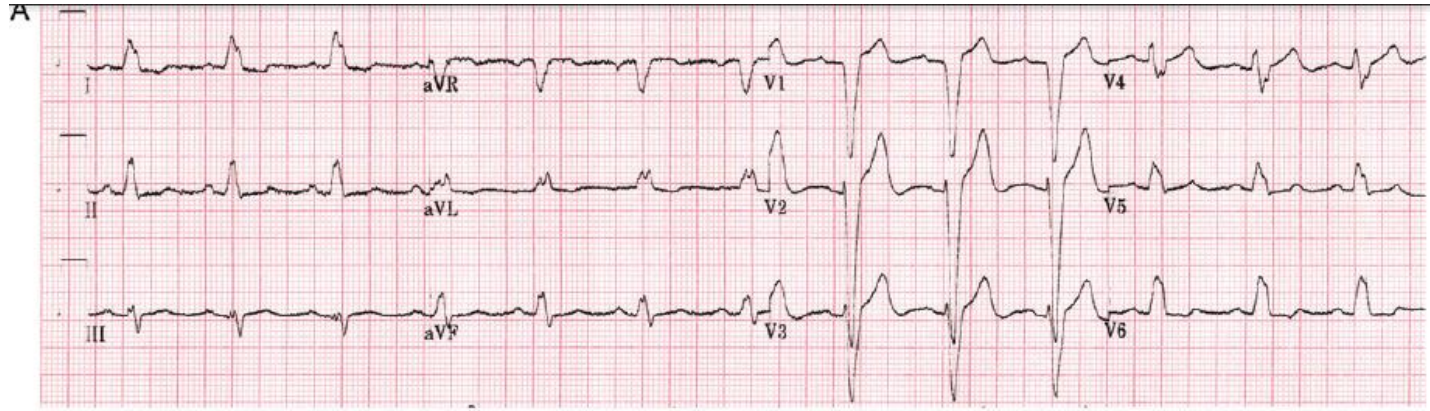


Diagnostic performance of different LBBB definition for prediction of CRT response (>15% reduction in end-systolic LV volume)

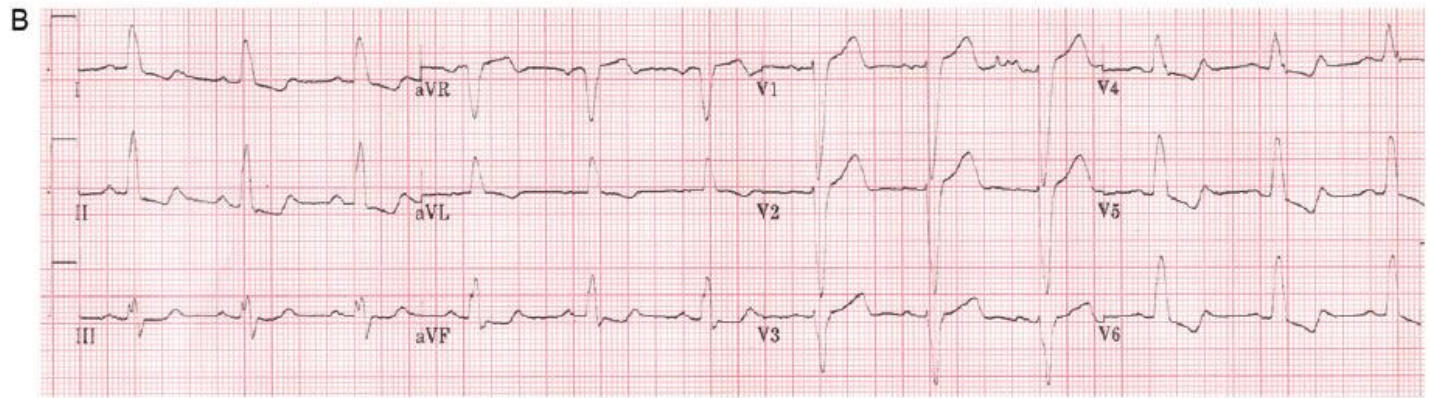
LBBB definition	OR	95% CI	<i>P</i> -value	Sensitivity	Specificity
ESC guideline	3.225	1.285–8.095	0.011	64%	64%
AHA/ACCF/HRS guideline	3.700	1.386–9.871	0.007	55%	75%
Strauss et al.	11.813	3.359–41.544	<0.001	94%	43%
MADIT-CRT	3.556	0.741–17.069	0.095	96%	14%
REVERSE	3.900	0.625–24.746	0.124	97%	11%

Presented are odds ratios (OR) with 95% confidence interval (CI) and, accordingly, *P*-values, sensitivity and specificity.

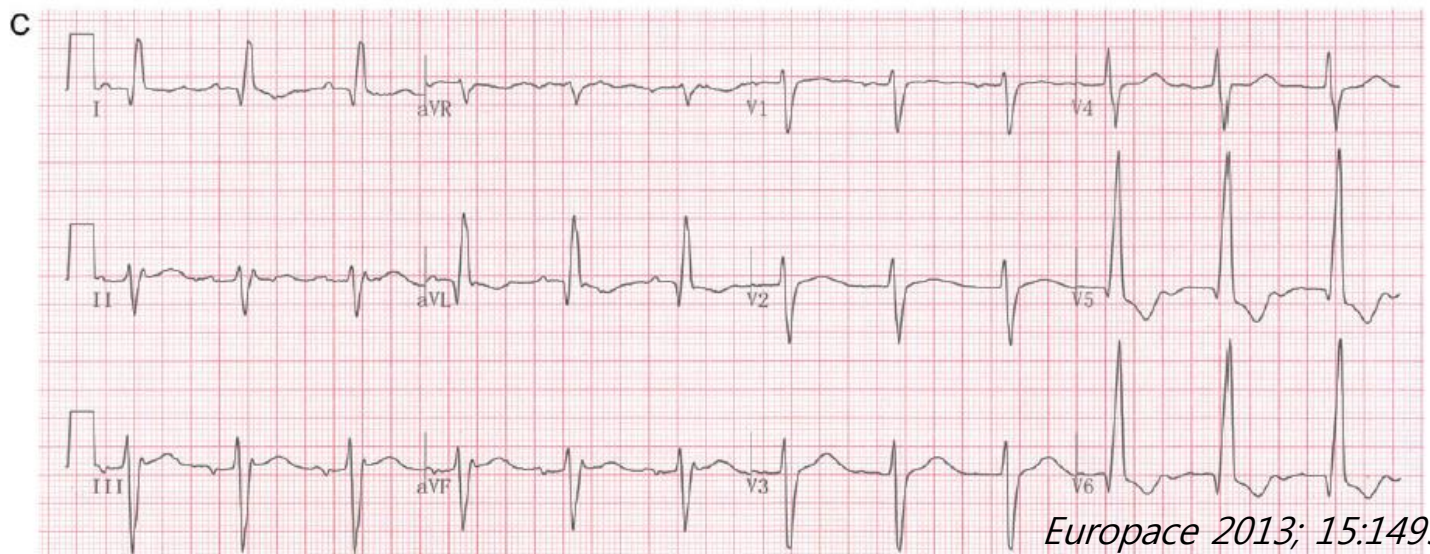
True LBBB



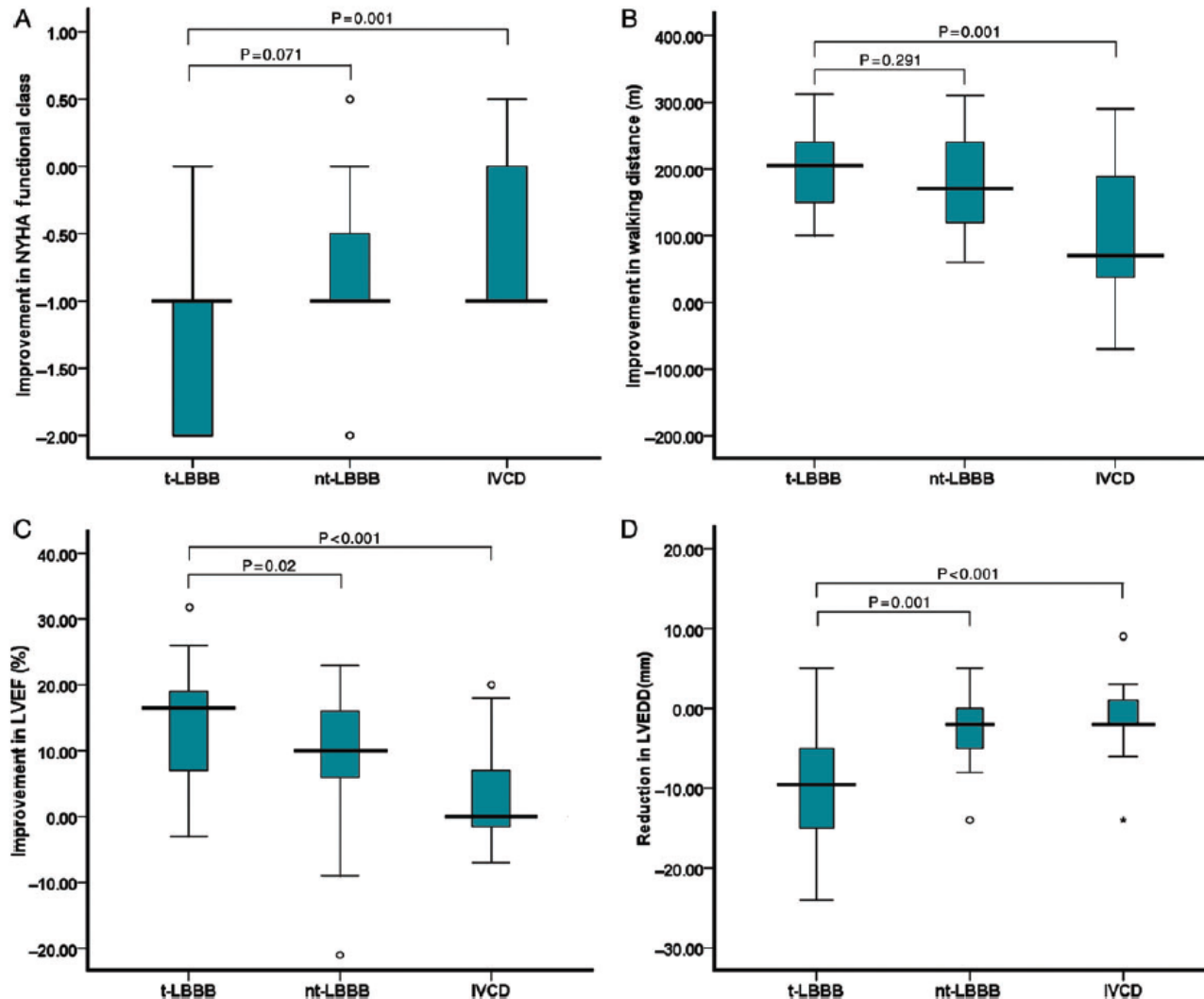
Non-true LBBB



IVCD



Clinical and Echocardiographic improvement 6 month after CRT



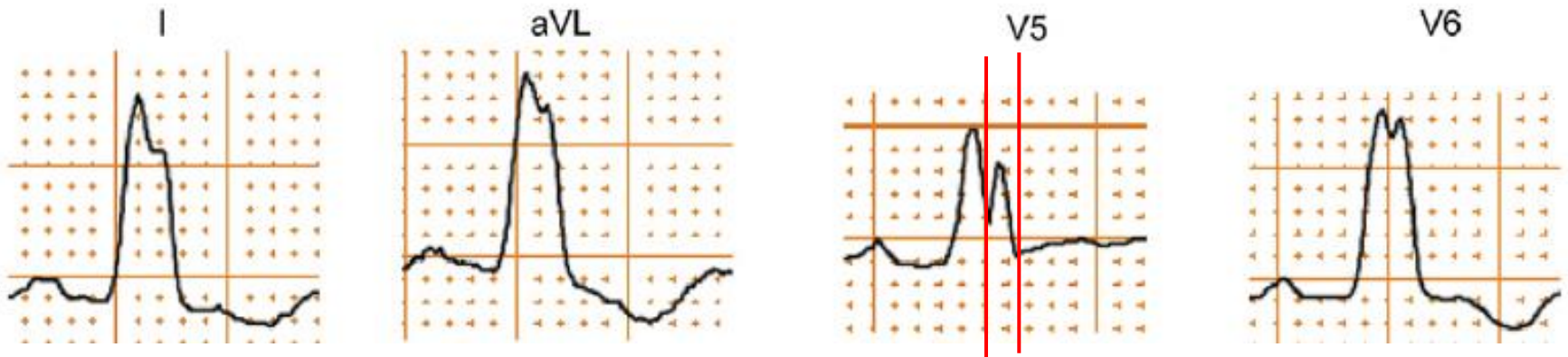
Difference in relevant characteristics of super-responder, responder, and non-responder

Variable	Super, <i>n</i> = 10	Responders, <i>n</i> = 31	Non-responders, <i>n</i> = 17
Gender (male)	7 (70.0%)	27 (87.1%)	16 (94.1%)
Age (years)	56.8 ± 13.3	61.0 ± 9.8	59.0 ± 14.1
HF duration (months)	21.1 ± 28.4	40.9 ± 32.9	56.5 ± 53.0
Non-ischaemic aetiology (<i>n</i>)	10 (100.0%)	23 (74.2%)	13 (76.5%)
NYHA class	2.9 ± 0.6	3.2 ± 0.5	3.5 ± 0.5
QRS duration pre-CRT (ms)	165.0 ± 17.2	161.0 ± 19.0	157.9 ± 15.3
QRS duration post-CRT (ms)	127.8 ± 29.1	132.5 ± 20.9	147.4 ± 22.9
QRS shortening after CRT (<i>n</i>)	10 (100.0%)	27 (79.4%)	12 (70.6%)
IVCD	1 (10%)	6 (22.6%)	12 (70.6%)
t-LBBB (<i>n</i> =22)	8 (80.0%)	14 (45.2%)	0 (0%)
LVEDD (mm)	68.2 ± 8.2	76.7 ± 10.4	76.3 ± 12.0
LVEF (%)	29.0 ± 5.3	26.5 ± 4.2	27.5 ± 7.8
MR (grade)	0.9 ± 0.6	1.7 ± 0.8	1.7 ± 0.8

Analysis of Ventricular Activation using ECG

LV activation time

- RVAT(RV activation time) :
- LVAT(LV activation time) :

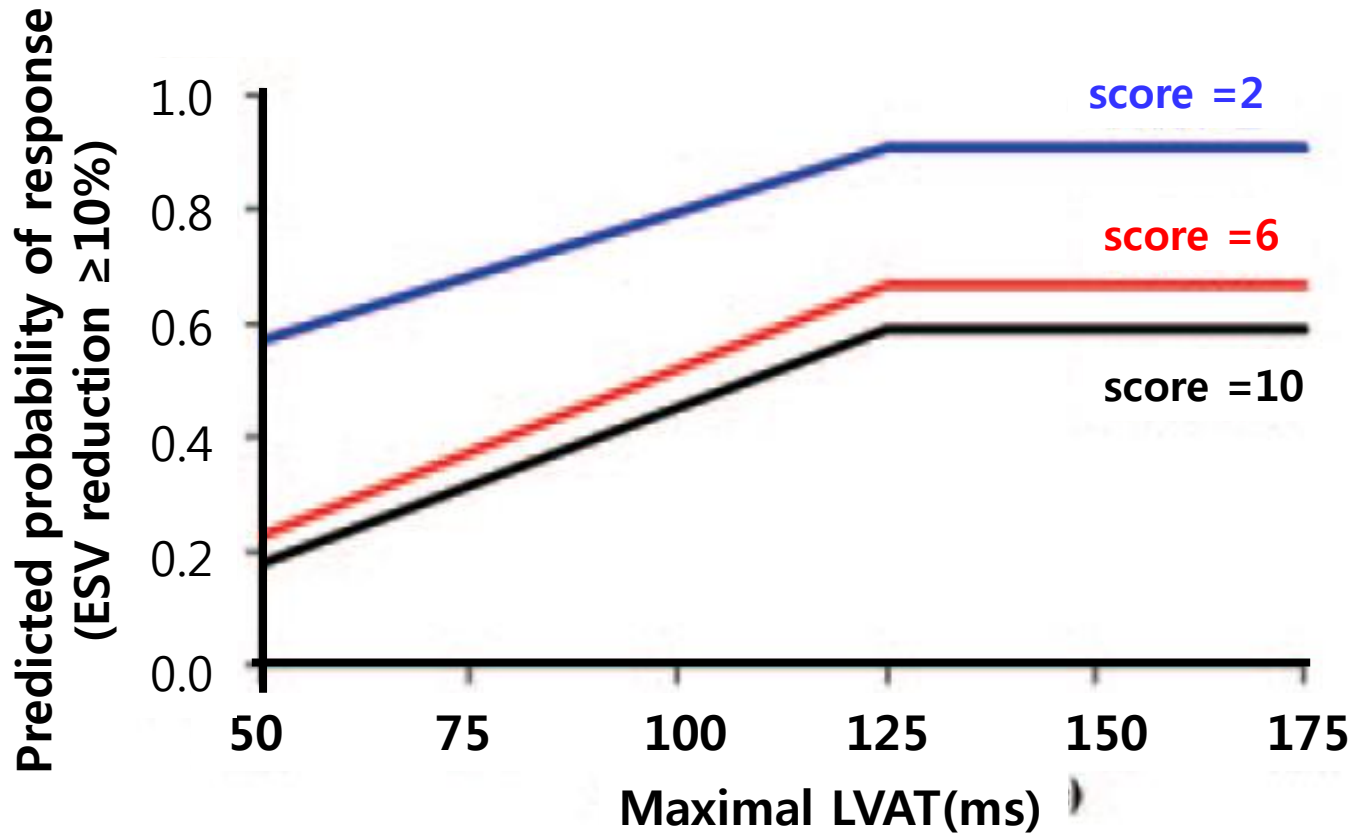


QRS Score for LV Scar

Points	Lead	Criterion
Inferior		
1	II	Q \geq 30 ms
2		Q \geq 40 ms
3	aVF	Q \geq 30 ms
4		Q \geq 40 ms
5		Q \geq 50 ms
6		Q/R \geq 0.5
7		Q/R \geq 1
Anterior		
1	V ₁	Any Q
2		S \geq 1.8 mV
3	V ₂	Any Q or R \leq 0.1 mV or R \leq 10 ms or RV ₂ \leq RV ₁
4	V ₃	Any Q or R \leq 0.2 mV or R \leq 20 ms
5	V ₄	Q \geq 20 ms
6		Q/R \geq 1 or R \leq 0.7 mV or R/S \leq 1 or notched R
7		Q/R \geq 2 or R/S \leq 0.5
Anterolateral		
1	I	Q \geq 30 ms
2		Q/R \geq 1 or R \leq 0.2
3	aVL	Q \geq 30 ms
4		Q/R \geq 1
Apical		
1	V ₅	Q \geq 30 ms
2		Q/R \geq 0.5 or R \leq 0.7 mV or R/S \leq 2 or notched R
3		Q/R \geq 1 or R/S \leq 1
4	V ₆	Q \geq 30 ms
5		Q/R \geq 1/3 or R \leq 0.6 mV or R/S \leq 3 or notched R
6		Q/R \geq 1 or R/S \leq 1
Posterolateral		
1	V ₁	R/S \geq 1
2		R \geq 0.6 mV or R \geq 40 ms
3		R \geq 1 mV or R \geq 50 ms
4		S \leq 0.3 mV
5	V ₂	R \geq 1.5 mV or R \geq 50 ms
6		R \geq 2 mV or R \geq 60 ms
7		R/S \geq 1.5
8		S \leq 0.4 mV

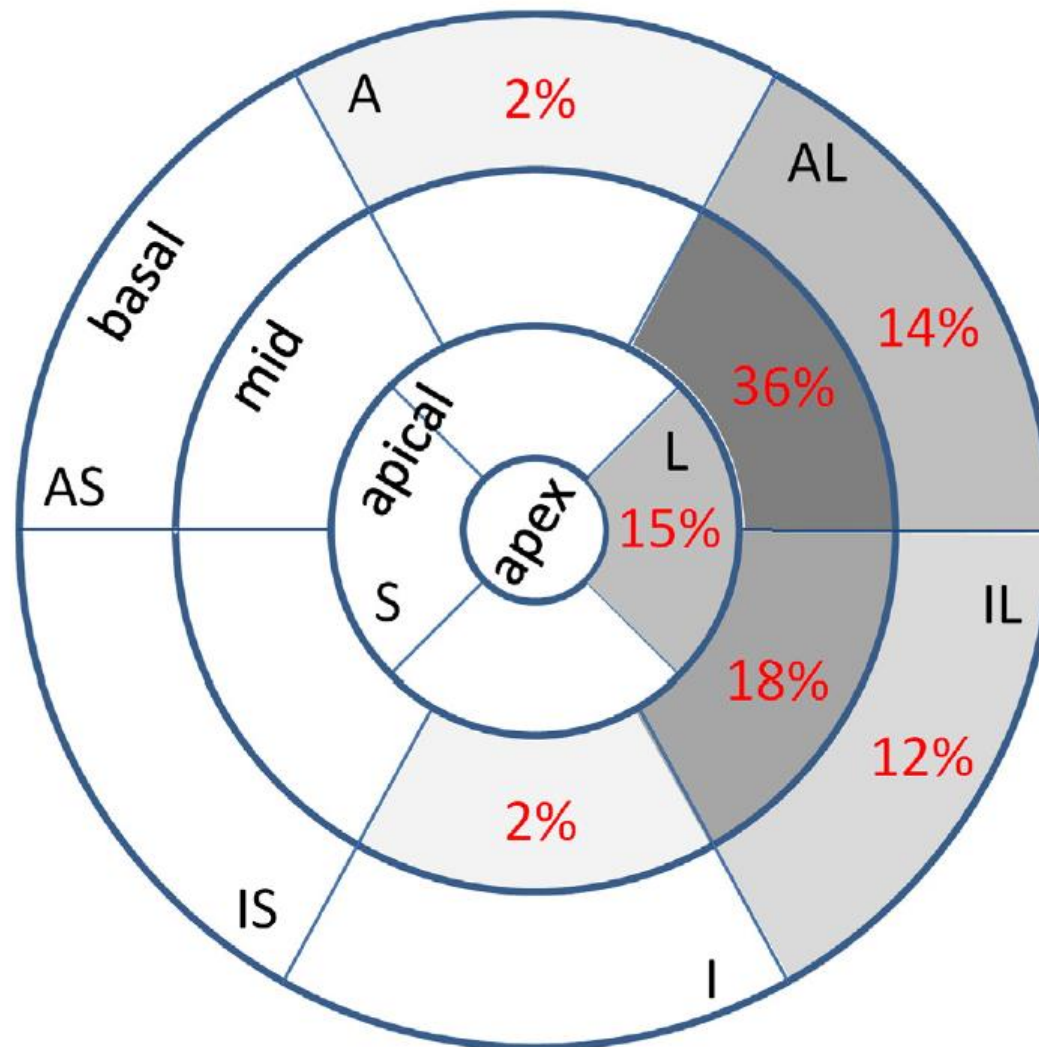
One point is awarded for each criterion met. Notched R indicates a notch that begins within the 1st 40 ms; Q = Q wave; Q/R = ratio of Q wave to R wave; R = R wave; R/S = ratio of R wave to S wave; RV₂ < RV₁ = R wave in lead V2 less than or equal to R wave in lead V1; S = S wave.

Probability of reverse remodeling by baseline QRS score and $LVAT_{max}$

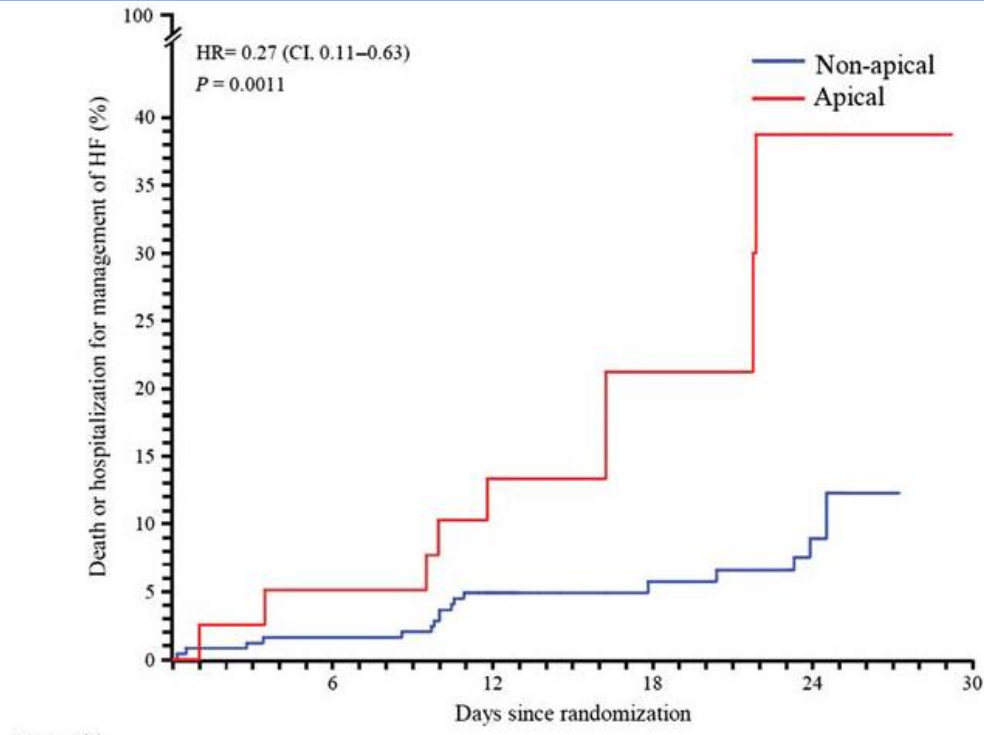
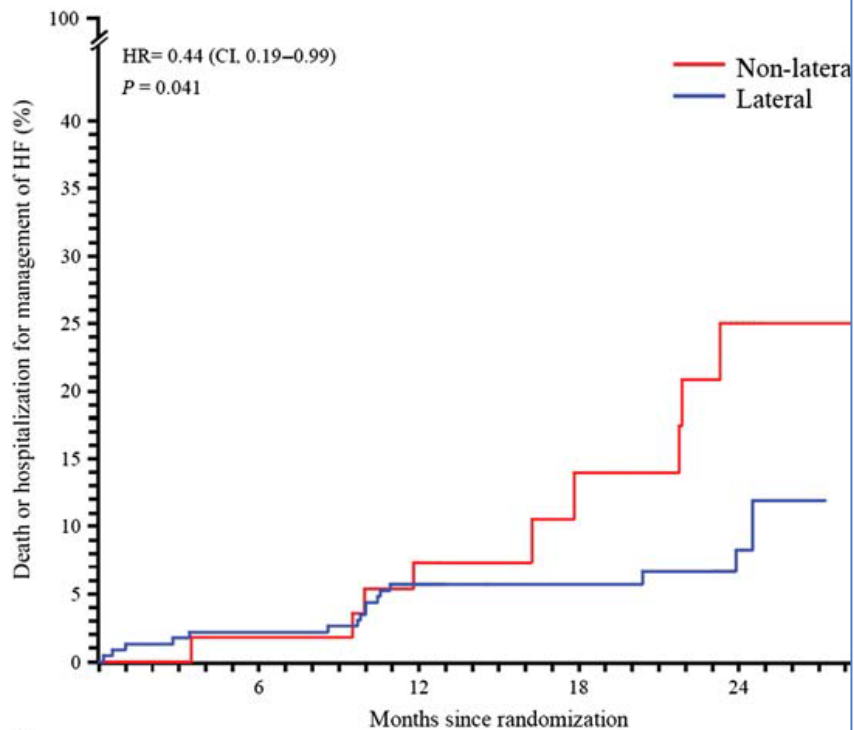


Concept of identification of the LV
Lead position by LV-paced QRS
morphology

Distribution of LV lead positions in 98 consecutive CRT patient by AHA standardized 17-segment Model

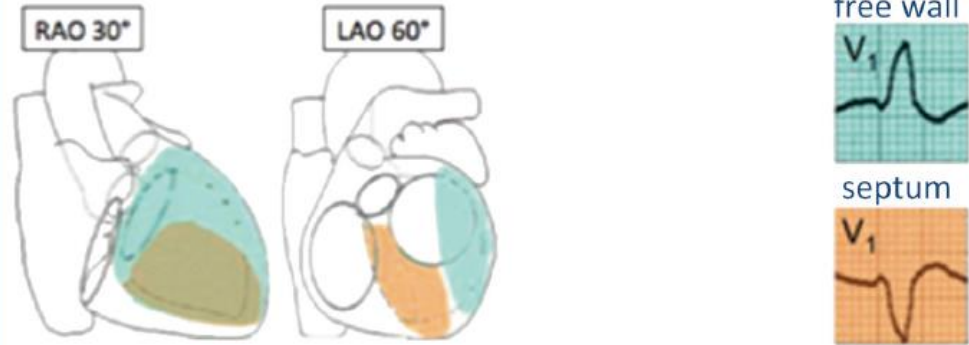


Site of left lead vs response to CRT



LV Pacing

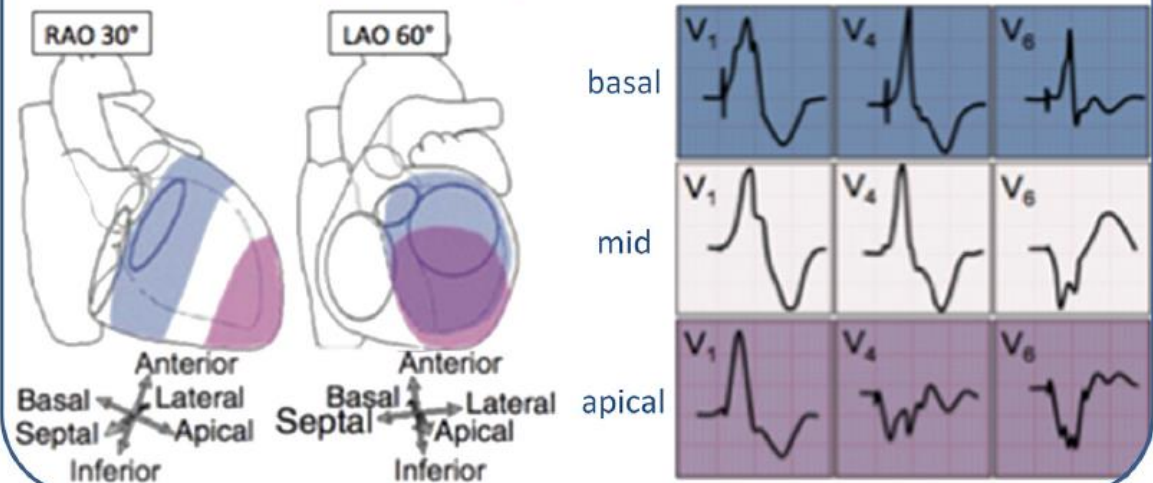
1. Check LV free wall lead position using V₁



2. Determine LV lead position in circumferential direction using aVF

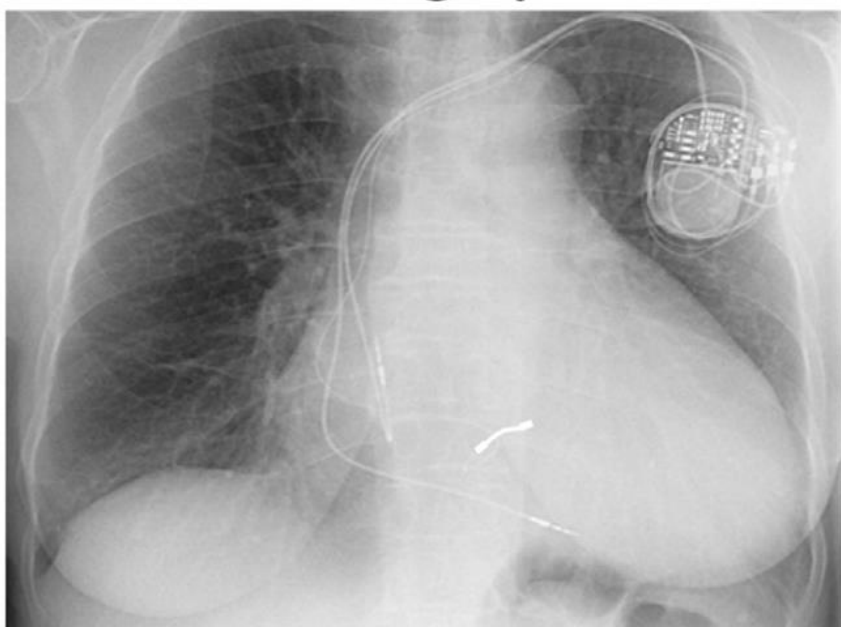


3. Determine LV lead position in apico-basal direction using precordial leads



Radiograph and ECG -basal inferolateral

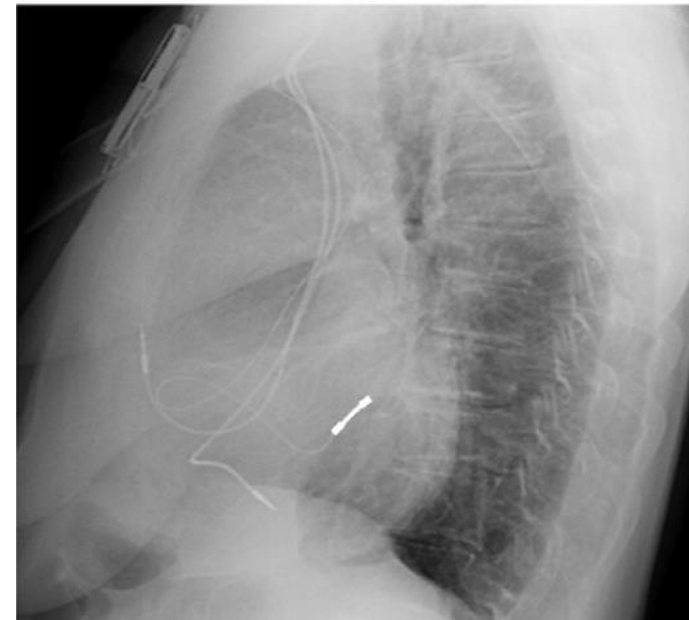
**Antero-posterior chest
radiograph**



**LV paced 1
lead ECG**

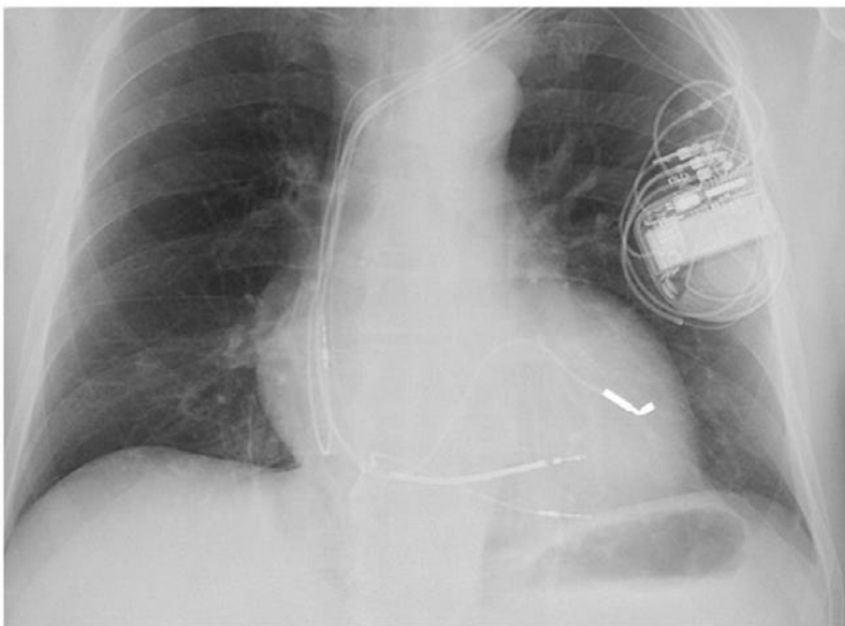


**Lateral chest
radiograph**



Radiograph and ECG -mid anterolateral

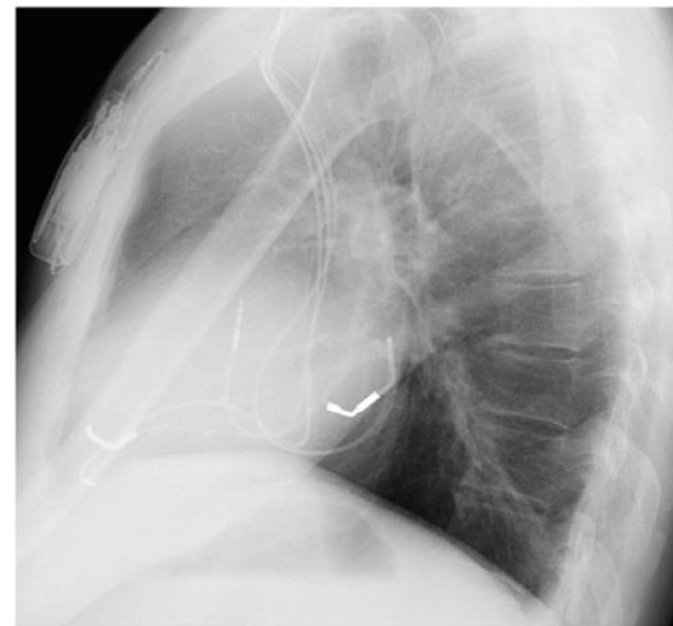
**Antero-posterior chest
radiograph**



**LV paced 12-
lead ECG**

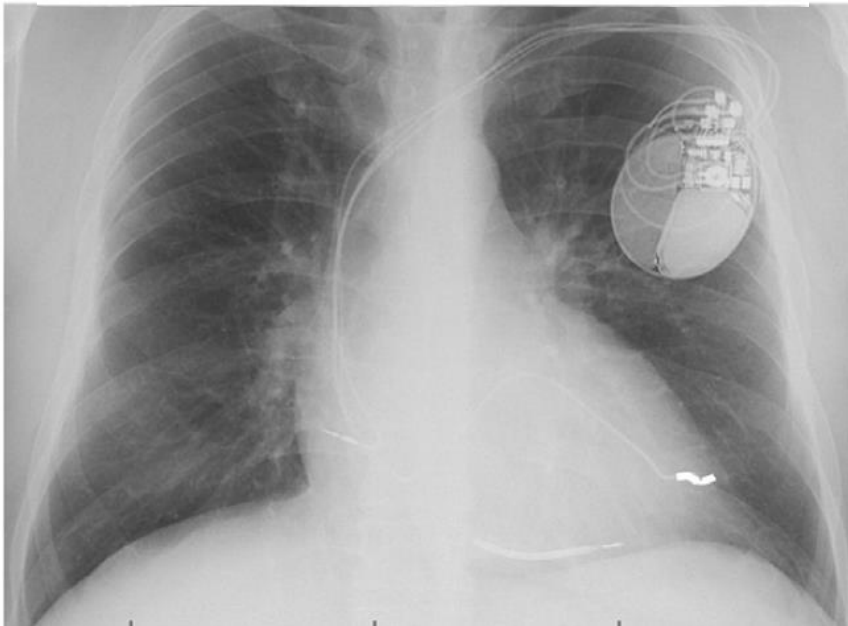


**Lateral chest
radiograph**

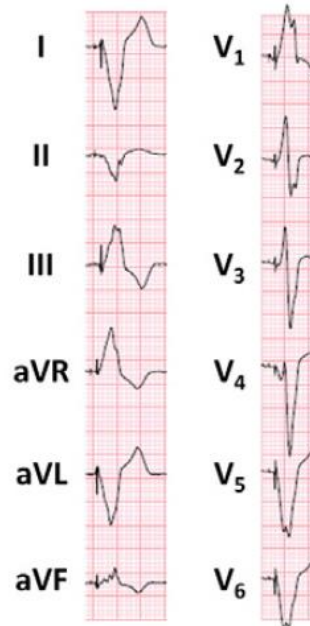


Radiograph and ECG -apical lateral

**Antero-posterior chest
radiograph**



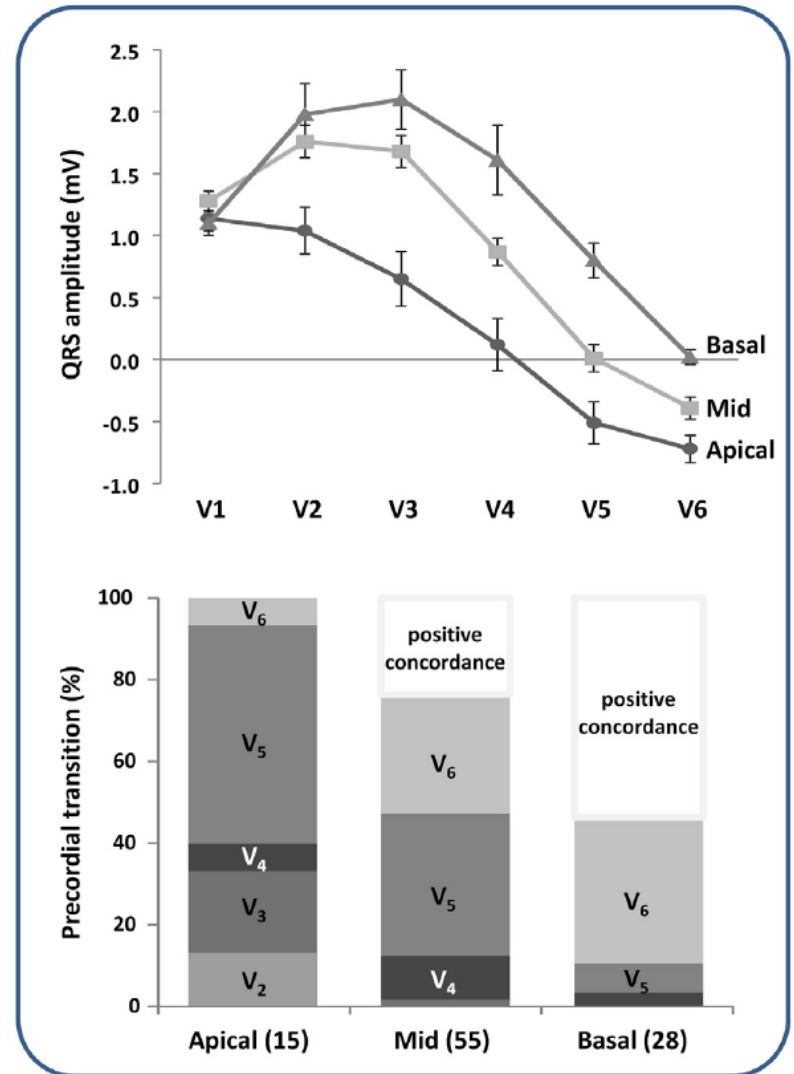
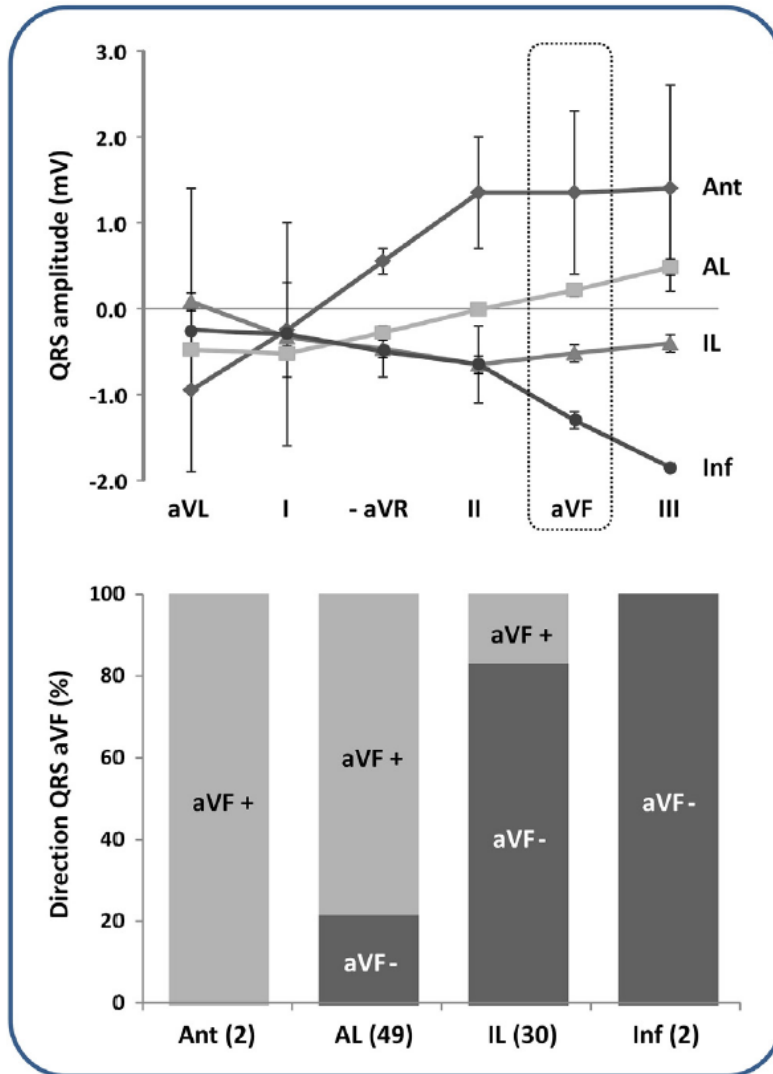
**LV paced 12-
lead ECG**



**Lateral chest
radiograph**



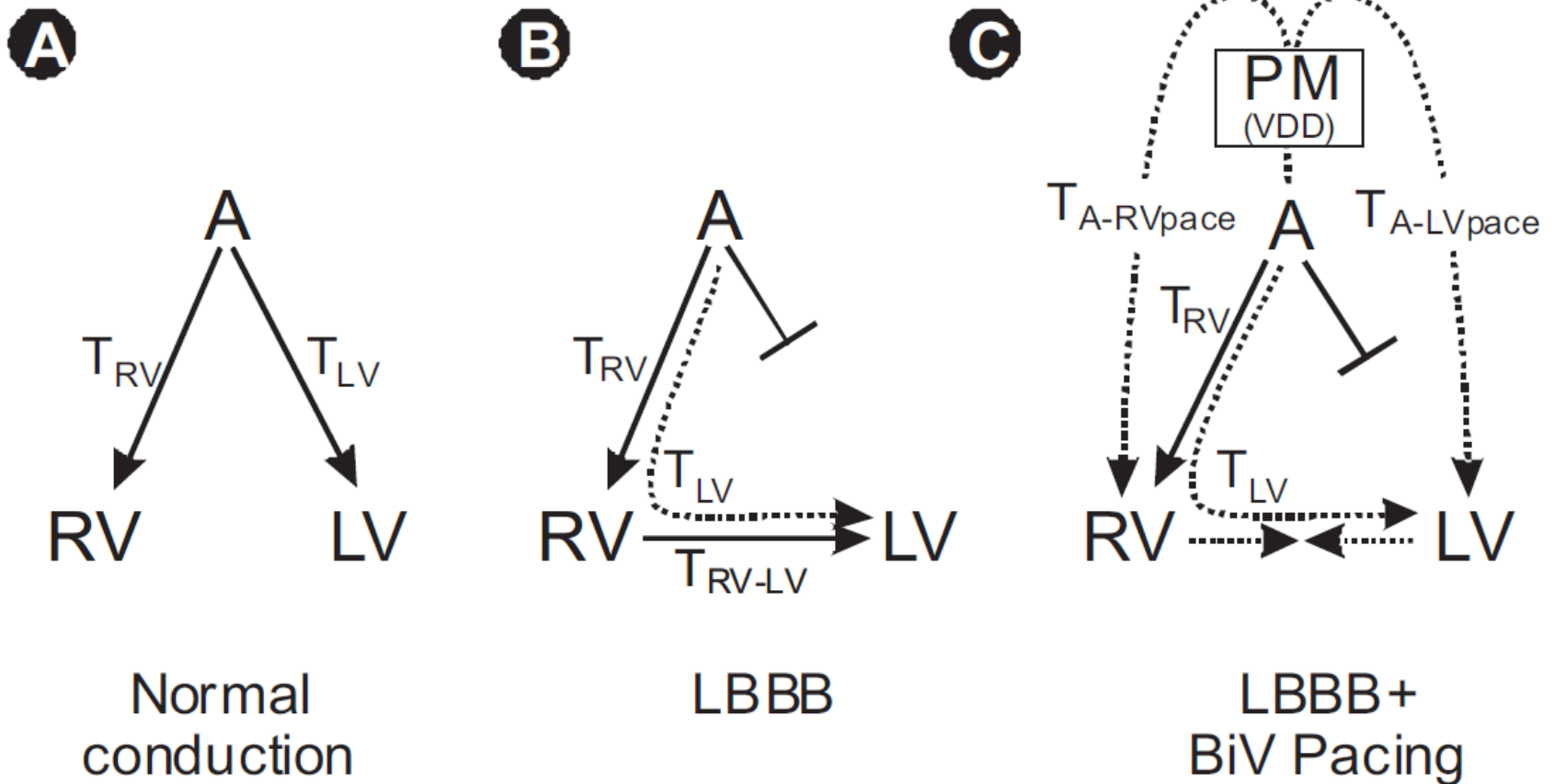
Evaluation of the concept of ECG-based estimation of LV lead position



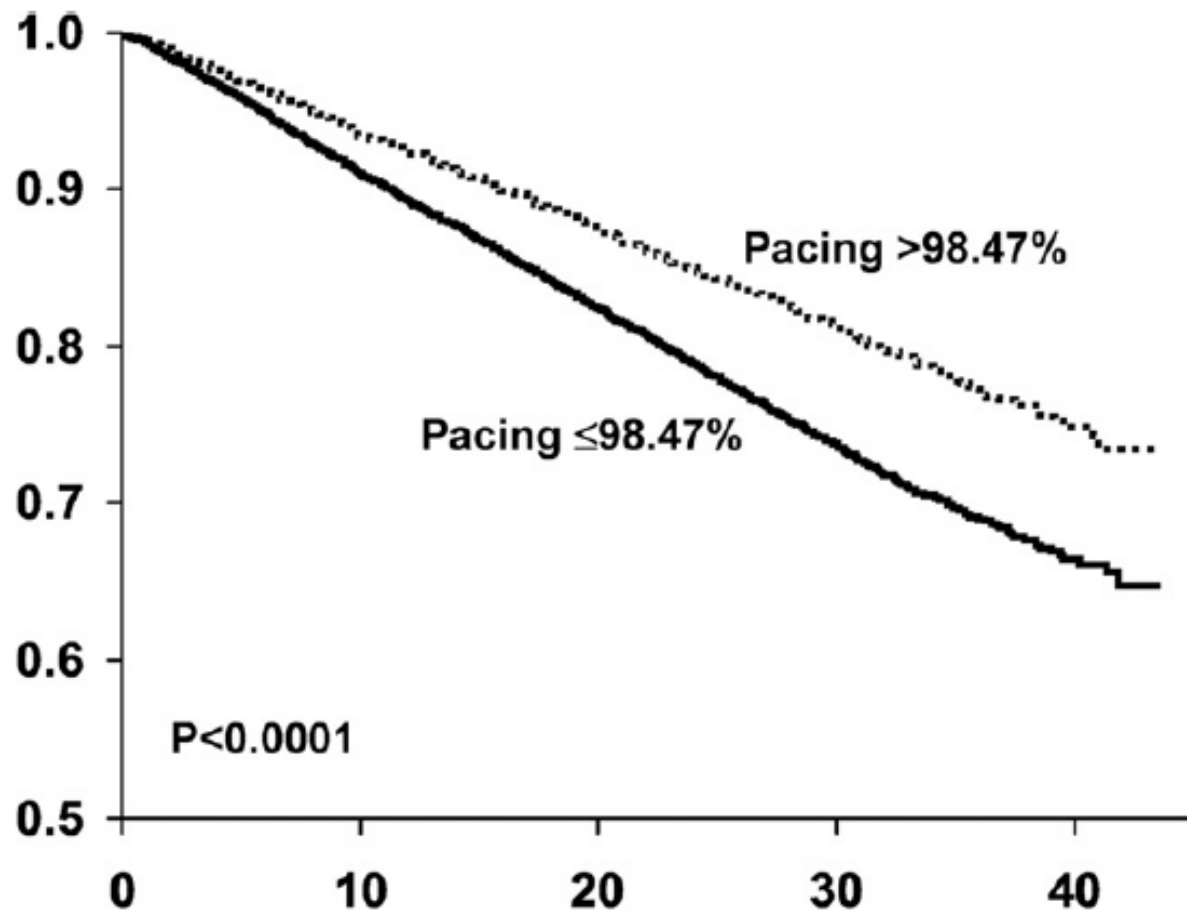
- Both avoidance of fractionation and large QRS width can be indirect tools to prevent LV pacing in a region of poor conduction due to scar or fibrosis

AV-delay optimization using the biventricular-paced ECG

Concept of CRT

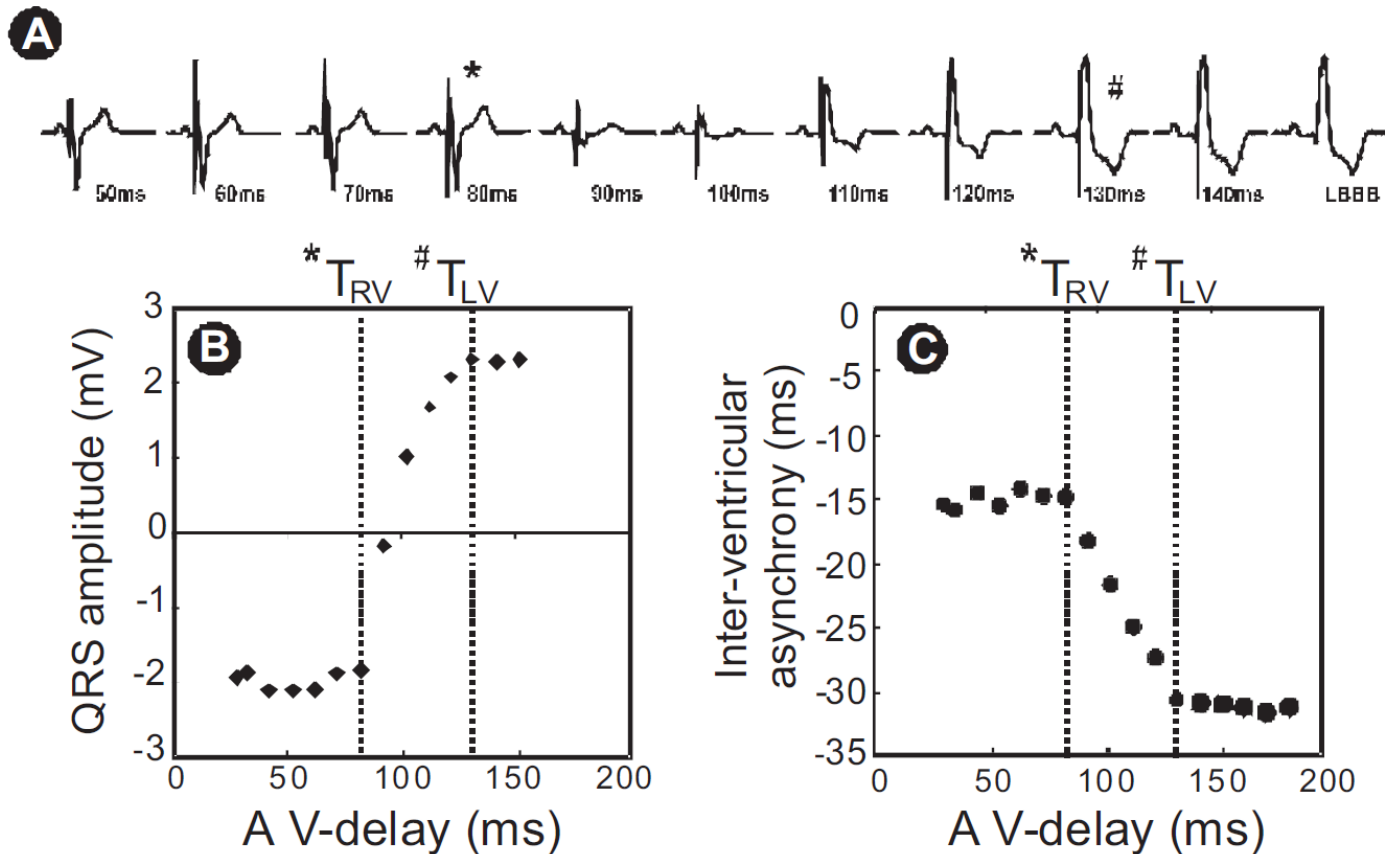


Survival vs Pacing rate

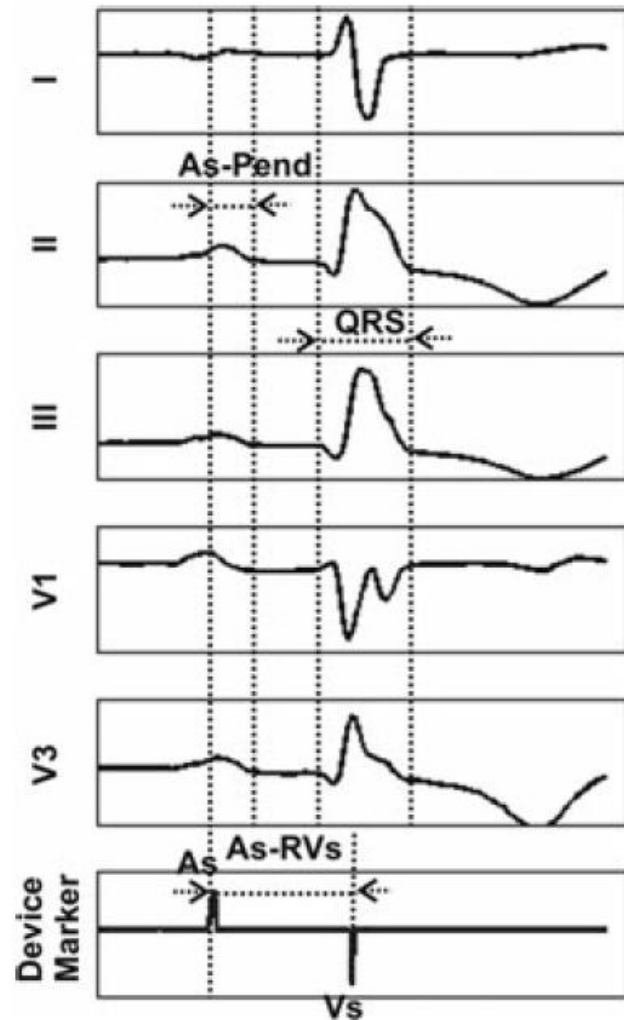


Identification method of the onset of intrinsic ventricular activation

The AV delay before the change in QRS morphology (**asterisk**) indicates the time of onset of intrinsic ventricular activation during stepwise increasing the AV delay with simultaneous BiV pacing

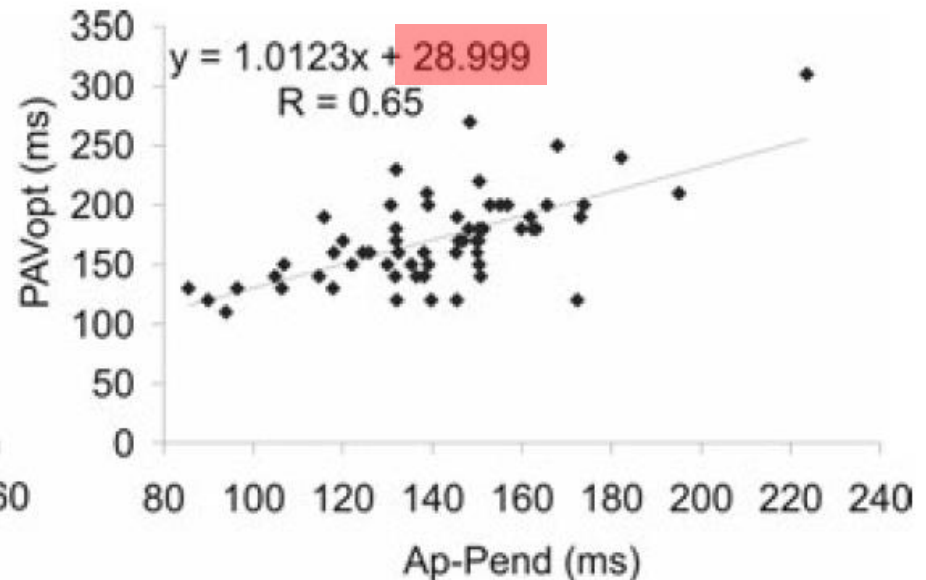
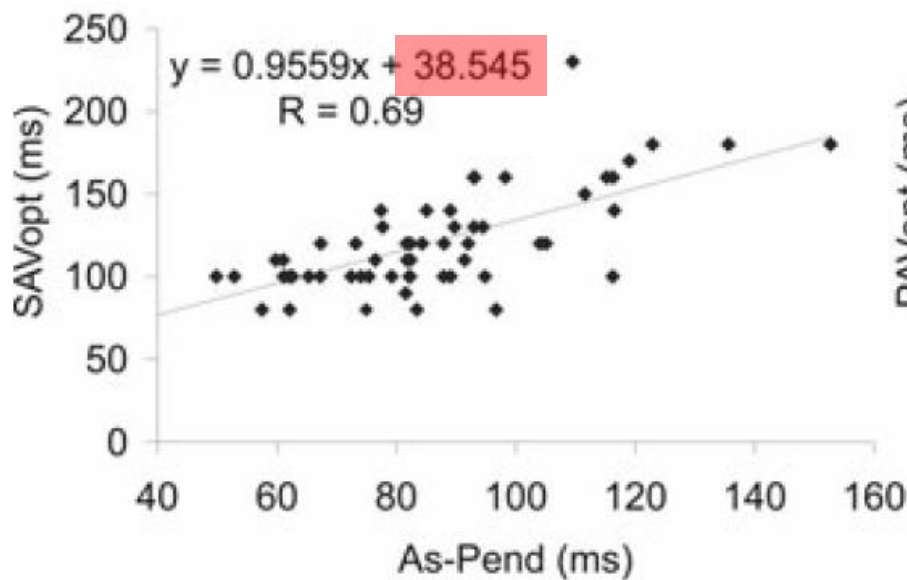


Correlation between the Echo-optimal SAV and PAV delays and electrocardiographic parameters



Correlation with SAVopt			
Parameter	Mean \pm SD	Correlation Coefficient (R)	P
As-Pend, ms	86 \pm 22	0.69	<0.0001
As-RVs, ms	215 \pm 58	0.45	0.0003
QRS, ms	154 \pm 30	0.24	0.014
Correlation with PAVopt			
Parameter	Mean \pm SD	Correlation Coefficient (R)	P
Ap-Pend, ms	141 \pm 25	0.65	<0.0001
Ap-RVs, ms	278 \pm 63	0.60	<0.0001
QRS, ms	155 \pm 29	0.20	0.038

Simple method of providing 30-40ms separation between the end of the P-wave and Ventricular pacing pulse for optimal AV delay

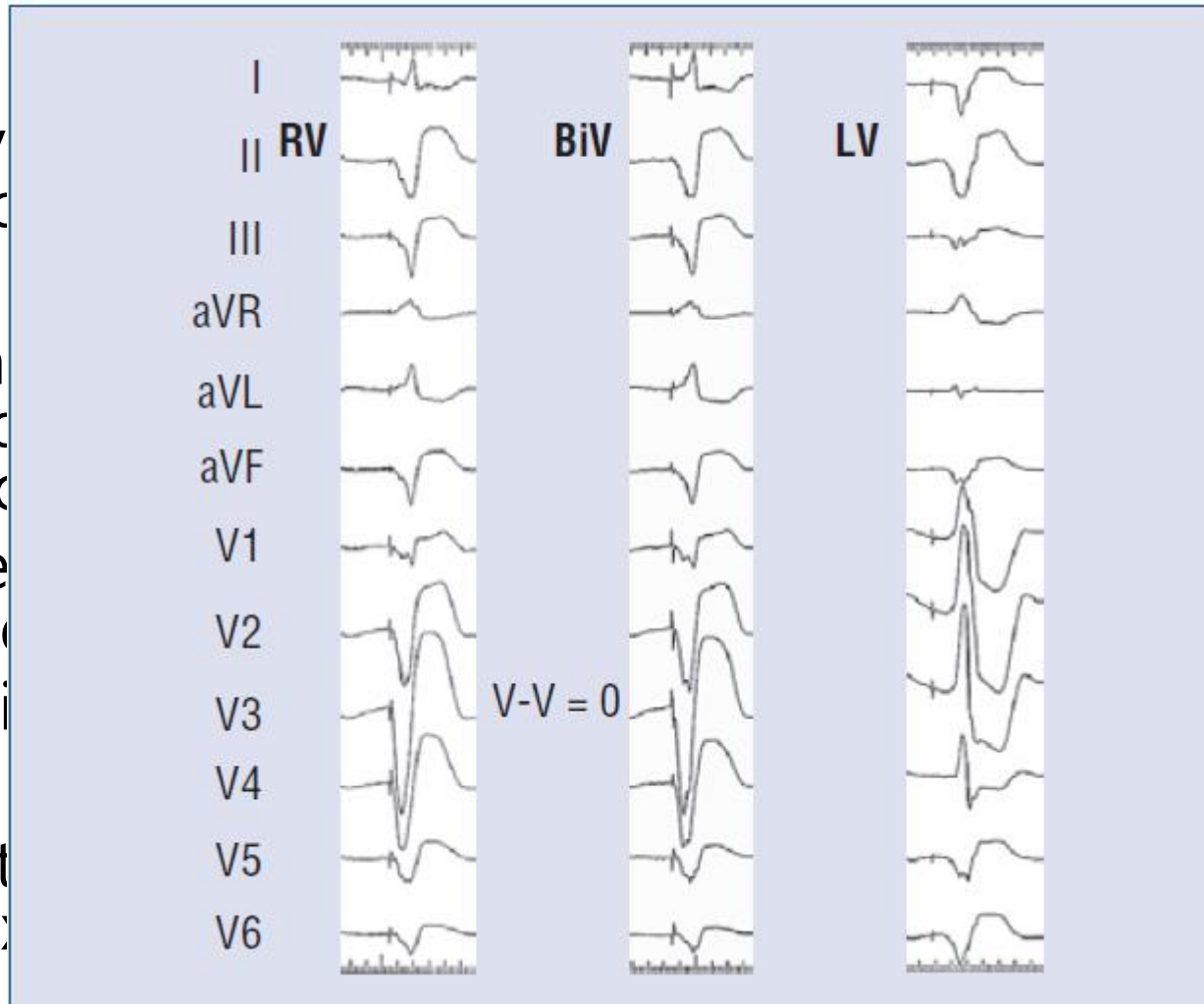


- $SAV_{ECG} = \min \{ As-Pend + 40ms, As-RVs-50ms \}$
- $PAV_{ECG} = \min \{ Ap-Pend + 30ms, Ap-RVs-50ms \}$

VV-interval optimization using the biventricular-paced ECG

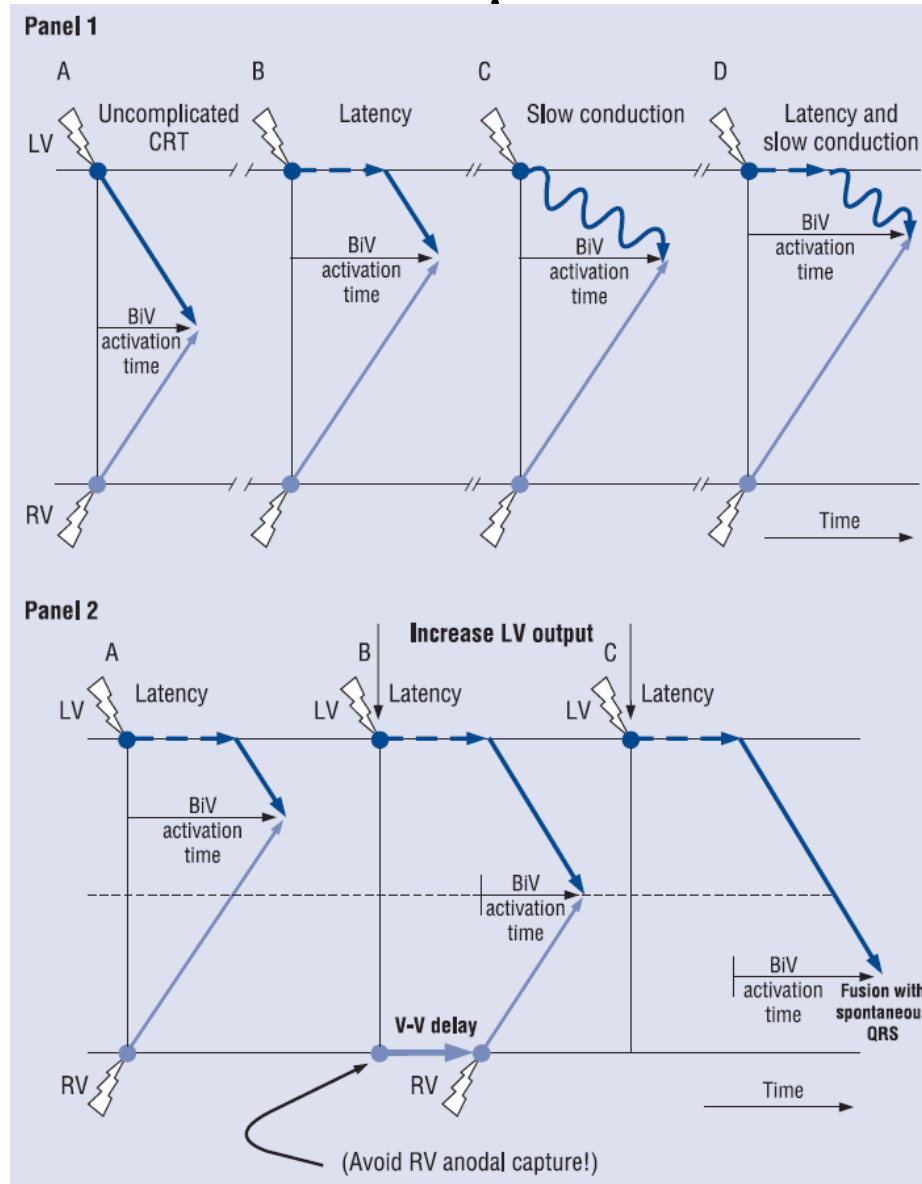
Simultaneous BiV pacing don't always result in best synchronization

- Delay
latenc
- LV pa
latenc
QRS c
- slowe
to enc
- high i
- So, it
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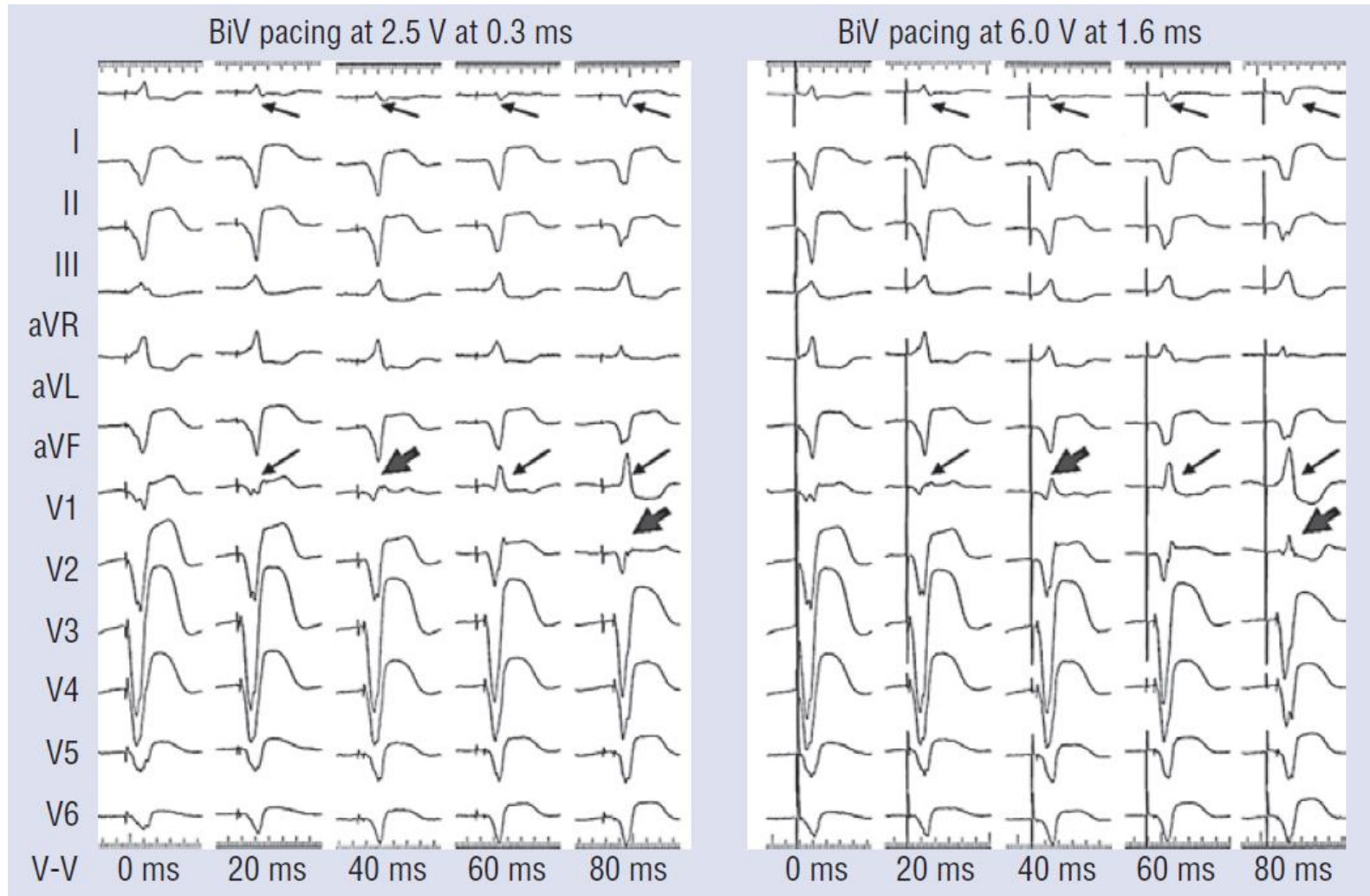


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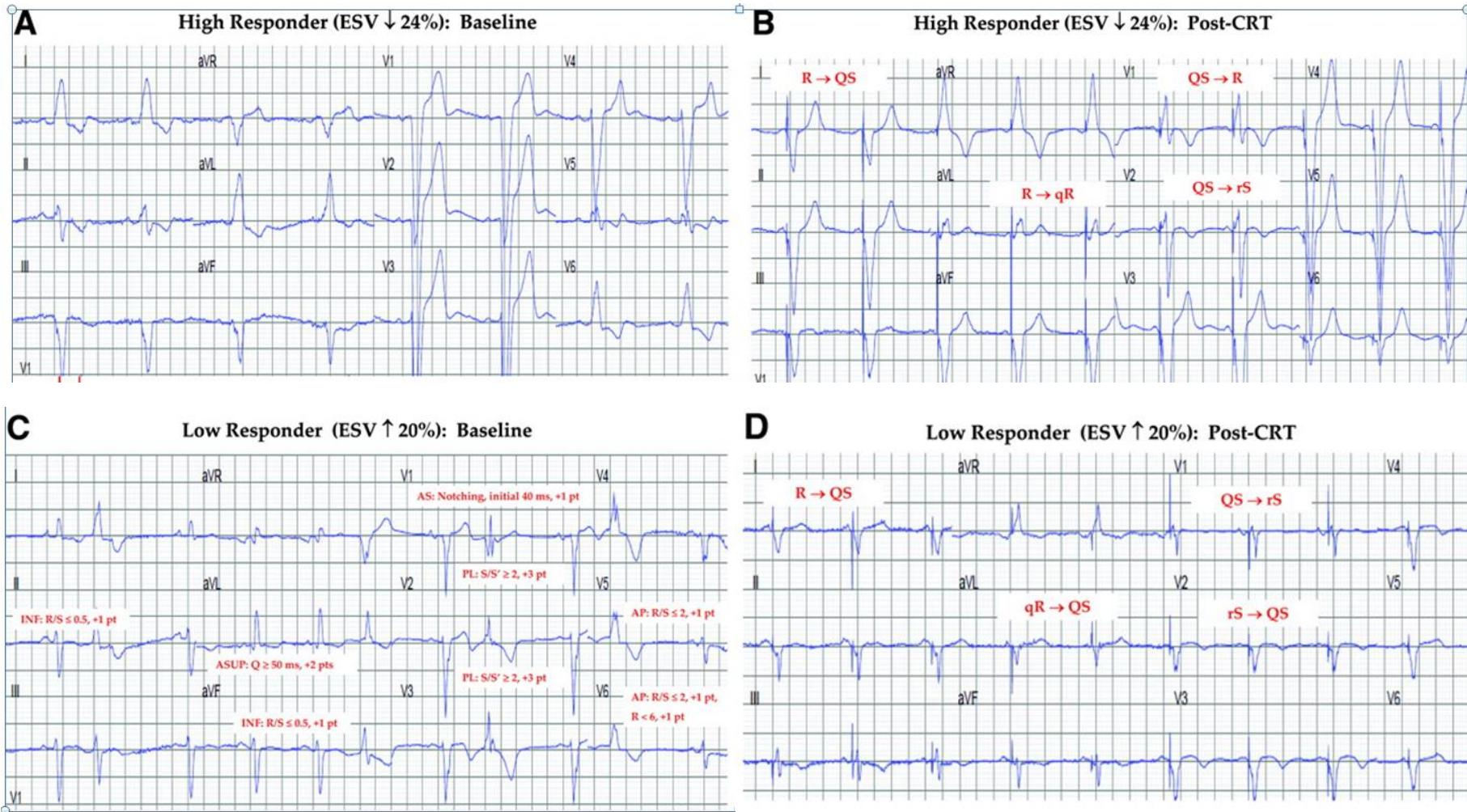
Diagrammatic representation of the significance of LV latency and slow conduction during Bi-V



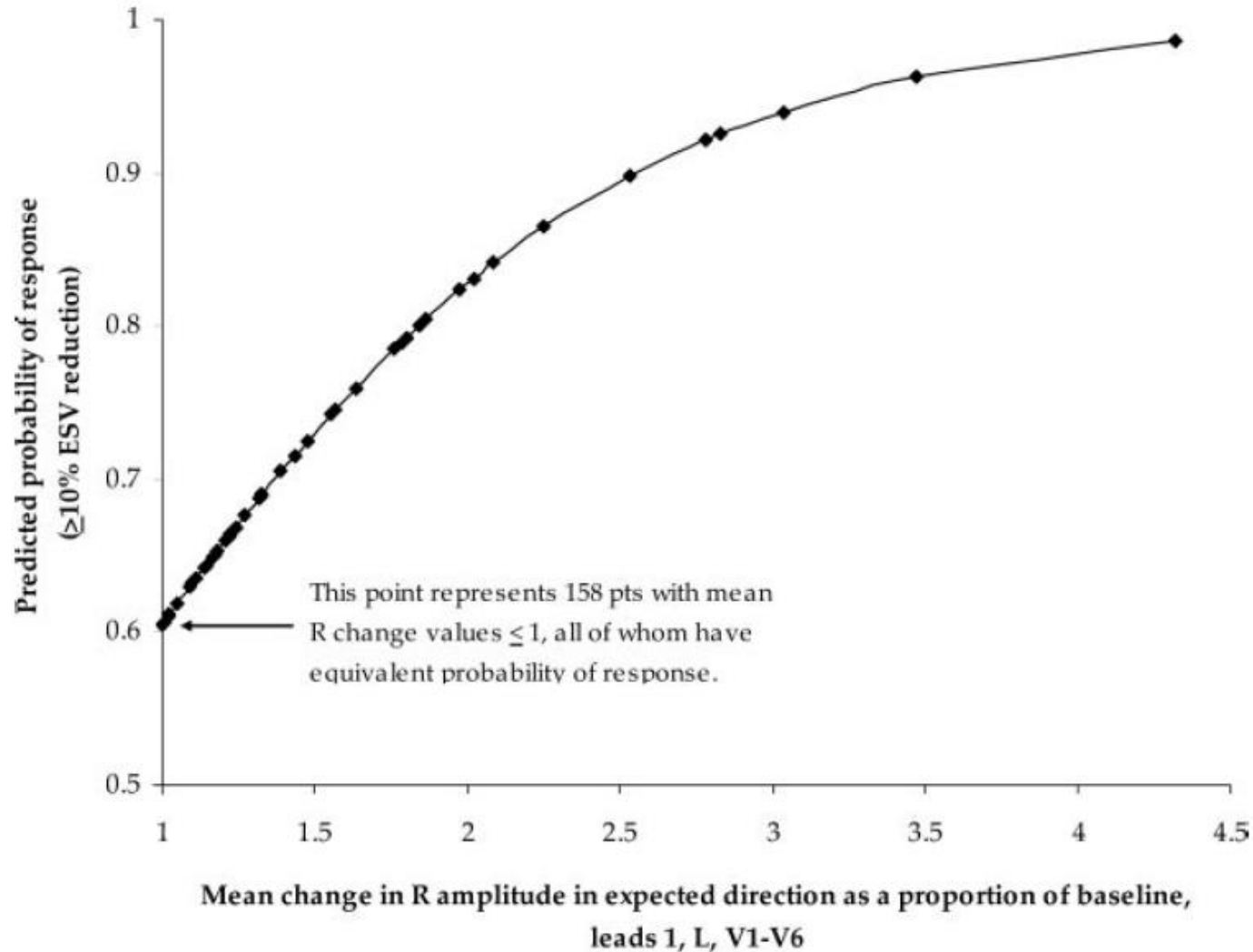
Impact of progressive LV pre-excitation during Bi-V pacing



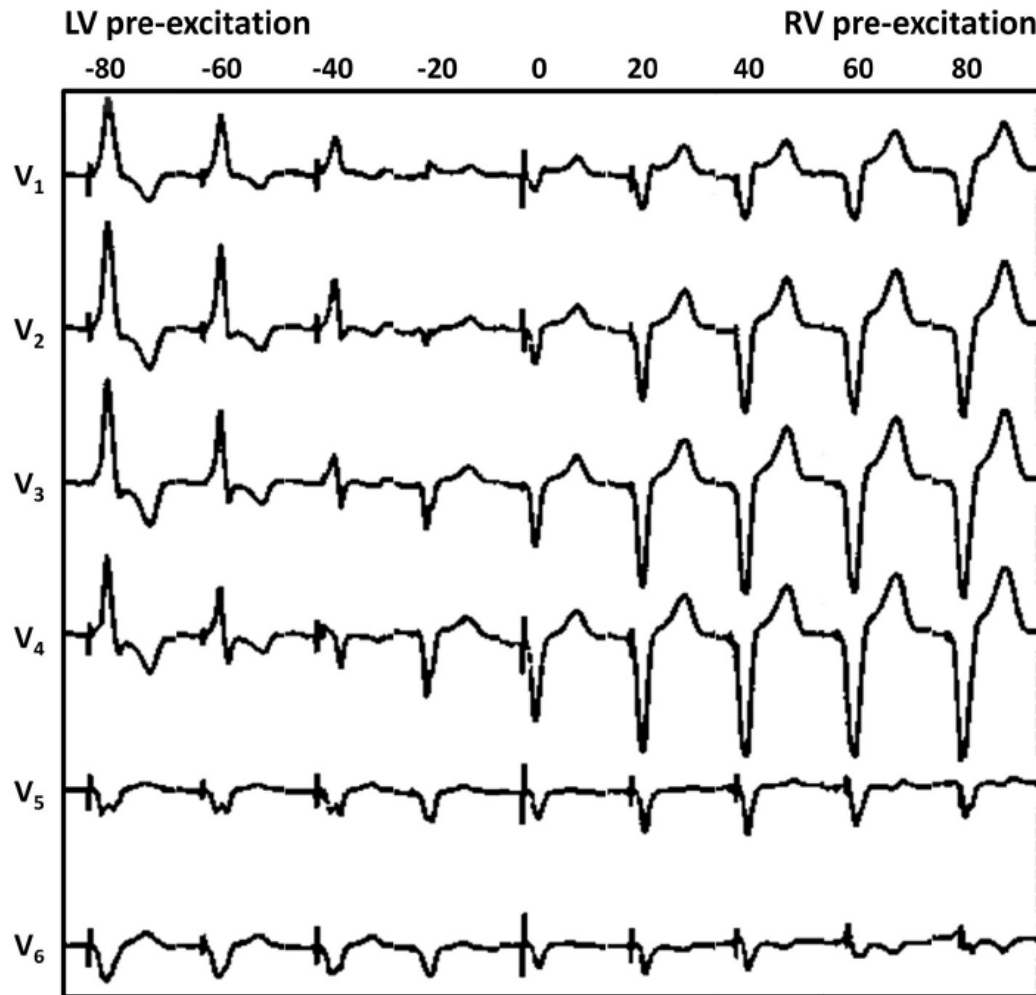
I. Best predictor was an increase in R-wave amplitude in V_1 and V_2



Greater changes in R-wave amplitudes after CRT, indicative of wavefront fusion, predict higher probability of response



II. QRS pattern during BiV pacing at various VV intervals – QRS vector



Lead V₁-V₃ help to identify contribution from LV pacing and show the gradual change of the QRS complex with changing VV interval

VV interval optimization

- **QRS vector** reflect electrical dyssynchrony.
- **QRS vector amplitude halfway** in between RV and LV pacing and minimal QRS vector area both reflect optimal resynchronization and timing of stimulation interval in CRT.
- The QRS morphology in lead V1-V3 that resembles a value halfway in between RV and LV pacing should probably be aimed for.

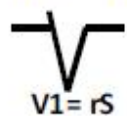
III. QRS fusion complex analysis

LBBB/ RV Pacing	LV Pacing	BV Pacing	QRSdiff (QRS _{BV} - QRS _{LBBB})	QRS Fusion Type*
			$(+, 0, -)$	1 QRS conformational change
				$(-)$
			$(+)$	3 Persistent LBBB

From leads V₁-V₂

A**LBBB**QRS_{LBBB} 168ms; LVAT_{max} 131 ms

QRS Score 24 (nonischemic)



V1 = rS

→ Lateral Lead
QRS Type 1**B****Simultaneous BV Pacing**QRS_{BV} 158 ms

QRSdiff = -10 ms



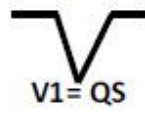
V1 = RS

Predicted response probability = 80%

Actual change in LVESV = -16

QRS_{LBBB} 190ms; LVAT_{max} 153 ms

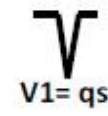
QRS Score 18 (ischemic)



V1 = QS

→ Lateral Lead
QRS Type 2QRS_{BV} 136 ms

QRSdiff = -54 ms



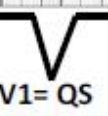
V1 = qS

Predicted response probability = 96%

Actual change in LVESV = -46

QRS_{LBBB} 152ms; LVAT_{max} 94 ms

QRS Score 24 (nonischemic)



V1 = QS

→ Non-Lateral Lead
QRS Type 3QRS_{BV} 165 ms

QRSdiff = +13 ms

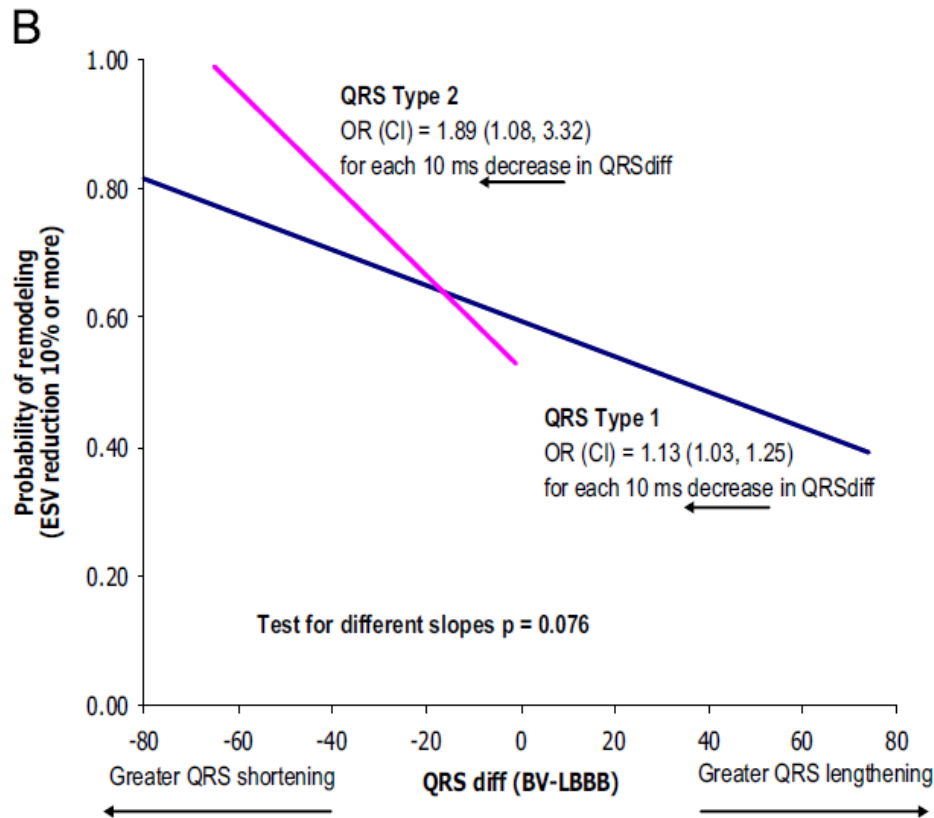


V1 = QS

Predicted response probability = 32%

Actual change in LVESV = +48

LV reverse remodeling by QRS fusion types



LVESV measure	QRS type 1 (n = 267)	QRS type 2 (n = 66)	<i>p</i>
LVESV—baseline	173 (130, 224)	182 (134, 229)	.54
LVESV—6 mo	135 (94, 196)	145 (94, 172)	.94
Δ LVESV	-26 (-56, -3.0)	-43 (-68, -18)	.035
Δ LVESV (% of baseline)	-16 (-34, -2.0)	-23 (-39, -10)	.15
LVESV reduction $\geq 10\%$	166 (62)	50 (76)	.034

Values are presented as mean \pm SD, n (%) or as median (25th, 75th percentile).

LV = left ventricular; LVESV = left ventricular end-systolic volume.

*Includes all LV lead sites.

Conclusion

Guide for CRT evaluation and optimization using the 12 lead ECG

1. Perform baseline ECG before CRT implantation

👉 Evaluate accurate Ix for CRT by identification of ventricular conduction disturbance like **LBBB, LVAT and QRS score**

2. Perform baseline ECG during

a) LV pacing (overdrive VVI mode to exclude fusion with intrinsic activation)

👉 Evaluate **LV lead position** from LV paced QRS morphology

👉 Evaluate **LV pacing latency interval**

b) Simultaneous Bi-V pacing(overdrive VVI mode)

👉 Evaluate **contribution of LV pacing** (latency interval)

Conclusion

Guide for CRT evaluation and optimization using the 12 lead ECG

3. **Identify onset of intrinsic ventricular activation** via stepwise increase of the AV-delay during simultaneous BiV pacing starting with a short AV-delay; the onset can be identified as the AV-delay where the QRS morphology change.
4. Program **sensed AV-delay** in a way that the pacing pulse occurs **40ms after the P wave**(special attention for the terminal negative part in V_1); the AV-delay must preferably be shorter than that with onset of intrinsic ventricular activation.
5. Increase lower rate above underlying rhythm and program **paced AV-delay** in a way that the pacing pulse occurs **30ms after the paced P wave**; AV delay must preferably be shorter than that with onset of intrinsic ventricular activation.

Conclusion

Guide for CRT evaluation and optimization using the 12 lead ECG

6. Program the VV-interval at a value with the QRS amplitude in lead V_{1-3} halfway in between RV and LV pacing; prefer a Rs over a rS pattern in lead V_1

Or

7. Program the VV-interval to lead the type 2 or type I BV fusion pattern in the lead V_1 If LV lead is in the lateral site.

경청해 주셔서 감사합니다.

