

## **Appendix 3 (as supplied by the authors): List of excluded studies**

### **B.1 Excluded – Wrong Population**

- (1) Beshai JF, Grimm RA, Nagueh SF, Baker JH, Beau SL, Greenberg SM et al. Cardiac resynchronization therapy in heart failure with narrow QRS complexes. *New England Journal of Medicine* 357 (24):2461 -71 , 2007.
- (2) Pham PP, Balaji S, Shen I, Ungerleider R, Li X, Sahn DJ. Impact of conventional versus biventricular pacing on hemodynamics and tissue Doppler imaging indexes of resynchronization postoperatively in children with congenital heart disease. *Journal of the American College of Cardiology* 46 (12):2284 -9 , 2005.

### **B.2 Excluded – Wrong Intervention and/or Comparator**

- (1) A comparison of antiarrhythmic with a defibrillatortrug therapy with implantable defibrillators in patients resuscitated from near-fatal ventricular arrhythmias. The Antiarrhythmics versus Implantable Defibrillators (AVID) Investigators. *New England Journal of Medicine* 337 (22):1576 -83 , 1997.
- (2) Anderson JL, Karagounis LA, Roskelley M, Osborn JS, Handrahan D. Effect of prophylactic antiarrhythmic therapy on time to implantable cardioverter with a defibrillatorefibrillator discharge in patients with ventricular tachyarrhythmias. *American Journal of Cardiology* 73 (9):683 -7 , 1994.
- (3) Auricchio A, Stellbrink C, Sack S, Block M, Vogt J, Bakker P et al. The Pacing Therapies for Congestive Heart Failure (PATH-CHF) study: rationale, design, and endpoints of a prospective randomized multicenter study. *American Journal of Cardiology* 83 (5B):130D -135D, 1999.
- (4) Bansch D, Antz M, Boczor S, Volkmer M, Tebbenjohanns J, Seidl K et al. Primary prevention of sudden cardiac death in idiopathic dilated cardiomyopathy: the Cardiomyopathy Trial (CAT). *Circulation* 105 (12):1453 -8, 2002.
- (5) Bansch D, Steffgen F, Gronefeld G, Wolpert C, Bocker D, Mletzko RU et al. The 1+1 trial: a prospective trial of a dual- versus a single-chamber implantable defibrillator in patients with slow ventricular tachycardias. *Circulation* 110 (9):1022 -9, 2004.
- (6) Bansch D, Kottkamp H, Gronefeld G, Vogt J, Israel C, Bocker D et al. The quick-implantable with a defibrillatorefibrillator trial. *Europace* 9 (12):1144 -50 , 2007.
- (7) Bardy GH, Poole JE, Kudenchuk PJ, Dolack GL, Kelso D, Mitchell R. A prospective randomized repeat-crossover comparison of antitachycardia pacing with low-energy cardioversion. *Circulation* 87 (6):1889 -96, 1993.

- (8) Bardy GH, Lee KL, Mark DB, Poole JE, Packer DL, Boineau R et al. Amiodarone or an implantable cardioverter with a defibrillatorefibrillator for congestive heart failure.[Erratum appears in N Engl J Med. 2005 May 19;352(20):2146]. New England Journal of Medicine 352 (3):225 -37, 2005.
- (9) Belardinelli R, Capestro F, Misiani A, Scipione P, Georgiou D. Moderate exercise training improves functional capacity, quality of life, and endothelium with a defibrillatoredependent vasodilation in chronic heart failure patients with implantable cardioverter defibrillators and cardiac resynchronization therapy. European Journal of Cardiovascular Prevention & Rehabilitation 13(5):818 -25, 2006.
- (10) Bigger JT, Jr. Prophylactic use of implanted cardiac defibrillators in patients at high risk for ventricular arrhythmias after coronary-artery bypass graft surgery. Coronary Artery Bypass Graft (CABG) Patch Trial Investigators. New England Journal of Medicine 337 (22):1569 -75 , 1997.
- (11) Borian G, Lubinski A, Capucci A, Niederle R, Kornacewicz-Jack Z, Wnuk-Wojnar AM et al. A multicentre, double-blind randomized crossover comparative study on the efficacy and safety of dofetilide vs sotalol in patients with inducible sustained ventricular tachycardia and ischaemic heart disease. European Heart Journal 22 (23):2180 -91, 2001.
- (12) Borian G, Kranig W, Donal E, Calo L, Casella M, Delarche N et al. A randomized double-blind comparison of biventricular versus left ventricular stimulation for cardiac resynchronization therapy: the Biventricular versus Left Univentricular Pacing with implantable defibrillator Back-up in Heart Failure Patients (B-LEFT HF) trial. American Heart Journal 159 (6):1052 -1058 e1, 2010.
- (13) Borian G, Muller CP, Seidl KH, Grove R, Vogt J, Danschel W et al. Randomized comparison of simultaneous biventricular stimulation versus optimized interventricular delay in cardiac resynchronization therapy. The Resynchronization for the hemodynamic Treatment for Heart Failure Management II implantable cardioverter defibrillator (RHYTHM II implantable defibrillator) study. American Heart Journal 151 (5):1050 -8, 2006.
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- (15) Buxton AE, Lee KL, Fisher JD, Josephson ME, Prystowsky EN, Hafley G. A randomized study of the prevention of sudden death in patients with coronary artery disease. Multicenter Unsustained Tachycardia Trial Investigators.[Erratum appears in N Engl J Med 2000 Apr 27;342(17):1300]. New England Journal of Medicine 341 (25):1882 -90, 1999.
- (16) Capucci A, Romano S, Puglisi A, Santini M, Pagani M, Cazzin R et al. Dual chamber pacing with optimal AV delay in congestive heart failure: a randomized study. Europace 1(3):174 -8 , 1999.

- (17) Connolly SJ, Gent M, Roberts RS, Dorian P, Roy D, Sheldon RS et al. Canadian implantable defibrillator study (CIDS) : a randomized trial of the implantable cardioverter defibrillator against amiodarone. *Circulation* 101 (11):1297 -302 , 2000.
- (18) Connolly SJ, Dorian P, Roberts RS, Gent M, Bailin S, Fain ES et al. Comparison of beta-blockers, amiodarone plus beta-blockers, or sotalol for prevention of shocks from implantable cardioverter defibrillators: the OPTIC Study: a randomized trial. *JAMA* 295 (2):165 -71 , 2006.
- (19) Doll N, Piorkowski C, Czesla M, Kallenbach M, Rastan AJ, Arya A et al. Epicardial versus transvenous left ventricular lead placement in patients receiving cardiac resynchronization therapy: results from a randomized prospective study. *Thoracic & Cardiovascular Surgeon* 56 (5):256 -61 , 2008.
- (20) Dorian P, Philippon F, Thibault B, Kimber S, Sterns L, Greene M et al. Randomized controlled study of detection enhancements versus rate-only detection to prevent inappropriate therapy in a dual-chamber implantable cardioverter with a defibrillatorefibrillator. *Heart Rhythm* 1(5):540 -7 , 2004.
- (21) Doshi RN, Daoud EG, Fellows C, Turk K, Duran A, Hamdan MH et al. Left ventricular-based cardiac stimulation post AV nodal ablation evaluation (the PAVE study). *Journal of Cardiovascular Electrophysiology* 16 (11):1160 -5, 2005.
- (22) Garrigue S, Bordachar P, Reuter S, Jais P, Haissaguerre M, Clementy J. Comparison of permanent left ventricular and biventricular pacing in patients with heart failure and chronic atrial fibrillation: a prospective hemodynamic study. [Review] [46 refs]. *Cardiac Electrophysiology Review* 7 (4):315 -24 , 2003.
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- (24) Gasparini M, Bocchiardo M, Lunati M, Ravazzi PA, Santini M, Zardini M et al. Comparison of 1-year effects of left ventricular and biventricular pacing in patients with heart failure who have ventricular arrhythmias and left bundle-branch block: the Bi vs Left Ventricular Pacing: an International Pilot Evaluation on Heart Failure Patients with Ventricular Arrhythmias (BELIEVE) multicenter prospective randomized pilot study. *American Heart Journal* 152 (1):155 e1 -7 , 2006.
- (25) Greene HL, Poole JE, Kudenchuk PJ, Dolack GL, Bardy GH, Cobb LA et al. Randomized antiarrhythmic drug therapy in survivors of cardiac arrest (the CASCADE study). *American Journal of Cardiology* 72 (3) (pp 280 -287 ), 1993;1993.
- (26) Hohnloser SH, Kuck KH, Dorian P, Roberts RS, Hampton JR, Hatala R et al. Prophylactic use of an implantable cardioverter with a defibrillatorefibrillator after acute myocardial infarction. *New England Journal of Medicine* 351 (24 ):2481 -8 , 2004.

- (27) Hoijer C-J, Meurling C, Brandt J. Upgrade to biventricular pacing in patients with conventional pacemakers and heart failure: A double-blind, randomized crossover study. *Europace* 8 (1) (pp 51 -55 ), 2006;Jan.
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- (29) Kalbfleisch SJ, Williamson B, Man KC, Vorperian V, Hummel JD, Hasse C et al. Prospective, randomized comparison of conventional and high dose loading regimens of amiodarone in the treatment of ventricular tachycardia. *Journal of the American College of Cardiology* 22 (6 ):1723 -9 , 1993.
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- (32) Kuck K-H, Cappato R, Siebels J, Ruppel R. Randomized comparison of antiarrhythmic drug therapy with implantable defibrillators in patients resuscitated from cardiac arrest: The cardiac arrest study hamburg (CASH). *Circulation* 102 (7 ) (pp 748 -754 ), 2000;2000.
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- (34) Leclercq C, Gadler F, Kranig W, Ellery S, Gras D, Lazarus A et al. A randomized comparison of triple-site versus dual-site ventricular stimulation in patients with congestive heart failure. *Journal of the American College of Cardiology* 51 (15 ):1455 -62 , 2008.
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- (36) Leclercq C, Cazeau S, Lellouche D, Fossati F, Anselme F, Davy JM et al. Upgrading from single chamber right ventricular to biventricular pacing in permanently paced patients with worsening heart failure: The RD-CHF Study.[Erratum appears in *Pacing Clin Electrophysiol*. 2007 Nov;30(11):1424]. *Pacing & Clinical Electrophysiology* 30 Suppl 1:S23 -30 , 2007.

- (37) Maynard C. Rehospitalization in surviving patients of out-of-hospital ventricular fibrillation (the CASCADE study). *American Journal of Cardiology* 72 (17) (pp 1295 - 1300 ), 1993;1993.
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- (40) Moss AJ, Hall WJ, Cannom DS, Daubert JP, Higgins SL, Klein H et al. Improved survival with an implanted defibrillator in patients with coronary disease at high risk for ventricular arrhythmia. Multicenter Automatic Defibrillator Implantation Trial Investigators. *New England Journal of Medicine* 335 (26 ):1933 -40 , 1996.
- (41) Moss AJ, Zareba W, Hall WJ, Klein H, Wilber DJ, Cannom DS et al. Prophylactic implantation of a defibrillator in patients with myocardial infarction and reduced ejection fraction. *New England Journal of Medicine* 346 (12 ):877 -83 , 2002.
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- (43) Occhetta E, Bortnik M, Magnani A, Francalacci G, Piccinino C, Plebani L et al. Prevention of ventricular desynchronization by permanent para-Hisian pacing after atrioventricular node ablation in chronic atrial fibrillation: a crossover, blinded, randomized study versus apical right ventricular pacing. *Journal of the American College of Cardiology* 47 (10):1938 -45 , 2006.
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- (50) Raviele A, Bongiorni MG, Brignole M, Cappato R, Capucci A, Gaita F et al. Early EPS/implantable defibrillator strategy in survivors of acute myocardial infarction with severe left ventricular dysfunction on optimal beta-blocker treatment. The beta-blocker strategy plus implantable defibrillator trial. *Europace* 7 (4):327 -37 , 2005.
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- (75) Wonisch M, Lercher P, Scherr D, Maier R, Pokan R, Hofmann P et al. Influence of permanent right ventricular pacing on cardiorespiratory exercise parameters in chronic heart failure patients with implanted cardioverter defibrillators. *Chest* 127 (3):787 -93 , 2005.

### B.3 Excluded – No Mortality Outcome

- (1) Schlosshan D, Barker D, Lewis N, Pepper C, Tan LB. A mechanistic investigation into how long-term resynchronization therapy confers ongoing cardiac functional benefits and improved exercise capacity. *American Journal of Cardiology* 103 (5):701-8, 2009.
- (2) Yu CM, Lin H, Fung WH, Zhang Q, Kong SL, Sanderson JE. Comparison of acute changes in left ventricular volume, systolic and diastolic functions, and intraventricular synchronicity after biventricular and right ventricular pacing for heart failure. *American Heart Journal* 145 (5):E18, 2003.
- (3) Auricchio A, Stellbrink C, Butter C, Sack S, Vogt J, Misier AR et al. Clinical efficacy of cardiac resynchronization therapy using left ventricular pacing in heart failure patients stratified by severity of ventricular conduction delay. *Journal of the American College of Cardiology* 42 (12 ):2109 -16 , 2003.

### B.4 Excluded – Wrong Study Design

- (1) Albert CM, Rosenthal L, Calkins H, Steinberg JS, Ruskin JN, Wang P et al. Driving and implantable cardioverter with a defibrillatorefibrillator shocks for ventricular arrhythmias: results from the TOVA study. *Journal of the American College of Cardiology* 50 (23):2233 -40 , 2007.
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- (5) Buxton AE, Lee KL, dicarlo L, Gold MR, Greer GS, Prystowsky EN et al. Electrophysiologic testing to identify patients with coronary artery disease who are at risk for sudden death. Multicenter Unsustained Tachycardia Trial Investigators. *New England Journal of Medicine* 342 (26 ):1937 -45 , 2000.

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- (7) Ermis C, Seutter R, Zhu AX, Benditt LC, vanheel L, Sakaguchi S et al. Impact of upgrade to cardiac resynchronization therapy on ventricular arrhythmia frequency in patients with implantable cardioverter with a defibrillatorefibrillators. Journal of the American College of Cardiology 46 (12 ):2258 -63 , 2005.
- (8) Fung JW, Zhang Q, Yip GW, Chan JY, Chan HC, Yu CM. Effect of cardiac resynchronization therapy in patients with moderate left ventricular systolic dysfunction and wide QRS complex: a prospective study. Journal of Cardiovascular Electrophysiology 17(12 ):1288 -92 , 2006.
- (9) Ganz DA, Lamas GA, Orav EJ, Goldman L, Gutierrez PR, Mangione CM. Age-related differences in management of heart disease: a study of cardiac medication use in an older cohort. Pacemaker Selection in the Elderly (PASE) Investigators. Journal of the American Geriatrics Society 47 (2):145 -50 , 1999.
- (10) Leclercq C, Cazeau S, Ritter P, Alonso C, Gras D, Mabo P et al. A pilot experience with permanent biventricular pacing to treat advanced heart failure. American Heart Journal 140 (6 ):862 -70 , 2000.
- (11) Leon AR, Abraham WT, Brozena S, Daubert JP, Fisher WG, Gurley JC et al. Cardiac resynchronization with sequential biventricular pacing for the treatment of moderate-to-severe heart failure. Journal of the American College of Cardiology 46 (12 ):2298 -304 , 2005.
- (12) Leon AR, Abraham WT, Curtis AB, Daubert JP, Fisher WG, Gurley J et al. Safety of transvenous cardiac resynchronization system implantation in patients with chronic heart failure: combined results of over 2,000 patients from a multicenter study program. Journal of the American College of Cardiology 46 (12 ):2348 -56 , 2005.
- (13) Nagele H, Bohlmann M, Eck U, Petersen B, Rodiger W. Combination therapy with carvedilol and amiodarone in patients with severe heart failure. European Journal of Heart Failure 2(1):71 -9, 2000.
- (14) Higgins SL, Hummel JD, Niazi IK, Giudici MC, Worley SJ, Saxon LA et al. Cardiac resynchronization therapy for the treatment of heart failure in patients with intraventricular conduction delay and malignant ventricular tachyarrhythmias. Journal of the American College of Cardiology 42 (8 ):1454 -9 , 2003.

## B.5 Excluded – Sub-study / Sub-analysis / Extension Study

- (1) Alonso C, Ritter P, Leclercq C, Mabo P, Bailleul C, Daubert JC et al. Effects of cardiac resynchronization therapy on heart rate variability in patients with chronic systolic heart

- failure and intraventricular conduction delay. American Journal of Cardiology 91 (9):1144 -7 , 2003.
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