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Sudden Death After Transcatheter Aortic Valve Implantation. Are Bradyarrhythmias Always The Cause?

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Abstract

Transcatheter Aortic-Valve Implantation (TAVI) is considered to be highly effective in the treatment of high-risk patients with severe aortic stenosis. After TAVI, the rate of pacemaker implantation is 6.5%-40%. Some reports of sudden death after TAVI are mostly attributed to bradyarrhythmias.

We report the case of three patients who experienced sudden cardiac death or aborted sudden cardiac death after TAVI. All patients were affected from ischemic heart disease with an ejection fraction of approximately 40% and underwent pacemaker implantation (PM) after the procedure due to 1rst degree atrioventricular block (AV) and left bundle branch block (LBBB). One of the patients died suddenly 30 days after the procedure. The PM interrogation revealed many episodes of non sustain ventricular tachycardias (NSVT) and one episode of ventricular fibrillation (VF) that led to death. The other two patients had syncope and during PM interrogation episodes of ventricular tachycardia >12 sec were recorded. Patients affected by ischemic heart disease undergoing TAVI, especially with borderline coronary lesions should receive particular attention in order to avoid potentially lethal ventricular arrhythmias. In addition, the physiopathologic mechanism of sudden arrhythmic death in these patients needs to be clarified.

Introduction

Transcatheter Aortic-Valve Implantation (TAVI) is considered to be highly effective in the treatment of patients with severe aortic stenosis who are inoperable. After TAVI, the rate of pacemaker implantation is 6.5%-40%.¹⁻³ Some reports of sudden death after TAVI are mostly attributed to bradyarrhythmias. Nevertheless, ventricular arrhythmias should always be taken in to consideration in patients with borderline ejection fraction.

Case Presentation

Patients with severe aortic stenosis at increased risk for surgical aortic valve replacement who underwent TAVI to our institution from October 2010 to May 2013 were retrospectively analyzed. Pa-

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Disclosures:

None.

Corresponding Author: Lida Pieretta Papavasileiou, M.D., Ph.D. Electrophysiology, Pacemaker and ICD Unit, Hygeia Hospital 9, Erythrou Stavrou Str., GR15123, Marousi, Athens, Greece. tients qualified for TAVI in the presence of an indication for aortic valve replacement according to current guidelines⁴ but unsuitable for surgical procedure due to high risk of morbidity and mortality as calculated with STScore and logistic Euroscore⁵⁻⁷

TAVI was performed either with the Medtronic Core Valve or the Edwards-SAPIENXT bioprosthesis with standard retrograde technique.¹ Vascular access was obtained either percutaneous transfemoral or surgical transaortic. As required, a temporary pacemaker was inserted before the procedure in all patients and kept in site for a minimum of 48 hr. Prior to the procedure all patients performed screening by coronary angiogram and/or (in rare cases) stress test if considered necessary and ECG-gated multidetector computed tomography.

Data Analysis

From each patient, different parameters were collected with respect to patients' characteristics, demographic and echocardiographic parameters as well as clinical and procedural data.

A 12-lead electrocardiogram (ECG) was recorded before the procedure, immediately after, every day until discharge and during every follow-up (FU) visit. Electrocardiographic characteristics of pre and postprocedural ECGs were analyzed by 2 independent investigators



(MC, AC) in order to assess ECG changes. Follow-up visits were scheduled at 1, 3, 6 and 12 months after discharge.

In patients with prior pacemaker implantation complete device interrogation was performed before the procedure and all arrhythmic events were recorded.

All data were collected in a database that was updated after every FU visit; causes and timing of syncope or sudden death were also recorded.

For the purpose of this study we retrospectively analyzed the recorded data in order to access the incidence, time occurrence, predictive factors and causes of sudden cardiac death in patients after TAVI.

Results

Overall

A total of 146 consecutive patients were enrolled, (mean age 79±9y.o., 59% male, mean EUROSCORE 26,12±11,9, mean ST-Score 6,54±3,1). Seventeen patients (12%) had a previous pacemaker implantation. Ninety-one (62,3%) patients had coronary artery disease; in seventy (77%) of them revascularization was performed before o during the procedure. Baseline demographic and clinical, characteristics of the study population are reported in Table 1. Baseline ECG characteristics are reported in table 2.

Electrocardiographic Characteristics And Permanent Pacemaker Implantation (PPM)

Immediately after the procedure new onset left bandle branch block (LBBB) was observed in 39 (27%) patients. Complete atrio-ventricular block (AVB) was present in 14 patients (9,6%). Detailed electrocardiographic characteristics are reported in Table 2. Twenty five patients underwent permanent pacemaker implantation (17,1%) before discharge (11% in the Edward Sapien group and 21% in the Corvalve group); due to 3rd degree AV block (56%), right bandle branch block (RBBB) and LBBB alternans (8%), 2nd degree AV block with 2: 1 conduction (4%) and 1rst degree AV block and LBBB (32%). No syncope was recorded during hospitalization. In all patients with

Baseline demographic and clinical, characteristics of the Table 1: study population. Mean age 79±9 Ejection fraction (%) 51.9±12.14 Gender (male) 59% 26,12±11,9 Euroscore 6.54 + 3.1Atrial fibrillation 45% STScore mortality NYHA Hypertension 3.12±0.56 66.4% Diabetes 47% Corvalve 82.9% Coronary artery 62,3% In hospital stay (days) 5,4±2,2 disease PPM before 12% PPM post 25/146 (17.1%)

Case Report

complete AV block immediately after TAVI or during hospitalization a pacemaker was implanted. Regarding 1rst degree AV block associated with new onset LBBB, only patients who presented progression of the conduction disturbance (progressive prolongation of the PR interval and increase of the duration of the QRS), recorder during hospital stay by ECG, received a pacemaker.

Syncope, Mortality And Ventricular Arrhythmias

In hospital mortality related to procedural complications, heart failure or multi organ failure was 4%, 30 day mortality was 2%. One sudden death occurred and 3 patients experienced syncope (excluding in hospital mortality).

Syncope was present in three patients, in one (with new onset LBBB after the procedure) due 3rd degree AV block so a PPM was implanted while in two other patients, with previous PPM, syncope was related to ventricular tachyarrhythmia's. [fig 1,2] Sudden death (during sleep) due to ventricular fibrillation occurred in one patient implanted with PPM after TAVI due to 1rst degree AV block and LBBB. Intracavitary eletrogram (IEGM) and coronary angiogram of the patient is available in figure 3 A, B. All patients were affected by coronary artery disease and all procedures were performed by transfemoral approach. As reported in table 3 patients were assessed prior to the TAVI procedure and treated if necessary after adequate evaluation with coronary angiogram and/or stress test. Demographic and clinical characteristics of patients are reported in table 3.

All events were recorded in less than 30 days after the procedure. Sudden cardiac death was at day 21, 3rd degree AV block at day 19, and NSVT with syncope at 25 and day 29 respectively.

Discussion

Transcatheter aortic valve implantation (TAVI) for severe aortic stenosis (AS) has become a valid alternative for the treatment of older patients with severe co-morbidities.^{2,3,8-11} Despite the obvious benefits of the procedure, the experience acquired overtime revealed several intra and post procedural complications that can neutralize the potential benefits of valve replacement.² Other than possible vascular, kidney, coronary complications, stroke and paravalvular regurgitation¹² major interest is the occurrence of post-operative conduc-

 Table 2:
 Baseline ECG characteristics before and after TAVI.

	Before TAVI	After TAVI
Sinus Rhythm	112/146 (77%)	99/146 (68%)
Atrial fibrillation	17/146 (12%)	16/146 (11%)
1rst degree AV block	21/146 (14%)	30/146 (20,5%)
RBBB	11/146 (7,5%)	9/146 (6%)
LBBB	8/146 (5,5%)	39/146 (27%)
LAH	32/146 (22%)	19/146 (13%)
Total AV block	0	14/146 (9,6%)
Previous PPM	17/146 (12%)	

Demographic and clinical characteristics of patients with Table 3: syncope and sudden death.

patient	Symptom	Age	Gender	CAD	Treated CAD	EF %	ECG
1	syncope	65	Male	+	-	50	3rd degree AVB
2	syncope	75	male	+	+	50%	VT
3	SCD	78	Female	+	-	35-40%	VF
4	syncope	80	male	+	+	35%	VT

tion disorders that require permanent pacemaker implantation and the occurrence of new left bundle branch block. Several studies have reported rates of new PPM up to 40% of cases^{2,3,13-14} with the occurrence being 6-7 times higher in case of the Medtronic CoreValve system (MCV; Medtronic) implantation.¹⁵ The variations of rate implantation most probably depends on different practice and threshold among physicians.¹²

The new onset LBBB is a common complication after both surgical aortic valve replacement (SAVR) and TAVI procedure. Regarding LBBB after SAVR this is known to be associated with higher rates of complete AV block, syncope and cardiac arrest at long term and it is mostly related to surgical trauma of the conduction system during debridement of the calcified annulus.¹⁶⁻¹⁹ Less is known about the potential effects of new LBBB in the TAVI population. The course of post TAVI-LBBB is not well known, and several cases of spontaneous resolution have been described.²⁰⁻²³ Recent reports correlate new LBBB with increased mortality²⁴ mostly related to progression to high-degree atrioventicular conduction disorders, progression of heart failure due to left ventricular remodeling induced by abnormal activation contraction pattern.²⁴⁻²⁵ In such causes some benefit may be provided by cardiac resynchronization therapy (CRT-D).²⁰

A gray zone of the TAVI population regards sudden cardiac death (SCD). The rate of SCD varies between 4% and 17%.²⁶⁻²⁸ but in most cases is related to bradyarrhythmias. Potential proarrhythmic effect of the procedure may be correlated more than complete heart block. In fact, as reported ventricular arrhythmias occurred in patients without elevated risk of sudden death according to current knowledge. The potential mechanism is not clear, QT dispersion or alterated ventricular activation sequence as reported in patients with CRT-D³⁰⁻³¹ and PPM³² cannot be implicated in such patients. Potential increase of adrenergic tone due to improvement of functional class or myocardial ischemia due to increased demand or potential coronary ostia obstruction by the frame may be implicated.

In our cohort three patients experienced ventricular tachycardia/ ventricular fibrillation after transcatheter aortic valve implantation. All patients were affected by ischemic heart disease with borderline ejection fraction. A probable proarrhythmic effect of new onset LBBB or aggressive left ventricular remodelling after TAVI can be





assumed to be the trigger of ventricular arrhythmias in these patients with borderline left ventricular function. In addition, the time of occurrence of events in less than 30 days after the procedure collocates these events as procedural mortality, thus directly related to the procedure. In such scenario the reports of sudden death, reported to be up to 17%,26-28 should be reconsidered. The exact mechanism of such event cannot be clearly identified as, by definition is sudden and abrupt and in most cases there is no clear documentation of the event, only the result is assessed, but is clearly related to the new hemodynamics created in the left ventricle after the procedure. Regarding late occurrence of complete AV block this event is also less predictable as the TAVI population is older and most likely to receive a pacemaker for degenerative conduction disturbance irrelevant with the TAVI and the occurrence of such event shortly after the procedure might be circumstantial.

In any case, the occurrence of SCD in the TAVI population is a devastating event especially after the high expectations regarding lower rate of morbidity and mortality related to the procedure. Appropriate selection of patients, adequate medical treatment and more studies are necessary in order to completely understand such mechanism and potentially reduce its occurrence.

Conclusions

The TAVI is a valid alternative to SAVR in high risk or inoperable patients. Nevertheless, potential post procedural complications, especially arrhythmic, may reduce its potential benefit. Permanent pacemaker implantation is a frequently reported complication and even if many risk factors are reported none of them is currently considered to be a gold standard. Less is known about sudden cardiac death related to ventricular arrhythmias after TAVI. More research is necessary in order to improve outcomes of the TAVI population.

References

- 1. Clinical outcome and predictors for adverse events after transcatheter aortic valve implantation with the use of different devices and access routes. Wenaweser P, Pilgrim T, Roth N, Kadner A, Stortecky S, Kalesan B, Meuli F, Büllesfeld L, Khattab AA, Huber C, Eberle B, Erdös G, Meier B, Jüni P, Carrel T, Windecker S. Am Heart J. 2011 Jun;161(6):1114-24. doi: 10.1016/j.ahj.2011.01.025. Epub 2011 May 11.
- 2. Thomas Schroeter, Axel Linke, Martin Haensig, Denis R. Merk, Michael A. Borger, Friedrich W. Mohr, and Gerhard Schuler. Predictors of permanent pacemaker implantation after Medtronic CoreValve bioprosthesis implantation. Europace (2012) 14, 1759-1763 doi:10.1093/europace/eus191.
- Spargias K, Toutouzas K, Chrissoheris M, Synetos A, Halapas A, Paizis I, Latsios 3. G, Stathogiannis K, Papametzelopoulos S, Zanos S, Pavlides G, Zacharoulis A, Antoniades A, Stefanadis The Athens TAVR Registry of newer generation

transfemoral aortic valves: 30-day outcomes. C. Hellenic J Cardiol. 2013 Jan-Feb;54(1):18-24.

- Guidelines on the management of valvular heart disease (version 2012). The Joint Task Force on the Management of Valvular Heart Disease of the European Society of Cardiology (ESC) and the European Association for Cardio-Thoracic Surgery (EACTS). European Heart Journal (2012) 33, 2451–2496
- Roques F, Michel P, Goldstone AR, Nashef SA. The logistic EuroSCORE. Eur Heart J. 2003 May;24(9):882-3
- Nashef SA, Roques F, Hammill BG, Peterson ED, Michel P, Grover FL, Wyse RK, Ferguson TB. Validation of European System for Cardiac Operative Risk Evaluation (EuroSCORE) in North American cardiac surgery. Eur J Cardiothorac Surg. 2002 Jul;22(1):101-5.
- http://www.sts.org/quality-research-patient-safety/quality/risk-calculator-andmodels.
- Cribier A, Eltchaninoff H, Tron C, Bauer F, Agatiello C, Sebagh L et al. Early experience with percutaneous transcatheter implantation of heart valve prosthesis for the treatment of end-stage inoperable patients with calcific aortic stenosis. J Am Coll Cardiol 2004;43:698–703.
- Cribier A, Eltchaninoff H, Tron C, Bauer F, Agatiello C, Nercolini D et al. Treatment of calcific aortich stenosis with the percutaneous heart valve: mid-term follow-up from the initial feasibility studies: the French experience. J Am Coll Cardiol 2006;47:1214–23.
- 10. Grube E, Schuler G, Buellesfeld L, Gerckens U, Linke A, Wenaweser P et al. Percutaneous aortic valve replacement for severe aortic stenosis in high-risk patients using the second- and current third-generation self-expanding CoreValve Prosthesis. J Am Coll Cardiol 2007;50:69–76.
- Webb JG, Chandavimol M, Thompson CR, Ricci DR, Carere RG, Munt BI et al. Percutaneous aortic valve implantation retrograde from the femoral artery. Circulation 2006;113:842–50.
- Philippe Ge'ne'reux, Stuart J. Head, David A. Wood, Susheel K. Kodali, Mathew R. Williams, Jean-Michel Paradis, Marco Spaziano, A. Pieter Kappetein, John G. Webb, Alain Cribier, and Martin B. Leon. Transcatheter aortic valve implantation: 10-year anniversary. Part II: clinical implications. European Heart Journal (2012) 33, 2399–2402 doi:10.1093/eurheartj/ehs223.
- Piazza N, de Jaegere P, Schultz C, Becker AE, Serruys PW, Anderson RH. Anatomy of the aortic valvar complex and its implications for transcatheter implantation of the aortic valve. Circ Cardiovasc Interv 2008;1:74–81.
- Bleiziffer S, Ruge H, Horer J, Mazzitelli D, Schreiber C, Hutter A et al. Postoperative AV block and pacemaker implantation after transcatheter aortic valve implantation. Circulation 2008;118:S904–5.
- Michael Liang, Gerard Devlin, Sanjeevan Pasupati. The Incidence of Transcatheter Aortic Valve Implantation-Related Heart Block in Self-Expandable Medtronic CoreValve and Balloon-Expandable Edwards Valves. J INVASIVE CARDIOL 2012;24(4):173-176
- I. Akin, S. Kische, H. Schneider, A. Liebold, J. Ortak, D. Ba¨nsch, T. C. Rehders, O. Thiele, R. Schneider, G. Kundt, H. Krenz, T. Chatterjee, C. A. Nienaber, H. Ince. Surface and intracardiac ECG for discriminating conduction disorders after CoreValve implantation. Clin Res Cardiol (2012) 101:357–364
- El-Khally Z, Thibault B, Stainloae C, Theroux P, Dubuc M, Roy D, Guerra P, Macle L, Talajic M (2004) Prognostic significance of newly acquired bundle branch block after aortic valve replacement. Am J Cardiol 94:1008–1011
- Marchenese K, Schenk EA (1972) Atrioventricular conduction system lesion following cardiac valve replacement. Circulation 45–46(Suppl. II):II-188
- Copeland JG, Griepp RB, Stinson EB, Shumway NE (1977) Long-term followup after isolated aortic valve replacement. J Thorac Cardiovasc Surg 74:875–889
- 20. Kentaro MeguroNicolas Lellouche, MD, PhD, Emmanuel Teiger, MD, PhDCardiac Resynchronization Therapy Improved Heart Failure After Left Bundle Branch Block During Transcatheter Aortic Valve Implantation J

INVASIVE CARDIOL 2012;24:132-133

- Gutierrez M, Rodes-Cabau J, Bagur R, et al. Electrocardiographic changes and clinical outcomes after transapical aortic valve implantation. Am Heart J. 2009;158(2):302-308.
- 22. Calvi V, Puzzangara E, Pruiti GP, et al. Early conduction disorders following percutaneous aortic valve replacement. Pacing Clin Electrophysiol. 2009;32(Suppl 1):S126-S130.
- Aktug O, Dohmen G, Brehmer K, et al. Incidence and predictors of left bundle branch block after transcatheter aortic valve implantation. Int J Cardiol. 2011 Mar 30 [Epub ahead of print].
- 24. Patrick Houthuizen, Leen A.F.M. Van Garsse, Thomas T. Poels; Peter de Jaegere, Robert M.A. van der Boon; Ben M. Swinkels, Jurrie n M. ten Berg, Frank van der Kley, Martin J. Schalij, Jan Baan, Jr, Ricardo Cocchieri, Guus R.G. Brueren, Albert H.M. van Straten, Peter den Heijer, Mohamed Bentala, Vincent van Ommen, Jolanda Kluin, Pieter R. Stella, Martin H. Prins, Jos G. Maessen, Frits W. Prinzen. Left Bundle-Branch Block Induced by Transcatheter Aortic Valve Implantation Increases Risk of Death. Circulation. 2012;126:720-728.
- 25. Kourouklis SP, Manolis AG. Cardiac resynchronization therapy in asymptomatic and mildly symptomatic chronic systolic heart failure: a new era of systolic heart failure management? Hellenic J Cardiol. 2008 Sep-Oct;49(5):349-51. Review.
- 26. Johan M. Bosmansa,*, Joe"lle Keferb, Bernard De Bruynec, Paul Herijgersd, Christophe Duboise, Victor Legrandf, Stephan Verheyeg, Inez Rodrigusa, for the Belgian TAVI Registry ParticipantsProcedural, 30-day and one year outcome following CoreValve or Edwards transcatheter aortic valve implantation: results of the Belgian national registry. Interactive CardioVascular and Thoracic Surgery 12 (2011) 762–767
- 27. Gian Paolo Ussia, Marco Barbanti, Anna Sonia Petronio, Giuseppe Tarantini, Federica Ettori, Antonio Colombo, Roberto Violini, Angelo Ramondo, Gennaro Santoro, Silvio Klugmann, Francesco Bedogni11, Francesco Maisano, Antonio Marzocchi, Arnaldo Poli, Marco De Carlo, Massimo Napodano, Claudia Fiorina, Federico De Marco, David Antoniucci, Emanuela de Cillis, Davide Capodanno, Corrado Tamburino, on behalf of the CoreValve Italian Registry Investigators. Transcatheter aortic valve implantation: 3-year outcomes of self-expanding CoreValve prosthesis European Heart Journal (2012) 33, 969–976.
- Moreno R, Calvo L, Salinas P, Dobarro D, Santiago JV, Sanchez-Recalde A, Galeote G, Riera L, Moreno-Gomez I, Mesa J, Plaza I, Lopez-Sendon J. Causes of peri-operative mortality after transcatheter aortic valve implantation: a pooled analysis of 12 studies and 1223 patients. J Invasive Cardiol. 2011 May;23(5):180-4.
- 29. Arsenos P, Gatzoulis K, Dilaveris P, Manis G, Tsiachris D, Archontakis S, Vouliotis AH, Sideris S, Stefanadis C. Arrhythmic sudden cardiac death: substrate, mechanisms and current risk stratification strategies for the post-myocardial infarction patient. Hellenic J Cardiol. 2013 Jul-Aug;54(4):301-15.
- Indranill Basu Ray, Lahn Fendelander, Jagmeet P. Singh. Cardiac Resynchronization Therapy and its Potential Proarrhythmic Effect. Clin. Cardiol. 30, 498–502 (2007)
- Osama Tayeh, Waleed Farouk, Abdo ElAzab, Hassan Khald , Antonio Curnis. Potential pro-arrhythmic effect of cardiac resynchronization therapy. J Saudi Heart Assoc 2013;25:181–189.
- Wataru Shimizu. Proarrhythmic effect of altered ventricular activation sequence in patients with permanent pacemaker. Heart Rhythm. 2007 Dec;4(12):1487-8.