openheart How much does a heart valve implantation cost and what are the health care costs afterwards?

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ABSTRACT

Objective In the era of limited healthcare budgets, healthcare costs of heart valve implantations need to be considered to inform cost-effectiveness analyses. We aimed to provide age group-specific costs estimates of heart valve implantations, related complications and other healthcare utilisation following the intervention.

Methods We performed retrospective analyses of healthcare costs of patients who had undergone heart valve implantations in 2010–2013 and controls using claims data from Dutch health insurers. Heart valve implantations included surgical valve replacement and transcatheter valve implantation in all heart valve positions. Patients were divided in four age groups. Control groups were created by taking random samples of the Dutch population stratified by age, gender, socioeconomic status and comorbidities. We applied non-parametric bootstrapping to address uncertainty of the cost estimates. The association of patient and intervention characteristics with costs was determined by (multilevel) generalised linear models.

Results The baseline characteristics of 18 903 patients and 188 925 controls were comparable. The annual healthcare costs were substantially higher for surgical heart valve replacement patients than for controls, especially in the year of heart valve implantation. Factors associated with increased annual healthcare costs for patients were older age, female gender, comorbidities, low socioeconomic status and complications.

Conclusions We provided a comprehensive overview of age group-specific incidence of heart valve implantations, subsequent survival and complications as well as associated healthcare costs of all patients in the Netherlands. Our results provide real-world costs estimates that can be used as a benchmark for costs of future innovative heart valve implantations.

INTRODUCTION

Heart valve disease has a profound impact on the use and costs of healthcare. This impact is even greater for heart valve disease than for coronary heart disease, despite the higher prevalence of coronary heart disease.¹ In developed countries, the prevalence of heart valve disease is 2.5%, and this prevalence is the highest in patients aged ≥75 years (13.3%).² Due to an ageing population,

Key questions

What is already known about this subject?

- Due to the ageing of the population, the number of patients with heart valve disease is expected to increase as is the number of heart valve replacements. The scarcity of resources for healthcare requires an analysis of the impact of this development on costs and cost-effectiveness.
- Comprehensive and comparable cost estimates for the entire target population and its subgroups are lacking. The costs of surgical valve replacement and transcatheter valve implantation and subsequent healthcare use have only been estimated for specific risk groups such as elderly high or intermediate surgical risk patients. Moreover, these estimates include only short-term costs.

What does this study add?

- This study adds valuable real-world, age group-specific cost estimates of all costs associated with heart valve implantations (in all valve positions), including the costs of heart valve implantations itself, complications and healthcare use in the years following the heart valve implantation. Furthermore, this study provides a unique insight in the differences in incidence, health outcomes and (predictors of the) associated healthcare costs of heart valve implantations between patients with different ages.

How might this impact on clinical practice?

- Cost-effectiveness is increasingly becoming an important hurdle for introduction of new interventions in clinical practice. Our results can be used as a benchmark for the costs of new technologies in the field of heart valve implantations that will be introduced in clinical practice in the future, such as tissue-engineered heart valves. To estimate the cost-effectiveness, healthcare decision makers need information about the cost-effectiveness in the real world. Our results provide valuable input for the costs in cost-effectiveness analyses of heart valve implantations based on data generated in routine care instead of under experimental conditions.

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the number of patients with heart valve disease requiring valve replacement is expected to rise, reaching more than 800 000 annual procedures worldwide by 2050.³ As a result, healthcare expenditures and societal burden of heart valve disease will increase.

In patients with severe heart valve disease, replacement of the native valve with a heart valve substitute may be required. There are different effective options, but every heart valve substitute type also has its limitations.⁴ In the future, these limitations may be reduced with the many emerging technologies in the field of heart valve interventions, such as tissue-engineered heart valves and less invasive implantation methods. Before these new technologies can be introduced in clinical practice, it is important to establish that they are effective and cost-effective, considering the scarcity of resources for healthcare. To determine whether a new intervention is cost-effective, the costs and effects of the new intervention need to be compared with current care. Since the choice of heart valve implantation and its outcomes differs substantially among age groups,⁵ it is likely that healthcare costs will also be influenced by patients' age. Hence, having robust age group-specific estimates of the costs of current care is important.

The objective of this study was to estimate the costs of currently used heart valve implantations and also costs of complications as well as healthcare use outside hospitals in the years after heart valve implantations in different age groups. We estimated these costs by retrospective analyses of health insurance claims of patients who had undergone heart valve implantations. This provides valuable real-world age group-specific cost estimates of all costs associated with heart valve implantations, in comparison with previous studies that focused on costs of the heart valve implantation and short-term follow-up in specific age groups only.^{6,7}

METHODS

Patients

We used health insurance claims databases ('Vektis') that contain the healthcare expenditures of all the insured in the Netherlands, which is 99% of all Dutch residents (± 17 million people). Patients were selected using Diagnosis Related Group (DRG) codes of heart valve implantations (online supplementary table S1). We could distinguish DRG codes for isolated or multiple surgical heart valve replacement (SVR) and transcatheter heart valve implantation (TVI) in every heart valve position with or without concomitant procedures. In the pulmonary and tricuspid valve position, DRG codes did not distinguish between valve repair and replacement; therefore, patients who have undergone repairs in these valve positions are also included in our patient population.

We used data from Vektis for the years 2010–2013. Data before 2010 are generally considered less valid. On the date of data extraction for this study (January 2016), data from the years 2014 and further were incomplete due

to time lags in administrations. Before 2013, there was no specific DRG code for TVI; therefore, we could only include TVI patients in the year 2013.

Controls

To calculate the excess healthcare costs due to the heart valve implantation, we compared annual healthcare costs of the patient group with a control group. The control group was created by stratified sampling from the remainder of the Dutch insured population in the Vektis databases to ensure that the distributions of person characteristics across the strata were similar to that in the patient population. The control group was 10 times as large as the patient population. Strata were based on age class, gender, socioeconomic status (SES) and comorbidities. Age was divided into nine age classes: 0–1, 2–18, 19–30, 31–40, 41–50, 51–60, 61–70, 71–80 and >80 years. SES was based on status scores reflecting the SES of a district based on characteristics of its residents: education, income and position on the labour market.⁸ The status scores were divided in four groups based on percentiles, with lower percentiles representing lower SES. Comorbidities were based on Pharmacy Cost Groups, which is an outpatient morbidity measure based on prior use of prescribed drugs as marker for chronic conditions.⁹ The strata used for comorbidities differed per age class because of differences in prevalence of comorbidities (see online supplement).

Healthcare costs

Patients were followed over time from the opening date of the heart valve implantation DRG in the financial administrative system until death or until 31 December 2013, whichever occurred first, to assess whether they experienced complications after the heart valve implantation and to collect their other healthcare costs besides the costs of heart valve implantation itself.

For every patient, the costs of the initial heart valve implantation and the healthcare costs during the first postintervention year and subsequent (max. 4) years were determined. Costs were defined as expenditures reimbursed by health insurers and expressed in euros (€). The procedure costs included the costs of the DRG and intensive care unit (ICU) stay. We assumed that the costs of ICU stay in the first postintervention year excluding the ICU costs related to specific complications were related to the heart valve implantation.

Complications were extracted from the Vektis database using DRG codes for treatment of complications (online supplementary table S2). The complications of interest were based on the conceptual model we have developed previously.⁵ The following complications were available in the Vektis database: acute kidney injury, atrial fibrillation (AF), stroke, transient ischaemic attack, prosthetic valve endocarditis (conservative treatment), myocardial infarction, pacemaker implantation and reintervention (redo heart valve implantation). For every complication, costs (DRG+ICU costs), proportions of

patients admitted to the ICU and number of ICU days were determined.

For both patients and controls, annual healthcare costs were determined. The starting point of the calculation of annual healthcare costs of patients is the quarter in which the heart valve implantation was performed. The annual healthcare costs were classified into costs of general practitioners, specialised medical care (both inpatient and outpatient care, including costs of DRGs, ICU stay, medicines on the expensive drugs list, primary care diagnostics and other costs), pharmaceuticals, paramedical care, patient transport, home care, nursing homes and geriatric rehabilitation care. Cost of medicines on the expensive drug list, home care, nursing homes and geriatric rehabilitation care were only available in the Vektis database from the year 2012 onwards.

Statistical analysis

The statistical analyses were performed with SAS V.9.4 using SAS Enterprise Guide V.7.1.

Descriptive analyses

Patients and controls were divided in four age groups: children (0–18 years), young adults (19–60 years), middle aged (61–70 years) and elderly patients (>70 years). For every age group, we performed descriptive analyses of person and intervention characteristics, occurrence of complications and healthcare costs. Continuous variables were depicted as means and standard deviation (SD) or 95% confidence interval (CI) and discrete variables as counts and proportions. We assessed survival after SVR and TVI using Kaplan-Meier estimates. The difference between mean healthcare costs of patients and controls was calculated; mean costs of controls were based on the same calendar years as patients (eg, for patients postintervention year 2 could be between 2011 and 2013; therefore, these costs were compared with the average costs of controls in 2011–2013). As the cost data were skewed, non-parametric bootstrapping (2000 replications) was used to address uncertainty (with 95% CI based on 2.5th and 97.5th percentile) in the annual healthcare costs. To be able to report annual healthcare costs, only patients who were followed the entire year of interest (including patients who died during this year) were included in these analyses.

Association analyses

To estimate the association between healthcare costs and patient and intervention characteristics, we developed (multilevel) generalised linear models ((M)GLM) for intervention, complication and annual healthcare costs. For annual healthcare costs of patients after heart valve implantations, we estimated a MGLM for children and for adults with normal distributions and identity links and with several observation periods per patient. All patients with at least 1-year complete follow-up were included. We excluded patients with incomplete follow-up in the first post intervention year (except for patients who died) to avoid overestimation of costs per day in post intervention

year one due to the high costs in the period after the intervention. To correct for differences in total duration of follow-up, the total healthcare costs were divided by the follow-up duration to estimate average costs per day during the specific year. The independent variables included in the MGLMs for children and adults were time (ie, intervention period (no defined length; includes costs of heart valve implantation and ICU stay), remaining postintervention year 1 and postintervention years 2, 3 and 4), gender and SES. In addition, the model of adults included age groups, comorbidities, mortality and complications. Mortality and the occurrence of complications were time dependent. Comorbidities, mortality and the occurrence of complications were not included in the model for children because there were no children with comorbidities and only a small proportion of children who experienced complications (including mortality). Details about the GLMs in which the costs of interventions and complications were analysed separately can be found in the online supplementary material.

RESULTS

Study population

In total, we included 18903 patients (SVR: n=17991, TVI: n=912) and 188925 controls. The baseline characteristics of patients and controls were comparable (table 1). The mean (median) follow-up was 1.9 (1.9) and 0.4 (0.4) years for SVR and TVI patients, respectively. Figure 1 illustrates the Kaplan-Meier survival curves after SVR and TVI. During follow-up, 41 children (10.0%), 841 young adults (26.2%), 1424 middle-aged patients (29.5%) and 3036 elderly patients (31.9%) experienced one or more complications after SVR. After TVI, six young adults (18.2%), two middle aged (2.9%) and 157 elderly patients (19.4%) experienced complications.

Table 2 presents the occurrence of complications per postintervention year, the proportion of patients with ICU stay and the mean length of ICU stay after complications.

Healthcare costs: descriptive analyses

Table 3 and figure 2 summarise the costs of heart valve implantations, complications and total annual healthcare costs during the first three postintervention years divided by age group. These costs could not be determined for TVI patients because their follow-up was less than 1 year. The annual healthcare costs of patients were substantially higher than the costs of controls in all age groups, especially in the year of implantation (figure 2; children €11 766 vs €796, young adults €15 060 vs €2944, middle aged €16 104 vs €4612 and elderly €18 255 vs €9236). The patients' annual healthcare costs were substantially higher than controls for most types of healthcare across all age groups (online supplementary tables S3–S6). However, middle-aged and elderly patients had substantially lower costs of nursing homes than controls in postintervention year 1 (€866 vs €2761). The costs of nursing homes remained substantially lower in elderly patients in postintervention

Table 1 Person and intervention characteristics of patients and controls divided by age group

	Children (0–18 years)			Young adults (19–60 years)			Middle aged (61–70 years)			Elderly (>70 years)		
	Patients	Controls	SVR	Patients	SVR	TVI	Patients	SVR	TVI	Patients	SVR	TVI
Number of persons,* n (%)	411	4170		3213 (99.0)	33 (1.0)	33960	4826 (98.6)	70 (1.4)	50440	9527 (92.2)	809 (7.8)	10,0355
Follow-up in years,† mean±SD	2.2±1.2	<4.0		2.0±1.2	0.5±0.3	<4.0	1.9±1.2	0.4±0.3	<4.0	1.8±1.2	0.4±0.3	<4.0
Age at time of intervention, mean±SD, range	5.5±6.2 0–18	– 0–18		50.3±9.4 19–60	39.4±14.9 19–60	– 19–60	66.0±2.8 61–70	66.9±2.1 62–70	– 61–70	77.6±4.4 71–94	81.9±4.9 71–95	– >70
Male, n (%)	245 (59.6)	2490 (59.7)		2140 (66.6)	23 (69.7)	22730 (66.9)	3261 (67.6)	43 (61.4)	33770 (67.0)	5075 (53.3)	387 (47.8)	47 580 (47.4)
SES,‡ n (%)												
0–20	72 (17.5)	720 (17.3)		737 (22.9)	4 (12.1)	7650 (22.5)	987 (20.5)	23 (32.9)	10530 (20.9)	2042 (21.4)	202 (25.0)	21 780 (21.7)
21–40	80 (19.5)	810 (19.4)		681 (21.2)	6 (18.2)	7250 (21.3)	994 (20.6)	12 (17.1)	10310 (20.4)	2031 (21.3)	161 (19.9)	21 280 (21.2)
41–70	124 (30.2)	1250 (30.0)		897 (27.9)	11 (33.3)	9570 (28.2)	1587 (32.9)	16 (22.9)	16490 (32.7)	2984 (31.3)	228 (28.2)	31 180 (31.1)
71–100	135 (32.8)	1390 (33.3)		888 (27.6)	12 (36.4)	9380 (27.6)	1256 (26.0)	19 (27.1)	13090 (26.0)	2469 (25.9)	218 (26.9)	26 110 (26.0)
Comorbidities, n (%)												
COPD, DM, kidney disease and/or HF				780 (24.3)	9 (27.3)	8510 (25.1)	2086 (43.2)	47 (67.1)	22 160 (43.9)	5041 (52.9)	518 (64.0)	54 145 (54.0)
Hypertension				1106 (34.4)	6 (18.2)	11600 (34.2)	1659 (34.4)	17 (24.3)	17 330 (34.4)	3036 (31.9)	199 (24.6)	31 310 (31.2)
Other comorbidities				293 (9.1)	3 (9.1)	3170 (9.3)	416 (8.6)	3 (4.3)	4210 (8.3)	575 (6.0)	36 (4.4)	5880 (5.9)
No comorbidities	411 (100.0)	4170 (100.0)		1034 (32.2)	15 (45.5)	10680 (31.4)	665 (13.8)	3 (4.3)	6740 (13.4)	875 (9.2)	56 (6.9)	9020 (9.0)
Valve position, n (%)												
Aortic	29 (7.1)			2460 (76.6)	6 (18.2)		4133 (85.6)	25 (35.7)		8578 (90.0)	395 (48.8)	
Pulmonary	338 (82.2)			115 (3.6)			2 (0.0)			2 (0.0)		
Mitral	19 (4.6)			431 (13.4)	2 (6.1)		484 (10.0)	1 (1.4)		652 (6.8)		
Tricuspid	23 (5.6)			65 (2.0)			34 (0.7)			36 (0.4)		
Aortic and mitral	2 (0.5)			142 (4.4)			173 (3.6)			259 (2.7)		
Unknown					25 (75.8)			44 (62.9)			414 (51.2)	
Concomitant procedures, n (%)												
No concomitant procedures	214 (52.1)			1924 (59.9)	33 (100.0)		2485 (51.5)	70 (100.0)		4735 (49.7)	807 (99.8)	
CABG				407 (12.7)			1364 (28.3)			3409 (35.8)	2 (0.2)	
Valve repair	1 (0.2)			253 (7.9)			344 (7.1)			643 (6.7)		
Maze+CABG or valve repair				58 (1.8)			212 (4.4)			418 (4.4)		
Bentall	8 (1.9)			303 (9.4)			162 (3.4)			94 (1.0)		
Aortic ascendens				173 (5.4)			161 (3.3)			114 (1.2)		
Tetralogy of Fallot	187 (45.5)			6 (0.2)								
Aortic ascendens+valve repair				27 (0.8)			31 (0.6)			36 (0.4)		
HOCM				11 (0.3)			14 (0.3)			52 (0.5)		

Continued

Table 1 Continued	Children (0–18 years)		Young adults (19–60 years)		Middle aged (61–70 years)		Elderly (>70 years)	
	Patients	Controls	Patients	TVI	Controls	Patients	SVR	TVI
	SVR		SVR					
Aortic root			24 (0.7)		19 (0.4)		8 (0.1)	
Aortic root+CABG			17 (0.5)		17 (0.4)		13 (0.1)	
Left ventricle repair	1 (0.2)		10 (0.3)		17 (0.4)		5 (0.1)	

*Fourteen patients excluded because their age was unknown.

†Control group includes patients who died during the study period, but information on number of deaths or follow-up of controls was not available.

‡Higher percentiles represent higher SES.

CABG, concomitant coronary artery bypass grafting; COPD, chronic obstructive pulmonary disease; DM, diabetes mellitus; HF, heart failure; HOCM, hypertrophic obstructive cardiomyopathy; SES, socioeconomic status; SVR, surgical heart valve replacement; TVI, transcatheter heart valve implantation.

years 2 and 3 (€1763 (year 2) and €1990 (year 3) vs €2761 for controls). In addition, costs of home care were lower for elderly patients than controls in the first year after the intervention (€1199 vs €1330).

Healthcare costs: association analyses

Table 4 presents the results of the MGLMs of annual healthcare costs for children and adult SVR patients. Annual healthcare costs for adult SVR patients increased with older age at intervention (on average+€2441 for elderly vs middle aged patients), comorbidities (on average+€6543 for patients with chronic obstructive pulmonary disease, diabetes mellitus, kidney disease and/or HF vs patients without comorbidities) and lower SES (on average+€1160 for patients with lowest vs highest SES). Men had somewhat lower costs than women (€1110 on average). If patients experience a complication, their annual healthcare costs increase on average with €623 after AF to €30 094 after reintervention. If patients die, their costs in the year of death increase on average with €6106. For children, costs were not associated with gender or SES. The results of the GLMs for intervention and complication costs are reported in online supplementary tables S7 and S8.

DISCUSSION

Using the comprehensive Vektis databases, we were able to estimate the real-world age group-specific incidence of heart valve implantations, subsequent survival and complications as well as the associated healthcare costs of all patients in the Netherlands who had undergone a heart valve implantation during our study period. Although the estimates are specific to the Dutch healthcare system, the results regarding differences between age groups, distribution of costs over types of healthcare and associations between patient and intervention characteristics and healthcare costs are also relevant for other countries. Our results can help raise awareness of the costs associated with heart valve implantations among clinicians and healthcare policy makers, which is important in the current era of limited healthcare budgets. However, we want to emphasise that considerations about costs should not play a role in the treatment decision for individual patients. Instead the results can be used as a benchmark in cost-effectiveness analyses for new technologies that will be introduced in clinical practice in the future, such as tissue-engineered heart valves.⁵

This study has shown that Dutch health insurers spent over €120million per year on procedure costs for heart valve implantations, of which 2% is spent on children, 17% on young adults, 26% on middle aged and 56% on elderly patients. Although there were no substantial differences in procedure costs between age groups, the costs of SVR were generally higher in older patients, while costs of TVI were lower for older patients. There was no trend in complication costs in relation to age groups. In addition to procedure costs, patients had excess healthcare costs

after the heart valve implantation compared with controls in almost all types of healthcare. These excess healthcare costs were especially high in the year of heart valve implantation; 41 (children), 14 (young adults), 9 (middle aged) and 5 (elderly) fold higher in patients than controls. In the subsequent postintervention years, however, the excess healthcare costs decreased. This decrease may be explained by survival of the fittest patients.

In contrast to other types of healthcare, the costs of nursing homes were substantially lower for elderly patients than for controls. This may be caused by selection bias of relatively healthy elderly patients for SVR. Patients living in nursing homes may be less likely to undergo heart valve implantation due to other factors

influencing someone's health state, such as frailty or dementia. Since these factors could not be taken into account when defining the control sample, people living in nursing homes may be over-represented in the control group as compared with the patient group.

As expected, older age, female gender, comorbidities, low SES and/or experiencing complications (including death) were associated with higher annual healthcare costs. It should be noted that the aim of this study was to describe and predict costs and that it does not make casual claims. Nevertheless, some explanations for the associations can be considered. The association of lower SES and poor health has also been shown consistently in previous research.¹⁰ The association of gender and

Figure 1 Kaplan-Meier curves of survival after SVR (top) and TVI (bottom) divided by age group. SVR, surgical heart valve replacement; TVI, transcatheter heart valve implantation.

Table 2 Complications after initial heart valve intervention

	No. of patients with complications*, n (% of total patients)				ICU stay, n (% of patients with complication)			
	Year 1		Year 2		Year 3		Year 4	
	SVR	TVI	SVR	TVI	SVR	TVI	SVR	TVI
Children (0–18 years) (n)	411	0	406		405			
All-cause mortality	5 (1.2)		1 (0.2)					
Complications (total)	25 (6.1)		12 (3.0)		4 (1.0)		2 (0.5)	
Acute kidney injury	2 (0.5)							
Atrial fibrillation	4 (1.0)		3 (0.7)		1 (0.2)			
Stroke			1 (0.2)					
TIA								
Endocarditis	4 (1.0)		2 (0.5)					
Myocardial infarction								
Pacemaker implantation	1 (0.2)		1 (0.2)		2 (0.5)			
Reintervention	14 (3.4)		5 (1.2)		2 (0.5)		1 (0.2)	
Young adults (19–60 years) (n)	3213	33	3090	3056	3034		15 (68.2)	2.1±1.1
All-cause mortality	123 (3.8)	2 (6.1)	34 (1.1)	22 (0.7)	5 (0.2)			
Complications (total)	680 (21.2)	6 (18.2)	189 (6.1)	98 (3.2)	37 (1.2)	3 (50.0)		
Acute kidney injury	26 (0.8)		6 (0.2)	1 (0.0)	1 (0.0)		6 (18.2)	13.5±18.6
Atrial fibrillation	268 (8.3)	1 (3.0)	63 (2.0)	40 (1.3)	17 (0.6)		8 (2.4)	2.0±2.4
Stroke	154 (4.8)		40 (1.3)	18 (0.6)	4 (0.1)		26 (12.8)	10.4±21.4
TIA	51 (1.6)		29 (0.9)	14 (0.5)	2 (0.1)		1 (1.2)	2.0±
Endocarditis	118 (3.7)	2 (6.1)	37 (1.2)	15 (0.5)	8 (0.3)	1 (50.0)	29 (18.2)	5.0±4.0
Myocardial infarction	25 (0.8)		8 (0.3)	4 (0.1)	1 (0.0)		5 (13.5)	17.8±32.6
Pacemaker implantation	90 (2.8)	2 (6.1)	10 (0.3)	9 (0.3)	1 (0.0)		11 (10.1)	3.7±2.5
Reintervention	39 (1.2)	2 (6.1)	18 (0.6)	6 (0.2)	3 (0.1)	2 (100.0)	41 (65.1)	9.1±26.6
Middle aged (61–70 years) (n)	4826	70	4545	4472	4413			8.5±4.9
All-cause mortality	281 (5.8)	5 (7.1)	73 (1.6)	59 (1.3)	15 (0.3)			
Complications (total)	1167 (24.2)	2 (2.9)	300 (6.6)	176 (3.9)	67 (1.5)	1 (50.0)	153 (10.7)	
Acute kidney injury	57 (1.2)		17 (0.4)	5 (0.1)	2 (0.0)		10 (12.7)	8.3±7.0
Atrial fibrillation	534 (11.1)		138 (3.0)	87 (1.9)	31 (0.7)		10 (1.6)	2.7±2.2
Stroke	259 (5.4)		53 (1.2)	27 (0.6)	11 (0.2)		43 (12.9)	5.8±5.1
TIA	113 (2.3)		41 (0.9)	32 (0.7)	10 (0.2)		8 (4.3)	3.9±2.0
Endocarditis	153 (3.2)		33 (0.7)	17 (0.4)	4 (0.1)		29 (15.7)	7.0±9.4
Myocardial infarction	45 (0.9)		16 (0.4)	9 (0.2)	2 (0.0)		8 (11.4)	7.6±7.4

Continued

Table 2 Continued

	No. of patients with complications*, n (% of total patients)				ICU stay, n (% of patients with complication)			
	Year 1		Year 2		Year 3		Year 4	
	SVR	TVI	SVR	TVI	SVR	TVI	SVR	TVI
Pacemaker implantation	158 (3.3)	1 (1.4)	27 (0.6)		12 (0.3)		7 (0.2)	
Reintervention	49 (1.0)	1 (1.4)	10 (0.2)		5 (0.1)		5 (0.1)	
Elderly (>70years) (n)	9527	809	8580		8315		8146	
All-cause mortality	947 (9.9)	87 (10.8)	265 (3.1)		169 (2.0)		61 (0.7)	
Complications (total)	2436 (25.6)	157 (19.4)	678 (7.9)		426 (5.1)		125 (1.5)	
Acute kidney injury	135 (1.4)	8 (1.0)	14 (0.2)		9 (0.1)		29 (18.4)	15.0±15.5
Atrial fibrillation	994 (10.4)	26 (3.2)	340 (4.0)		211 (2.5)		67 (0.8)	6.9±13.1
Stroke	670 (7.0)	33 (4.1)	155 (1.8)		87 (1.0)		114 (12.9)	6.5±6.4
TIA	220 (2.3)	11 (1.4)	66 (0.8)		52 (0.6)		13 (0.2)	2.8±1.5
Endocarditis	184 (1.9)	8 (1.0)	57 (0.7)		23 (0.3)		9 (0.1)	8.9±9.9
Myocardial infarction	112 (1.2)	5 (0.6)	33 (0.4)		22 (0.3)		7 (0.1)	4.4±4.8
Pacemaker implantation	428 (4.5)	57 (7.0)	67 (0.8)		47 (0.6)		13 (0.2)	4.9±4.6
Reintervention	76 (0.8)	35 (4.3)	9 (0.1)		11 (0.1)		2 (0.0)	7.8±17.8
							54 (55.1)	3.2±5.9

*Early complications are included in intervention DRG and therefore not included here.

DRG, Diagnosis Related Group; ICU, intensive care unit; MI, myocardial infarction; SVR, surgical heart valve replacement; TIA, transient ischaemic attack; TVI, transcatheter heart valve implantation.

Table 3 Costs of initial heart valve intervention and complications

	Children (0–18 years)		Young adults (19–60 years)		Middle aged (61–70 years)		Elderly (>70 years)	
	n	Costs (€, mean, CI)	n	Costs (€, mean, CI)	n	Costs (€, mean, CI)	n	Costs (€, mean, CI)
Intervention (including ICU*)								
SVR (total)	399	21 941 (20 543 to 23 811)	3172	25 050 (24 446 to 25 711)	4727	25 502 (25 054 to 25 988)	9387	25 740 (25 414 to 26 058)
Aortic	29	20 068 (18 843 to 21 279)	2428	23 935 (23 350 to 24 592)	4050	24 553 (24 131 to 25 004)	8448	25 165 (24 845 to 25 482)
Pulmonary	328	21 800 (20 144 to 23 978)	114	19 442 (18 598 to 20 297)	2	14 483 (11 966 to 17 009)	2	23 702 (21 923 to 25 518)
Mitral	17	26 885 (19 920 to 35 138)	427	27 449 (25 691 to 29 491)	475	28 493 (26 779 to 30 386)	646	29 408 (27 634 to 31 510)
Tricuspid	23	22 409 (19 226 to 27 671)	64	33 306 (26 339 to 41 758)	32	26 858 (21 833 to 33 760)	35	23 611 (21 577 to 25 923)
Aortic+mitral	2	25 451 (20 148 to 30 753)	194	37 985 (32 581 to 44 947)	167	39 834 (34 892 to 45 874)	256	35 759 (33 074 to 38 668)
TVI (total)	0	–	29	33 385 (30 842 to 36 490)	64	32 440 (30 860 to 34 142)	744	32 209 (31 582 to 32 883)
Aortic	0	–	6	35 884 (30 552 to 43 785)	21	33 135 (29 843 to 36 786)	366	32 776 (31 812 to 33 842)
Mitral	0	–	2	33 838 (22 789 to 44 831)	1	36 661 (36 661 to 36 661)	0	–
Unknown	0	–	21	32 614 (30 134 to 36 300)	42	32 003 (30 193 to 33 909)	378	31 660 (30 915 to 32 563)
Complications (including ICU)								
Acute kidney injury	2	6007 (2617 to 9407)	36	9061 (5575 to 13 552)	81	8021 (6303 to 9988)	169	9533 (7597 to 11 769)
Atrial fibrillation	11	2702 (717 to 5789)	548	1418 (1295 to 1543)	1087	1229 (1147 to 1313)	2187	1210 (1119 to 1321)
Stroke	1	1418 (1418 to 1418)	257	3264 (2458 to 4197)	403	2627 (2222 to 3129)	1115	3017 (2731 to 3341)
TIA	0	–	104	1213 (990 to 1470)	220	1311 (1122 to 1522)	387	1267 (1155 to 1394)
Endocarditis	6	7971 (3764 to 13 057)	254	7418 (6449 to 8401)	292	7543 (6689 to 8380)	380	8815 (7960 to 9722)
Myocardial infarction	0	–	49	6248 (4264 to 9517)	83	5421 (4524 to 6518)	203	5094 (4509 to 5911)
Pacemaker implantation	4	4884 (2175 to 6621)	118	10 987 (10 403 to 11 525)	209	12 395 (10 875 to 15 179)	619	11 596 (11 348 to 11 853)
Reintervention	22	20 057 (18 326 to 21 784)	70	25 328 (21 590 to 30 273)	72	21 340 (19 249 to 24 120)	135	25 622 (23 138 to 28 862)

*ICU costs during first year minus ICU costs after complications. Only patients with ≥30-days' follow-up or died ≤30 days included to ensure all ICU costs after intervention were considered.

ICU, intensive care unit; SVR, surgical heart valve replacement; TIA, transient ischaemic attack; TVI, transcatheter heart valve implantation.

Figure 2 Annual healthcare costs during the first three postintervention years of surgical valve replacement patients and controls divided by age group.

costs, even after adjusting for comorbidities and complications, is in line with previous research that found that women have higher healthcare costs than men.¹¹ In cardiovascular diseases, this might be due to the different

preoperative risks profiles of women compared with men,^{12 13} which may be caused by delayed presentation or diagnosis of valve problems and/or later referral to cardiothoracic surgery of women.¹² If these different risk

Table 4 Multilevel generalised linear model of total annual healthcare costs after SVR in postintervention years 1–4

Total costs Parameter	Children (0–18 years, n=325)			Adults (≥18 years, n=13944)		
	β	95% CI	P value	β	95% CI	P value
Intercept	16 931	–36 190 to 70 051	0.533	11 338	9906 to 12 770	<0.0001
Time (compared with year 1 excluding intervention costs)						
Intervention period*	21 841	20 857 to 22 825	<0.0001	25 492	25 248 to 25 736	<0.0001
Year 2	–11 519	–67 302 to 44 264	0.686	–2904	–3779 to –2030	<0.0001
Year 3	–14 952	–67 272 to 37 368	0.576	–1862	–3421 to –302	0.019
Year 4	–6170	–64 405 to 52 065	0.836	396	–1627 to 2420	0.701
Death				6106	4784 to 7428	<0.0001
Age (compared with elderly)						
Children (0–18 years)						
Young adults (19–60 years)				–1179	–2290 to –68	0.038
Middle aged (61–70 years)				–2441	–3359 to –1524	<0.0001
Male	1133	–30 369 to 32 635	0.944	–1110	–1911 to –310	0.007
Comorbidity (compared with no comorbidity)						
COPD, DM, kidney disease and/or HF				6543	5328 to 7757	<0.0001
Hypertension				1309	67 to 2550	0.039
Other comorbidities				1990	218 to 3761	0.028
SES† (compared with highest SES: 71–100)						
SES 0–20	8553	–36 202 to 53 308	0.708	1160	34 to 2285	0.044
SES 21–40	2878	–41 065 to 46 821	0.898	301	–823 to 1426	0.599
SES 41–70	2505	–37 038 to 42 048	0.901	887	–128 to 1901	0.087
Complications						
Atrial fibrillation				2985	1673 to 4296	<0.0001
Acute kidney injury				19 639	16 611 to 22 667	<0.0001
Stroke				7755	6181 to 9329	<0.0001
TIA				623	–2157 to 3403	0.661
Endocarditis				21 572	18 999 to 24 144	<0.0001
Myocardial infarction				13 192	9291 to 17 092	<0.0001
Pacemaker implantation				15 947	13 816 to 18 079	<0.0001
Reintervention				30 094	25 455 to 34 733	<0.0001

NB: 3622 SVR and all TVI patients were excluded because follow-up <1 year.

*Includes costs of heart valve implantation and ICU stay but no other costs of the first postintervention year (these are included in the reference group of this variable).

†Higher percentiles represent higher SES.

COPD, chronic Obstructive Pulmonary Disease; DM, diabetes mellitus; HF, heart failure; ICU, intensive care unit; SES, socioeconomic status; SVR, surgical heart valve replacement; TIA, transient ischaemic attack.

profiles result in slower or impaired recovery of women compared with men, this might result in more use of healthcare and thereby higher annual healthcare costs.

Strengths and limitations

An important strength of our study is the use of databases including the health insurance claims of 99% of Dutch residents. Therefore, almost all patients that have undergone heart valve implantations during our study period

were included, and we presented outcomes in a diverse study population that reflects the range and distribution of patients in clinical practice instead of focusing on specific age or risk groups.¹⁴ This resulted in comprehensive analyses of the real-world healthcare costs associated with heart valve implantations with high external validity and generalisability. Since healthcare decision makers need information about the cost-effectiveness in the

real world, our results provide valuable input for the costs in cost-effectiveness analyses based on data generated in routine care instead of under experimental conditions.¹⁴ Furthermore, this study provided a unique insight in the differences in incidence, health outcomes and associated healthcare costs of heart valve implantations of patients with all ages, divided over four informative age groups.

Our study also has some limitations. First, we could not separate our results for different types of valve prostheses (eg, mechanical and biological prostheses). However, although the type of prosthesis has impact on survival and complication rates, we do not expect that the type of prosthesis has a large impact on healthcare costs. Furthermore, since the DRG code for TVI was only available from 2013, the follow-up was too short to estimate annual healthcare costs in postintervention years for these patients. Additionally, since complications were identified using DRG codes, we could only determine the incidence and costs of complications for which patients were treated in the hospital (including outpatient treatment). Furthermore, not all in-hospital complications could be identified because for some complications the costs may be included in the DRG of the initial heart valve implantation instead of a separate DRG. In addition, the exact date of the heart valve implantation was unknown; instead, the opening date of the DRG in the financial administrative system was used as a proxy. It is possible that the heart valve implantation took place a few days/weeks before or after the opening date of the DRG. Furthermore, we could not calculate annual healthcare costs from the exact date of the intervention onwards but only from the quarter in which the intervention took place. Finally, the reported costs are expenditures reimbursed by health insurers based on agreements between healthcare providers and insurers, not actual costs.

CONCLUSION

This study provided a comprehensive overview of age group-specific incidence of heart valve implantations, subsequent survival and complications as well as the associated healthcare costs of all patients who had undergone a heart valve implantation in the Netherlands. We have shown that after heart valve implantation, patients have substantially higher healthcare costs than controls. The costs are higher in patients with comorbidities and patients who have experienced a complication. The costs estimated in this study can be used as a benchmark for future innovative heart valve implantations, such as tissue-engineered heart valves.

Contributors SAH was involved in the study design, data collection, statistical analysis and writing of the manuscript. JAVe was involved in the study design and data collection. LMAG contributed to the statistical analysis. JJMT and MPMHR-vm contributed to the study design, statistical analysis, interpretation of the data and writing of the manuscript. All authors read and approved the final manuscript.

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Correction: How much does a heart valve implantation cost and what are the health care costs afterwards?

Huygens SA, Goossens LMA, van Erkelens JA, *et al.* How much does a heart valve implantation cost and what are the health care costs afterwards? *Open Heart* 2018;5:e000672. doi: 10.1136/openhrt-2017-000672.

There is a mistake in figure 2 and it should be replaced with the following:

Figure 2 Annual healthcare costs during the first three postintervention years of surgical valve replacement patients and controls divided by age group.

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