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A review of health utilities using the EQ-5D in studies of cardiovascular disease

Matthew TD Dyer1,4*, Kimberley A Goldsmith2,3, Linda S Sharples2,3, Martin J Buxton1

Abstract

**Background:** The EQ-5D has been extensively used to assess patient utility in trials of new treatments within the cardiovascular field. The aims of this study were to review evidence of the validity and reliability of the EQ-5D, and to summarise utility scores based on the use of the EQ-5D in clinical trials and in studies of patients with cardiovascular disease.

**Methods:** A structured literature search was conducted using keywords related to cardiovascular disease and EQ-5D. Original research studies of patients with cardiovascular disease that reported EQ-5D results and its measurement properties were included.

**Results:** Of 147 identified papers, 66 met the selection criteria, with 10 studies reporting evidence on validity or reliability and 60 reporting EQ-5D responses (VAS or self-classification). Mean EQ-5D index-based scores ranged from 0.24 (SD 0.39) to 0.90 (SD 0.16), while VAS scores ranged from 37 (SD 21) to 89 (no SD reported). Stratification of EQ-5D index scores by disease severity revealed that scores decreased from a mean of 0.78 (SD 0.18) to 0.51 (SD 0.21) for mild to severe disease in heart failure patients and from 0.80 (SD 0.05) to 0.45 (SD 0.22) for mild to severe disease in angina patients.

**Conclusions:** The published evidence generally supports the validity and reliability of the EQ-5D as an outcome measure within the cardiovascular area. This review provides utility estimates across a range of cardiovascular subgroups and treatments that may be useful for future modelling of utilities and QALYs in economic evaluations within the cardiovascular area.
for heart failure patients and the Canadian Cardiovascular Society (CCS) grading scale for angina pectoris [4,5]. HRQoL measurement in CVD can be assessed using disease-specific instruments such as the Seattle Angina Questionnaire (SAQ); MacNew Heart Disease Health-related Quality of Life Questionnaire; and the Minnesota Living with Heart Failure score (MLHF) [6-8]. These questionnaires are particularly sensitive to changes in aspects of HRQoL directly related to CVD. Alternatively, commonly used generic measures of HRQoL including the SF-6D, Health Utilities Index (HUI) and the EQ-5D have also been used in CVD studies [9-11]. The main advantages of such generic multi-attribute health state classifiers are that they allow the calculation of Quality adjusted life years (QALYs) within cost-utility analyses as well as allowing comparison of HRQoL across different conditions and against age-sex matched population norms.

Among the available generic measures, the EQ-5D has gained widespread use due to its simplicity to administer, score and interpret. It also imposes minimal burden on the respondent as it is a brief, simple measure for patients to understand and to complete. The index-based score is generated by applying societal preference weights to the health state classification completed by the patient that consists of five dimensions (mobility, self-care, usual activities, pain/discomfort, and anxiety/depression), each with three levels of response or severity (no problems, some problems, or extreme problems). The ability to convert self classification responses into a single index score makes the EQ-5D practical for clinical and economic evaluation [11]. The index-based score is typically interpreted along a scale where 1 represents best possible health and 0 represents dead, with some health states valued as being worse than dead (<0). In addition to the index-based scoring system, the visual analogue scale (VAS) component of the EQ-5D enables the patient to place their current health state on a range from 0 (worst imaginable health state) to 100 (best imaginable health state). Algorithms have been developed based on societal preferences for health states, with the most popular being based on the UK-based population [12], although many other country-specific algorithms are also available [13-18].

The principle aims of this paper were: to synthesise the evidence on the validity and reliability of the EQ-5D in studies within the cardiovascular field; to summarise the EQ-5D based scores reported in studies within the CVD field; and to attempt to stratify mean utility scores according to level of disease severity.

**Methods**

**Data Collection and Assessment**

A computerised search of the current published literature was performed using MEDLINE and EMBASE for the period January 1988 to October 2008. The search strategy combined exploded or medical subject headings relating to the CVD field and the EQ-5D as follows: (‘cardiovascular’/exp OR ‘cardiovascular’) OR (‘cardiac’/exp OR ‘cardiac’) OR (cardiology/exp OR ‘cardiology’) AND ‘euroqol’ OR ‘EQ 5D’ OR ‘EQ5D’. The EuroQol website http://www.euroqol.org was also used to identify unique references, including working papers and conference proceedings that may not have been captured in the initial literature search. Only full-text published papers were included for analysis.

The inclusion criteria required that the paper was original research, and that it reported EQ-5D scores specific to cardiovascular disease or reported psychometric properties of the EQ-5D in a population with cardiovascular disease. Studies that only reported EQ-5D index or VAS scores graphically in terms of change over time were excluded from the analysis. When multiple studies used the same dataset, EQ-5D scores were only reported from one article to avoid double counting. No language restrictions were imposed. Study abstracts that potentially met the inclusion criteria were identified, and full-text articles were retrieved for further review. A standard data abstraction form was developed to facilitate the structured review, which included study design, patient characteristics, intervention information, published source of index-based preference weights and EQ-5D scores as well as details of any other clinical measures; disease-specific quality of life and generic HRQoL instruments. A summary of the results of the literature search is provided in figure 1.

**Data Analysis**

Initially, studies that reported EQ-5D index-based scores and/or VAS scores were sorted into cardiovascular subgroups (e.g. Angina/Myocardial Infarction/CHD etc) that were informed by the latest WHO International Classification of Diseases (ICD-10: 100-199 - diseases of the circulatory system) (Table 1 in Additional file 1) [19]. Confidence intervals were calculated from the sample size and standard deviation (SD) or the standard error when not reported directly in the paper. Scores that were not reported using the appropriate range of scale were transformed, i.e. index-based scores anchored by 0 (dead) and 1 (full health), VAS scores range from 0 (dead) to 100 (full health). If EQ-5D results were stratified (e.g. by age, sex and disease severity), results were only reported once, using the most clinically relevant stratification: CCS angina classification; NYHA heart failure classification or demographic characteristics (% of males/females and mean age of patient cohort). Error bars in Figures 2 and 3 represent 95% confidence intervals around the mean score, which were calculated from the reported SD and sample size. There was no attempt to combine
estimates from different studies in a formal meta-analysis since the main objective was to contrast studies with different features and to explain heterogeneity in the results. The degree of heterogeneity between studies was quantified using the $I^2$ statistic [20]. The $I^2$ statistic uses the sum of the squared differences of each study from the pooled estimate and the degrees of freedom of the test to provide a measure of the percent of total variation across studies due to heterogeneity between studies. A meta-analysis yielding a value of $I^2$ above 75% suggests a high level of heterogeneity between the studies. Psychometric properties were summarised according to the type of property assessed (validity/reliability/responsiveness), the comparison performed, and the statistical test result.

**Results**

The electronic search of databases returned 147 papers of which 66 met the selection criteria. 60 publications reported an EQ-5D index score, VAS score and/or responses to the self-classification system, whilst 10 papers presented evidence of the psychometric properties of the EQ-5D (Figure 1).

Overall, there was wide variation in terms of CVD subgroups, disease stage, age distribution and other methodological aspects (Table 1 in Additional file 1). Of studies reporting mode of administration (n = 41), 42% were filled out on-site by respondents, 52% were mailed-out questionnaires, and 6% were administered via telephone interview. Overall, there was an equal mix of randomised controlled trial (RCT) and observational study designs. Prospective observational study designs were more common than retrospective (69% vs. 31%) and there was an equal mix of longitudinal and cross-sectional studies. The majority of studies (52%) reported EQ-5D index scores using the UK-based algorithm although scores based on Czech, Danish, Dutch, German, US and European preferences were also used [13-18]. However, a number of studies (33%) did not explicitly state the algorithm used to calculate the index score.

Studies of cardiovascular patients that reported psychometric properties of the EQ-5D (n = 10) explored construct validity (convergent and discriminative), typically in terms of correlations with other disease-specific HRQoL measures as well as reliability and
Figure 2 EQ-5D Index Mean scores for Heart Failure patients - Stratified by baseline disease severity (NYHA class).

Figure 3 EQ-5D Index Mean scores for IHD patients - Stratified by baseline disease severity (CCS class).
responsiveness (Table 2 in Additional file 1). Evidence of validity and reliability were reported in studies of ischaemic heart disease (n = 3); cerebrovascular disease (n = 3); heart failure (n = 2) and peripheral vascular disease (n = 2). Convergent validity was the most common property assessed, using Spearman rank correlations to explore associations with another measure. Reliability and responsiveness were generally measured by test-retest statistics; intra-class correlation coefficients (ICC) and effect size (ES). In terms of construct validity, comparisons were made between the EQ-5D and disease specific questionnaires such as the Barthel Index (BI), Kansas City Cardiomyopathy Questionnaire (KCCQ), MacNew Heart Disease Quality of Life Questionnaire, NYHA and VascuQol as well as other generic measures such as the Health Utilities Index (HUI2; HUI3) and the RAND Short Form Health Survey (SF-36) and its derivatives (SF-6D; SF-12).

For convergent validity, moderate to strong agreement represented as significant correlation was generally found between EQ-5D Index and VAS scores and other generic HRQoL measures both at the domain and index level [21-23]. For discriminative validity, the EQ-5D was less able to detect clinical changes than other disease specific measures such as the KCCQ or NYHA and performed better when detecting large rather than small changes in disease severity [24]. There was also evidence of strong ceiling effects (i.e. inability to discriminate between comparatively good health states) across both domain and index values [21,25]. In general, the EQ-5D Index and VAS showed good reliability and responsiveness in comparison to other generic measures such as the SF-12 but were less responsive than disease-specific measures such as the KCCQ [26,27].

A wide range of mean and median EQ-5D scores were reported (Table 3 in Additional file 1). Studies of patients with ischaemic heart diseases (ICD codes I20-I25) reported mean index scores that ranged from 0.45 (SD 0.22) to 0.88 (no SD reported). Visual analogue scale (VAS) scores ranged from a mean of 45 (SD 17) to 82 (SD 13). Studies of heart failure (150) patients reported mean index scores ranging from 0.31 (no SD) to 0.78 (0.11) and mean VAS scores from 37 (21) to 73 (18). Studies of cerebrovascular diseases (160-169) reported mean index scores ranging from 0.24 (0.39) to 0.90 (0.16) and mean VAS scores from 51 (SD 20) to 89 (no SD). Studies of peripheral vascular diseases (I73) reported mean index scores ranging from 0.33 (no SD) to 0.78 (0.23) and mean VAS scores ranging from 49 (no SD) to 71 (8).

The lowest mean EQ-5D index scores were reported in female patients with intermittent claudication undergoing secondary amputation [28]; patients with a large deterioration in heart failure [24]; and post-stroke patients [29] (Table 3 in Additional file 1). The highest mean EQ-5D index scores were reported in elderly CHD patients one year after undergoing exercise training [30]; post-trans-ischaemic attack (TIA) patients at four-year follow-up [31] and patients with history of subarachnoid haemorrhage [32].

An attempt was made to stratify mean EQ-5D index or VAS scores by disease severity (for example by CCS angina grading scale or NYHA heart failure classification). Both the CCS and NYHA scales range from class I (mild symptoms) to class IV (severe symptoms) and CCS can also be graded as 0 for no symptoms (Table 4 in Additional file 1). A previously published study stratified mean EQ-5D scores across CCS grades for patients with stable angina [33]. The results showed mean EQ-5D scores decreasing as the severity of angina increased. EQ-5D index scores ranged from 0.36 (95% CI: 0.25 to 0.48) for CCS grade IV to 0.81 (95% CI: 0.77 to 0.85) for CCS grade 0. Here, there was sufficient data available to stratify mean EQ-5D index scores by NYHA class in heart failure patients and by CCS class in patients with ischaemic heart disease (IHD). EQ-5D index scores were stratified into three categories of NYHA or CCS class based on the percentage of patients in a given group in a study in class III/IV (0-33%; 34-67%; 68-100%). It was assumed here that 0-33% in class III/IV corresponds to mild HF/angina whilst 68-100% corresponds to moderate/severe HF/angina.

In almost all cases, mean EQ-5D index scores increased with an increase in the proportion of patients with mild disease (Figure 2). Mean EQ-5D index scores decreased from 0.78 (SD 0.18) for mild states to 0.51 (SD 0.21) for moderate/severe health states. In common with heart failure patients, mean EQ-5D index scores for IHD patients generally decreased with the increasing proportions of patients with moderate/severe angina (Figure 3). Here, scores decreased from a mean of 0.80 (SD 0.05) for mild angina to 0.45 (SD 0.22) for moderate/severe angina.

An initial attempt was made to summarise the burden of CVD for each disease subgroup by calculating pooled means across studies. Both fixed and random effects meta-analyses were carried out for studies that used reported EQ-5D index scores and disease severity in terms of either CCS Angina class or NYHA heart failure class at baseline. Fixed and random effects meta-analyses of heart failure patients stratified by NYHA class and IHD patients stratified by CCS angina class produced I² indices of between 82-96%, suggesting a high level of statistical heterogeneity between studies [34]. Such a degree of heterogeneity between studies ruled out any further estimation of pooled mean utility scores according to disease severity.
14 studies also provided detailed information on the dimension-specific burden of cardiovascular disease, exploring the distribution of scores across the five dimensions of the EQ-5D [21,23,35-46]. In examining the dimension-specific burden of disease among cardiovascular studies, the trend in the distribution of scores was fairly similar across all five dimensions. In general, problems with usual activities tended to be most common, followed by problems with mobility and pain/discomfort (Figures 4, 5, 6, 7 and 8).

Discussion

In recent years use of the EQ-5D to measure patient HRQoL in published studies within the cardiovascular field has increased. This largely reflects the growing requirement, over time, of clinical trials to consider cost-effectiveness alongside the clinical effectiveness of new interventions. As the “gold standard” form of economic evaluation in many health care systems, cost-utility analyses (CUA) rely on generic measures such as the EQ-5D for the calculation of QALYs. Increased use of the EQ-5D may also support the view that patient reported outcomes and quality of life are becoming more widely accepted as routine measures in clinical studies, with the EQ-5D being an internationally recognised generic measure of HRQoL. This summary of EQ-5D index and VAS scores in the cardiovascular field complements other published reports describing the use of the EQ-5D in the cancer and asthma/COPD literature and of utility scores associated with various conditions [47-50].

The review found that the majority of studies that included the EQ-5D were within IHD (I20 - I25) and cerebrovascular disease (I60 - I69), subgroups, reflecting the relative prevalence of these diseases worldwide. Stratification by disease severity (measured by CCS angina or NYHA heart failure scales) was possible for IHD patients and heart failure patients and illustrated a positive relationship with the EQ-5D when moving from severe to mild disease severity (Figures 2 and 3). However, calculation of pooled means across studies using meta-analytic techniques was not appropriate, given the high level of heterogeneity in terms of study design and patient characteristics. In general, evaluations of the validity and reliability of the EQ-5D suggested fairly strong convergent validity when assessed by correlations with other HRQoL measures and good discriminative abilities in detecting patients whose health status changed by a given clinical magnitude. However, there was evidence of strong ceiling effects across each domain for the index values. In terms of the dimension-specific burden of cardiovascular disease, problems with pain or discomfort were the most common, followed by problems with usual activities and mobility.

There was much heterogeneity in the scores observed across the studies, which was not necessarily entirely
explained by the range of cardiovascular subgroups. The diverse range of index and VAS scores was also related to stage of illness or treatment (for example baseline versus post-treatment measurements) as well as non-disease-related factors such as other co-morbidities and demographic characteristics. Furthermore, no *a priori* quality criteria were imposed on studies included for review in terms of sample size or methodological quality which may explain some of the heterogeneity. On the other hand, imposing stringent inclusion criteria in terms of study methodological quality would have reduced the potential availability of studies considered for analysis. It is difficult to predict to what extent the level of heterogeneity would have been reduced if more stringent inclusion criteria had been imposed for the literature review.
Overall, this study illustrated the difficulty in attempting to adequately deal with statistical heterogeneity based on aggregated data from published studies [51]. This would suggest that individual patient-level data is required in order to estimate mean utility scores according to disease stage, at least within the cardiovascular field. Furthermore, not all studies used the same algorithm to calculate index-based scores with a third of studies also failing to report which scoring system was used. The choice of algorithm used to convert self-classification scores can affect the index-based score, as shown in a recent study which compared UK and US scoring algorithms in patients undergoing percutaneous coronary intervention (PCI) [52]. However, whilst country-specific societal preferences may reduce the scope for comparing HRQoL estimates across studies from...
different countries, they are more helpful to local decision making, especially when allocating resources within national health care programmes.

Conclusion

HRQoL measures such as the EQ-5D can be useful tools to clinicians in terms of evaluating the impact of cardiovascular disease on patients and can help to inform decision making and resource allocation. The use of the EQ-5D in CVD studies has increased in recent years and published studies provide evidence of its validity and reliability. The variation in EQ-5D index and VAS scores reported here largely reflect systematic differences in terms of disease stage, treatment and patient characteristics. In the future, as more studies of CVD present EQ-5D scores according to disease severity, it may be possible to calculate pooled mean estimates that can be useful in modelling of CVD-related health outcomes in economic evaluations.

Abbreviations used in Tables/Figures

AAA: Abdominal aortic aneurysm; ACS: Acute coronary syndromes; ACT: Anticoagulation therapy; AF: Atrial fibrillation; AH-Drug: Anti-hypertensive drug therapy; AHF: Advanced heart failure; AMI: Acute myocardial infarction; Amp.: Amputation; Angio: Coronary angiography; AAA: Abdominal aortic aneurysm; ACT: Anticoagulation therapy; ASCOT-AHD: Anglo-Scandinavian cardiac outcomes trial - anti-hypertensive drug treatment; Asym./Sym.: Asymptomatic/Symptomatic; AUT: Austria; AVR: Aortic valve replacement; BI: Barthel Index; BOTH-CABG/PCI/MM: Patients who are suitable for both CABG and PCI and receive CABG/PCI/MM; CABG: Coronary artery bypass graft; CABG-80: Coronary artery bypass surgery in octogenarians; CABG-CABG/PCI/MM: Patients who are suitable for CABG and receive CABG/PCI/MM; CABG-CPB: CABG using heart lung machine; CAD: Coronary artery disease; C-Arrest: Cardiac arrest; CCR: Comprehensive cardiac rehabilitation; CCS: Canadian Cardiovascular Society; CCU: Coronary care unit; CES-D: Centre for Epidemiological Studies - Depression Scale; CHD: Coronary heart disease; CHD-PHARM/Control: Community pharmacy-led medicines management programme/control treatment for patients with CHD; CML: Case method learning supported lipid-lowering strategy; COMM - CVD/NOCVD: Community dwelling-based elderly patients with/without CVD; CR: Cardiac resynchronisation; CR-Home/Hosp: Home/Hospital-based cardiac rehabilitation; C-REHAB: Cardiac rehabilitation; CS: Conservative strategy; CVA: Cerebrovascular Accident; CVD: Cardiovascular disease; Duplex US: Duplex ultrasonography; Echo: Echocardiography; EHS-CR: Euro Heart Survey on coronary revascularisation; Endo: Endovascular AAA surgery; ES: Effect size; Exercise-Qol: Long-term effects of exercise training on quality of life; F-u: Follow-up; GRS: Guyatt’s responsiveness statistic; HeartMed: Lifestyle advice intervention by community pharmacists for heart failure patients; HF: Heart failure; HOSP - CVD: Hospital-based elderly patients with CVD; HRQoL: Health-related quality of life; HU12/3: Health Utilities Index mark 2/3; IC: Intermittent claudication; ICC: Intra-class correlation; ICD: Implantable cardioverter defibrillator; ICP: Integrated care pathway; IHD: Ischaemic heart disease; IQR: Interquartile range; IS: Interventional strategy; IV: Intravenous; KCCQ: Kansas City Cardiomyopathy Questionnaire; LV: Left ventricular; MacNew: MacNew Heart Disease Quality of Life Questionnaire; MCS: Mental component summary; MDT: Multi-disciplinary team; MEDMAN: Community pharmacy-led medicines management services; MEPS: Medical expenditure panel survey; MI: Myocardial infarction; MI - Self-help: Home-based self-help rehabilitation package for MI patients; MIDCAB: Minimally invasive direct CABG; MM: Medical management; MR Angio: Magnetic resonance angiography; MRI: Magnetic resonance imaging; MT: Medical therapy; MVPs: Mitral valve prolapse syndrome; NYHA: New York Heart Association; OP-CABG: Off-pump CABG; Open: Open AAA surgery; PAOD: Peripheral arterial occlusive disease; PCI: Percutaneous coronary intervention; PCI-BMS: PCI with bare-metal stents; PCI-CABG/MM/PCI: Patients who are suitable for PCI and receive CABG/PCI/MM; PCI-DES: PCI with drug-eluting stents; PCS: Physical component summary; PER: Peripheral endovascular revascularisation; Pre-op: Pre-operation; Proxy: HRQol questionnaire completed by spouse/family member; PSM: Patient self-management; P-PTCA: Primary PTCA; P-Stent: Primary stent placement; PTCA: Percutaneous transluminal coronary angioplasty; QLMI: Quality of Life after MI questionnaire; QoL: Quality of life; RCT: Randomised controlled trial; REV/NO REV: Eligible/Ineligible for Revascularisation; SAH: Subarachnoid haemorrhage; SCOPE-Drug/Control: Study on cognition and prognosis in the elderly - Drug/Control treatment; SD: Standard deviation; SES: Socioeconomic status; SF-36: Short-form 36-item health survey questionnaire; SF-6D: Short-form 6D; SF-12: Short-form 12-item health survey questionnaire; SPECT: Single photon emission computed tomography; SRM: Standardised response means; Stroke-4Y: Four years post-stroke; TIA: Transischaemic attack; Trans.: Heart Transplantation; Tx: Treatment; UC: Usual care; VAD: Ventricular assist device; VAS: Visual analogue scale; VascuQol: Vascular Quality of Life Questionnaire; -ve/+ve: Deterioration/Improvement in condition; WHO-ICD: World Health Organisation - International Classification of Diseases; W-list: Waiting-list.
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Author details
1Health Economics Research Group, Brunel University, Uxbridge, UK
2Papworth Hospital NHS Trust, Cambridge, UK
3MRC Biostatistics Unit, Institute of Public Health, Cambridge, UK
4National Collaborating Centre for Mental Health, The Royal College of Psychiatrists, London, UK.

Authors’ contributions
MD participated in the design of the study, carried out the systematic literature review, conducted any data analysis and drafted the manuscript. KG provided support in the statistical analysis and helped to draft the manuscript. LS participated in the design of the study, provided support in the statistical analysis and helped to draft the manuscript. MB conceived of the study, participated in the design of the study and helped to draft the manuscript. All authors read and approved the final manuscript.

Competing interests
The authors declare that they have no competing interests.

References
1. World Health Organisation: The World Health Report 2003: Shaping the Future. Geneva 2003.
2. Leal J, Luengo-Fernandez R, Gray A, Petersen S, Rayner M: Economic burden of cardiovascular diseases in the enlarged European Union. Eur Heart J 2006, 27:1610-1619.
3. Swenson JR, Clinch JJ: Assessment of quality of life in patients with cardiac disease - the role of psychosomatic medicine. J Psychosom Res 2000, 48:405-415.
4. The Criteria Committee of the New York Heart Association: Nomenclature and Criteria for Diagnosis of Diseases of the Heart and Great Vessels. Boston, MA: Little, Brown & Co. 9 1994.
5. Campeau L: The Canadian Cardiovascular Society grading of angina pectoris revisited 30 years later. Can J Cardiol 2002, 18:439-442.
6. Spertus JA, Winder JA, Dewhurst TA, Deyo RA, Prodzinski J, McDonell M, Fihn SD: Development and evaluation of the Seattle Angina Questionnaire: a new functional status measure for coronary artery disease. J Am Coll Cardiol 1995, 25:533-341.
7. Hofer S, Lim L, Guyatt G, Oldridge N: The Macnew Heart Disease health-related quality of life instrument: A summary. Health Qual Life Outcomes 2004, 2:3.
8. Guyatt G: Measurement of health related quality of life in heart failure. J Am Coll Cardiol 1993, 22(Suppl A):18A-19A.
9. Brazier J, Roberts J, Deverill M: The estimation of a preference-based measure of health from the SF-36. J Health Econ 2002, 21:271-292.
10. Furlong WJ, Feeny DH, Torrance GW, Barr RD: The Health Utilities Index (HUI) system for assessing health-related quality of life in clinical studies. Ann Med 2001, 33:375-384.
11. Raber R, de Charro F: EQ-SD: a measure of health status from the EuroQol Group, Ann Med 2001, 33:337-343.
12. Dolan PD: Modelling Valuations for EuroQol Health States. Med Care 1997, 35(11):1095-1108.
13. Dankova I, Dilouhy M: The measurement of health status with Czech version of European quality of life questionnaire: version EQ-SD. Czech Health Policy Econ 2001, 2.
14. Wittrup-Jensen K, Lauridsen J, Cudec C, Brooks R, Pedersen K: Estimating Danish EQ-SD tariffs using the time trade-off (TTO) and visual analogue scale (VAS) methods. Lund: Swedish Institute for Health Economics257-292.
15. Lamers L, Stalmeier P, Krabbe P, Busschbach J: The Dutch tariff: results and arguments for an effective design for national EQ-SD valuation studies. Health Econ 2006, 15:1121-1132.
16. Claas C, Greiner W, Uber A, Schülenberg JM: The new German version of the EuroQol quality of life questionnaire. Centre for Health Policy and Law, Erasmus University Rotterdam13-23.
17. Shaw NP, Johnson JAP, Coons SJP: US Valuation of the EQ-5D Health States: Development and Testing of the D1 Valuation Model. Med Care 2003, 41:203-220.
18. Greener W, Weijnen T, Neuenhuizen M, Oppe S, Badia X, Busschbach J, Buxton M, Dolan P, Krabbe P, Ohrnmaa A, Parkin D, Rose M, Sintonen H, Tsuichiy A, Charo F. A single European currency for EQ-5D health states. Eur J Health Econ 2003, 4:222-231.
19. WHO: International Statistical Classification of Diseases and Related Health Problems. Geneva 2007.
20. Higgins J, Thompson SG: Quantifying heterogeneity in meta-analysis. Stat Med 2002, 21:1593-1600.
21. Schweikert B, Hahmann H, Leidl R: Validation of the EuroQol questionnaire in cardiac rehabilitation. Heart 2006, 92:62-76.
22. Novels D, McGlinn J, Westfall JM, Holcomb S: Validation of the EQ-SD quality of life instrument in patients after myocardial infarction. Qual Life Res 2005, 14:95-105.
23. Pickard AS, Johnson JA, Feeny DH: Responsiveness of generic health-related quality of life measures in stroke. Qual Life Res 2005, 14:207-219.
24. Sperrus J, Peterson E, Conard M, Heidenreich P, Rumohlz M, Jones P, McCullogh PA, Pina I, Wooley J, Weintrab WS, Rumsfeld JS: Monitoring clinical changes in patients with heart failure: A comparison of methods. Am Heart J 2005, 150:707-715.
25. van Stel H, Buxens E: Comparison of the SF-6D and the EQ-5D in patients with coronary heart disease. Health Qual Life Outcomes 2006, 4:20.
26. Eurch D, Johnson J, Reid K, Spertus J: Assessing responsiveness of generic and specific health related quality of life measures in heart failure. Health Qual Life Outcomes 2006, 4:89.
27. Dorman P, Slattery J, Farell B, Dennis M, Sandercop P: Qualitative Comparison of the Reliability of Health Status Assessments With the EuroQol and SF-36 Questionnaires After Stroke. Stroke 1998, 29:63-68.
28. Tangelder MJ, McDonnell J, Van Busschbach JJ, Buxens E, Algra A, Lawson JA, Eikelboom BC: Quality of life after infrainguinal bypass grafting surgery. Journal of Vascular Surgery 1999, 29:913-919.
29. Pickard AS, Johnson JA, Feeny DH, Shuaib A, Cantiere KC, Nasser AM: Agreement Between Patient and Proxy Assessments of Health-Related Quality of Life After Stroke Using the EQ-5D and Health Utilities Index. Stroke 2004, 35:607-612.
30. Hage C, Mattsson E, Stahlle A: Long term effects of exercise training on physical activity level and quality of life in elderly coronary patients - A three- to six-year follow-up. Physiother Res Int 2003, 8:13-22.
31. Haacke C, Athuras A, Sporteke A, Sibbert U, Back T, Doder R: Long-Term Outcome After Stroke: Evaluating Health-Related Quality of Life Using Utility Measurements. Stroke 2006, 37:193-198.
32. Schaaf van der IC, Werner MHT, Velthuis BK, Buxens E, Bossyurt PM, Rinkel GJE: Psychosocial impact of finding small aneurysms that are left untreated in patients previously operated on for ruptured aneurysms. J Neurosurg Psychiatry 2006, 77:478-752.
33. Longworth L, Buxton M, Sclplpher M, Smith D: Estimating utility data from clinical indicators for patients with stable angina. Eur J Health Econ 2005, 6:347-353.
Influence of chronic cardiovascular disease and hospitalisation due to this disease on quality of life of community-dwelling elderly. Qual Life Res 2006, 15:1281-1289.

Lazen A, Schole-reimer W, Norkew DA, De Geest S, Fidruid R, Heinikki J, Jaarsma T, Mortensson J, Moore P, Smith K, Stewart S, Stromberg A, Thompson DR, Wiens W. Pharmacological treatment and perceived health status during 1-year follow up in patients diagnosed with coronary artery disease, but ineligible for revascularisation: Results from the Euro Heart Survey on Coronary Revascularisation. Eur J Cardiovasc Nurs 2006, 5:115-121.

Sollano JA, Rose EA, Williams DL, Thornton BM, Quint ER, Apfelbaum MM, Vasma GM, Cannavale GA, Smith CR, Reemstma KM, Greene R. Cost-Effectiveness of Coronary Artery Bypass Surgery in Octogenarians. Annals of Surgery 1999, 228(3):297-306.

Almenar-Pertejo M, Almenar L, Martinez-Dolz L, Campos J, Galan J, Grinos P, Ortega F, Ortega T, Rebollo P, Salvador A. Study on Health-Related Quality of Life in Patients With Advanced Heart Failure Before and After Transplantation. Transplant Proc 2006, 38:2524-2526.

Aust J, Williams WR, Ross L, Hutchison S. Five-year follow-up findings from a randomized controlled trial of cardiac rehabilitation for heart failure patients: Euro Heart Survey. Eur Heart J Cardiovass Prev Rehabil 2003, 15:162-166.

Calvert MJ, Freemantle N, Cleland JF. The impact of chronic heart failure on health-related quality of life data acquired in the baseline phase of the CARE-HF study. Eur J Heart Fail 2005, 7:243-251.

Hofer S, Kulhij W, Gruninger U, Brandl D, Gasser A, Klicpera M, Laimer H, Marko C, Schwann H, Moller R. Cardiac rehabilitation in Austria: short term quality of life improvements in patients with heart disease. Wien Klin Wochenschr 2006, 118:744-753.

Lottman PEM, Laheij RJF, Cuypers PVM, Bender M, Buth J. Health-related quality of life outcomes following elective open or endovascular AAA Repair: A randomised controlled trial. J Endovasc Surg 2004, 11:323-329.

Sulch D, Melbouron A, Perez I, Kalia L. Integrated Care Pathways and Quality of Life on a Stroke Rehabilitation Unit. Stroke 2002, 33:1600-1604.

Bush J, Hunink M. Comparison of the Health Utilities Index Mark 3 (HUI3) and the EuroQol EQ-5D in patients treated for intermittent dialysis. Qual Life Res 2000, 9:591-601.

Jowett S, Bryan S, Murray E, McCafran D, Rathery J, Hobbs F, Fitzmaurice D. Patient self-management of anticoagulation therapy: a trial-based cost-effectiveness analysis. Br J Haematol 2006, 134:632-639.

Granja C, Cabral G, Pinto A, Costa-Pereira A. Quality of life 6 months after cardiac arrest. Resuscitation 2002, 55:37-44.

Pickard AS, Wilkie CT, Lin HW, Lloyd A. Health Utilities Using the EQ-5D in Studies of Cancer. Pharmacoconomics 2007, 25:365-384.

Pickard AS, Wilkie C, Jung E, Patel S, Reimder IR, Salvador P. A use of a preference-based measure of health (EQ-5D) in COPD and asthma. Respir Med 2008, 102:519-536.

Sullivan PW, Lawrence WF, Gushchyan VM. A National Catalog of Preference-Based Scores for Chronic Conditions in the United States. Med Care 2005, 43:736-749.

Tengs TO, Wallace AM. One Thousand Health-Related Quality-of-Life Estimates. Med Care 2000, 38:683-687.

Higgins J, Thompson S, Deeks JJ, Altman DG. Statistical heterogeneity in systematic reviews of clinical trials: a critical appraisal of guidelines and practice. J Health Serv Res Policy 2002, 7:51-61.

Shrive FM, Ghali WA, Johnson JA, Donaldson CM, Manns BJ. Use of the U.S. and U.K. Scoring Algorithm for the EuroQol-5D in an Economic Evaluation of Cardiac Care. Med Care 2007, 45:269-276.

Ascierno R, Reeves BC, Taylor FC, Seehra HK, Angelini GD. Beating heart against cardiopulmonary arrest studies (BHACAS 1 and 2): quality of life at mid-term follow-up in two randomised controlled trials. Eur Heart J 2004, 25:765-770.

Rocca HBP-L, Kaiser C, Bernheim A, Zellweger MU, Jeger R, Buser PT, Osswald S, Pfisterer M. Cost-effectiveness of drug-eluting stents in patients with high or low risk of major cardiac events in the Basel Stent KostenEffektivitats Trial (BASKET): an 18-month analysis. Lancet 2007, 370:1552-1559.

Denvir MA, Lee AJ, Rysdale J, Walker A, Eteiba H, Starkey IR, Pelt JP. Influence of socioeconomic status on clinical outcomes and quality of life after percutaneous coronary intervention. J Epidemiol Community Health 2006, 60:1085-1088.

Dunning J, Waller JRL, Smith B, Pitts S, Kendall SWH, Khan K. Coronary Artery Bypass Grafting is Associated With Excellent Long-Term Survival and Quality of Life: A Prospective Cohort Study. Ann Thorac Surg 2008, 85:1988-1993.

Eefting F, Nathoe H, van Dijk D, Jansen E, Lahpor J, Stella P, Suyker W, Diephuis J, Suryaparanda H, Ernst S, Borst C, Buskens E, Grobbee D, de Jaegere P. Randomized Comparison Between Stenting and Off-Pump Bypass Surgery in Patients Referred for Angioplasty. Circulation 2003, 108:2870-2876.

Griffin SC, Barber JA, Manca A, Sculptor MJ, Thompson SG, Buxton MJ, Hemingway H. Cost effectiveness of clinically appropriate decisions on alternative treatments for angina pectoris: prospective observational study. BMJ 2007, 334:626-628.

Keseling A, Zethraeus N, Henriksson P. Cost of lipid lowering in patients with coronary artery disease by Case Method Learning. Int J Technol Assess Health Care 2005, 21:180-190.

Kim J, Henderson RA, Pocock SJ, Clayton T, Sculptor MJ, Fox KAA. Health-related quality of life after interventional or conservative strategy in patients with unstable angina or non-ST-segment elevation myocardial infarction: One-year results of the third randomized intervention trial of unstable angina (RITA-3). J Am Coll Cardiol 2005; 45:221-228.

Krase M, Hoehstrasser S, Zwicker AD, Keljberg J. Comprehensive cardiac rehabilitation: A cost assessment based on a randomized clinical trial. Int J Technol Assess Health Care 2006, 22:478-483.

Lacey EA, Musgrave RJ, Freeman JV, Tod AM, Scott P. Psychological morbidity after myocardial infarction in an area of deprivation in the UK: evaluation of a self-help package. Eur J Cardiovasc Nurs 2004, 3:129-134.

Legrand WMG, Serruys PW, Unger F, van Hout BA, Vrolijk MCM, Fransen GMP, Nielsen TT, Pausken PK, Gomes RS, de Queiroz de Melo J, Marques dos Santos Neves J, Lindeboom W, Backx F. Three-Year Outcome After Coronary Stenting Versus Bypass Surgery for the Treatment of Multivessel Disease. Circulation 2004, 109:1114-1120.

The Community Pharmacy Medicines Management Project Evaluation Team: The MEDSMAN study: a randomized controlled trial of community pharmacy-led medicines management for patients with coronary heart disease. Fam Pract 2007, 24:189-200.

Pulska JD, Williams WH, Mahoney EM, Huber PR, Block PC, Duke PG, Staples JR, Glas KE, Marshall ME, McCall SA, Petersen RJ, Bailey D, Weintraub WS, Guyton RA: Off-Pump vs Conventional Coronary Artery Bypass Grafting: Early and 1-Year Graft Patency, Cost, and Quality-of-Life Outcomes: A Randomized Trial. JAMA 2004, 291:1841-1849.

Reeves BC, Angelini GD, Bryan AJ, Taylor FC, Crpps T, Spyt TJ, Samani NJ, Roberts JA, Jackin P, Seehra HK, Cullford LF, Keenan DJM, Rowlands DJ, Clarke B, Stanbridge R, Fiaile R. A multi-centre randomised controlled trial of minimally invasive direct coronary bypass grafting versus percutaneous transluminal coronary angioplasty with stenting for proximal stenosis of the left anterior descending coronary artery. NCHTA Health Technol Assess 2004, 8:1-56.

Sharples L, Hughes V, Crean A, Dyer M, Buxton M, Goldsmith K, Stone D. Cost-effectiveness of functional cardiac testing in the diagnosis and management of coronary artery disease: a randomised controlled trial. The CECA trial. 11(49). NCHTA Health Technol Assess 2007, 11:1-136.

Shrive FM, Manns BJ, Galbraith PD, Knudtson ML, Ghali WA, For The APPROACH Investigators: Economic evaluation of sirolimus-eluting stents. CMAJ 2005, 172:945-951.

Taylor RS, Watt A, Dalal HM, Evans PH, Campbell JL, Read KLO, Mourant AJ, Wingham J, Thompson DR, Pereira Gray DJ. Home-based cardiac rehabilitation versus hospital-based rehabilitation: A cost-effectiveness analysis. Int J of Cardiol 2007, 119:196-201.

Xie J, Wu EQ, Zheng ZJ, Sullivan PW, Zhan L, Labarde DR. Patient-Reported Health Status in Coronary Heart Disease in the United States: Age, Sex, Racial, and Ethnic Differences. Circulation 2008, 118:491-497.

Cleland JGF, Daubert JC, Erdmann E, Freemantle N, Gras D, Kappenberger L, Tavazzi L. The Effect of Cardiac Resynchronization on Morbidity and Mortality in Heart Failure. N Engl J Med 2005, 352:1539-1549.

Holland R, Brooksby I, Lenaghan E, Ashton K, Hay L, Smith K, Shepstone L, Lipp A, Daly C, Howe A, Hall R, Harvey J. Effectiveness of visits from
community pharmacists for patients with heart failure: HeartMed randomised controlled trial. BMJ 2000, 334:1098.
73. Sharples L, Buxton M, Caine N, Cafferty F, Demnis N, Dyer M, Freeman C: Evaluation of the ventricular assist device programme in the UK. NCCHTA Health Technol Assess 2006, 10:1-138.
74. Spencer CA, Norman PE, Jamrozik K, Tsuhy R, Lawrence-Brown M: Is screening for abdominal aortic aneurysm bad for your health and well-being? ANZ J Surg 2004, 74(12):1069-1075.
75. Buxton M, Caine N, Chase D, Connelly D, Grace A, Jackson C, Parkes J, Sharplis L. A review of the evidence on the effects and costs of implantable cardioverter defibrillator therapy in different patient groups, and modelling of cost-effectiveness and cost-utility for these groups in a UK context. NCCHTA Health Technol Assess 2006, 10:1-132.
76. Groeneveld F, Matta M, Suh J, Yang F, Shea J: Quality of Life Among Implantable Cardiovore-Defibrillator Recipients in the Primary Prevention Therapeutic Era. Pacing Clin Electrophysiol. 2007, 30:463-471.
77. van Eck JW, van Hemel NM, den Boer A, Taku W, Grobbbee DE, Moons KG: Predictors of improved quality of life 1 year after percutaneous transluminal balloon angioplasty. Am Heart J 2008, 156:491-497.
78. Park J, White AR, James MA, Hemsley AG, Johnson P, Chambers J, Ernst E: Acupuncture for Subacute Stroke Rehabilitation: A Sham-Controlled, Subject- and Assessor-Blind, Randomized Trial. Arch Intern Med 2005, 165:2025-2031.
79. Ryan T, Endersby P, Rigby AS. A randomized controlled trial to evaluate intensity of community-based rehabilitation provision following stroke or hip fracture in old age. Clin Rehabil 2006, 20:123-131.
80. Xie J, Wu EQ, Zheng ZJ, Croft JB, Greenlund KJ, Mensah GA, Labarthe DR: Utility loss and indirect costs following cardiovascular events in hypertensive patients: the ASCOT health economic substudy. Arch Intern Med 2005, 165:2025-2031.
81. Innocenti AD, Elmfeldt D, Hansson L, Breteler M, James O, Lithell H, Olofsson B, Skoog I, Tenkwalder P, Zanchetti A, Wiklund I: Cognitive Function and Health-Related Quality of Life in Elderly Patients with Hypertension - Baseline Data from the Study on Cognition and Prognosis in the Elderly (SCOPE). Blood Press 2002, 11:157-165.
82. Krijnen P, van Jaarsveld BC, Hunink MGM, Habbema JDF: The effect of treatment on health-related quality of life in patients with hypertension and renal artery stenosis. J Hum Hypertens 2005, 19:467-470.
83. Lindgren P, Kahan T, Poullier N, Buxton M, Swaner P, Dahloff B, Jonsson B: Utility loss and indirect costs following cardiovascular events in hypertensive patients: the ASCOT health economic substudy. Eur J Health Econ 2007, 8:25-30.
84. Cook TA, Galland RB: Quality of life changes after angioplasty for claudication: medium-term results affected by comorbid conditions. Cardiovasc Surg 1997, 5:424-426.
85. de Vries M, Ouwendijk R, Flobbe K, Nelemans PJ, Kessels AG, Schurink GH, van Exel NJA, Scholte op Reimer WJM, Koopmanschap MA: Assessment of post-stroke quality of life in cost-effectiveness studies: The usefulness of the Barthel Index and the EuroQol-5D. Qual Life Res 2004, 13:427-433.
86. de Vries M, Ouwendijk R, Kessels AG, de Haan MW, Flobbe K, Hunink MGM, van Engelshoven JMA, Nelemans PJ: Comparison of generic and disease-specific questionnaires for the assessment of quality of life in patients with peripheral arterial disease. J Vasc Surg 2005, 41:261-268.
87. Bussbach Jv, McDonnell JM, Tangelder MJ, Eikelboom BCM, Buskens EM, Algra AM, Lawson JEM: EuroQol values for economic modelling quality of life after infrarenal bypass grafting surgery: a rectification. J Vasc Surg 1999, 30:1162-1163.

after mechanical aortic valve replacement. Cardiovasc Surg 2003, 11:381-387.
92. Spiraki C, Katselidou D, Papakonstantinou V, Perzerakos P, Maniadakis N: Health-related quality of life measurement in patients with coronary heart disease and heart failure to a cardiology department of a secondary urban hospital in Greece. Hellenic J Cardiol 2008, 49:241-247.
93. van Exel NJA, Scholte op Reimer WJM, Koopmanschap MA: Assessment of post-stroke quality of life in cost-effectiveness studies: The usefulness of the Barthel Index and the EuroQol-5D. Qual Life Res 2004, 13:427-433.

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