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Using a Discrete Choice Experiment Involving Cost to Value a Classification System Measuring the Quality-of-Life Impact of Self-Management for Diabetes

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ABSTRACT

Objectives: To describe the use of a novel approach in health valuation of a discrete choice experiment (DCE) including a cost attribute to value a recently developed classification system for measuring the quality-of-life impact (both health and treatment experience) of self-management for diabetes. Methods: A large online survey was conducted using DCE with cost on UK respondents from the general population (n = 1497) and individuals with diabetes (n = 405). The data were modeled using a conditional logit model with robust standard errors. The marginal rate of substitution was used to generate willingness-to-pay (WTP) estimates for every state defined by the classification system. Robustness of results was assessed by including interaction effects for household income. Results: There were some logical inconsistencies and insignificant coefficients for the milder levels of some attributes. There were some differences in the rank ordering of different attributes for the general population and diabetic patients. The WTP to avoid the most severe

state was £1118.53 per month for the general population and £2356.02 per month for the diabetic patient population. The results were largely robust. **Conclusions:** Health and self-management can be valued in a single classification system using DCE with cost. The marginal rate of substitution for key attributes can be used to inform cost-benefit analysis of self-management interventions in diabetes using results from clinical studies in which this new classification system has been applied. The method shows promise, but found large WTP estimates exceeding the cost levels used in the survey.

Keywords: cost, diabetes, discrete choice experiment, preference-based measures.

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Introduction

Discrete choice experiment (DCE) methods are increasingly being applied as a means to value the benefits of health care interventions. DCEs have usually been used to value the process of health care (either in isolation of or in combination with health outcomes) using bespoke or study-specific attributes developed for individual studies [1]. Recent DCE applications in diabetes include, for example, an investigation of patient preferences for insulin therapy and clinical outcomes in type 2 diabetes [2]. Recent work has extended the use of DCE to valuing classification systems for measuring health-related quality of life (HRQOL) such as the five-level EuroQol five-dimensional questionnaire (EQ-5D-5L) [3] and the six-dimensional health state short form (SF-6D) [4,5] on the 0 to 1 quality-adjusted life-year (QALY) scale

by adding an additional attribute for duration. This approach has been referred to as the DCE_{TTO} (TTO, time trade-off) approach in the literature [6–11]. These health state utility values can then be used to estimate QALYs for use in cost-utility analysis and for submission to regulatory agencies such as the National Institute of Health and Care Excellence in the United Kingdom [12] or the Pharmaceutical Benefits Advisory Committee in Australia [13].

An alternative to cost-utility analysis is cost-benefit analysis (CBA), in which the benefits of interventions are represented by monetary values. This approach has been used to capture the benefits of interventions beyond the health outcomes achieved including benefits gained from the process of care delivery. Monetary values of the benefits of interventions are often measured directly by asking respondents how much they would be willing to pay for one intervention over another (e.g., [14]). DCE

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methods may also be applied to obtain willingness-to-pay (WTP) estimates indirectly [15–17] by including cost as an additional attribute in the DCE. As mentioned previously, this approach has been applied recently in diabetes (e.g., in the study by Feher et al. [2]). Nevertheless, to our knowledge, to date this methodology has not been used to value a pre-existing classification system. Application of a DCE approach in this context provides an analogous way of valuing health to DCE $_{\rm TTO}$ used with the EQ-5D and the SF-6D, but provides monetary estimates of the WTP to avoid a health state that can potentially be used to inform CBA.

Diabetes costs across the globe are rising because of the increased prevalence of the disease and the increased complexity of its treatment. For example, in the United Kingdom, £936.7 million was spent on prescriptions for diabetes in 2015 [18], and the total cost of diabetes in the United Kingdom is estimated to be £23.7 billion [19]. Structured education in diabetes is one of nine key care process checks recommended by the National Institute of Health and Care Excellence [20]. It benefits patients by giving them the confidence and skills to self-manage their condition, but of those newly diagnosed, less than 6% have been recorded as attending such a course. Evaluating the true monetary value of interventions designed to improve self-management is urgently needed. Self-management of diabetes varies from one individual to another, and similarly the impact of self-management upon an individual is also very personal.

Currently, quality of life is linked to QALYs on the basis of questionnaires that are not diabetes-specific (e.g., the EQ-5D or the SF-12), and so the intended improvement in self-management skills of an intervention cannot be evaluated in economic terms. Likewise, measures that are diabetes-specific (e.g., Problem Areas in Diabetes [21]) are not linked to QALYs. The classification system used in this study was developed to provide a formal and consistent way to take account of self-management across different interventions, because existing measures do not consider the direct impact of different self-management regimes on patients' quality of life from their own perspective [22]. Without the use of a single widely applicable classification system, the change in processes is often measured using studyspecific descriptions or vignettes, rather than assessing the impact on quality of life through the use of patient-completed questionnaires in clinical studies.

This article describes the use of DCE including a cost attribute to value a classification system measuring the quality-of-life impact of self-management for diabetes. The article presents a DCE survey with a cost attribute conducted in general population and diabetic patient samples as well as the results of regression analyses to model the DCE data to provide monetary values of the WTP to avoid each state defined by the classification system for both general population and diabetic patient samples. We then discuss the results in terms of the implications for valuing this and other classification systems using this method.

Methods

Classification System

The Health and Self-Management in Diabetes classification system was developed to capture the impact of self-management on quality of life in diabetes (see Fig. 1). Four of the dimensions (mood, hypoglycemic attacks, vitality, and social limitations) represent HRQOL and the remaining four dimensions (control, hassle, stress, and support) represent self-management. The dimensions of HRQOL are taken from the Diabetes Health Profile-Five Dimension [23], a diabetes preference-based measure developed from the Diabetes Health Profile [24,25] and the short form 36 health survey (the vitality item) [26]. The development of

the classification system is reported in detail elsewhere [22] and research is ongoing to determine the psychometric properties of the measure and its performance relative to the EQ-5D-5L.

Valuation Technique

DCE tasks present two or more profiles, in which each profile consists of attribute levels selected from a classification system and respondents are asked to indicate their preferred profile. DCE was selected in this study because it enables WTP values to be generated for every state defined by the classification system through the inclusion of a cost attribute, and the technique is amenable to online data collection [1].

Selecting the Levels of the Cost Attribute

Limited guidance is provided in the DCE literature about how to choose levels for a cost attribute, and many published studies are either extremely brief in their details of how they determined the levels for the cost attribute or do not report details at all. Nevertheless, the levels should accurately capture the range of preferences for most of the respondents; otherwise, their inclusion will not add any useful information. It is important to ensure the levels are not too high or too low for the treatment or condition being valued, because otherwise cost would be either prohibitive or irrelevant [15]. It has been argued that the range for the cost attribute levels should incorporate values that are higher than the market price, because this may not be the maximum amount that people are willing to pay [16]. Typically, cost levels used in the literature reflect a range around mean cost that includes either a low cost or zero cost. In terms of wording the cost attribute, previous DCE experiments with a cost attribute in diabetes have used "personal cost to you each month" [27], "payment per month out of pocket," [2] and "cost of diabetes medicines each month" [28].

To empirically inform the selection of the levels of the cost attribute, an online binary choice survey of 400 members of the general population was conducted to assess people's WTP for hypothetical self-management and HRQOL states (recruitment followed the same process for the main general population survey reported later). Respondents completed experimental binary choice questions, where each question was a choice between a poor state with zero cost or a good state with nonzero cost that was randomly varied across different questions using various levels (across different survey versions the levels used were £10, £25, £50, £75, £100, £150, £200, £300, £400, and £600). The proportions of respondents choosing the better state with nonzero cost were compared to determine how the different costs impacted on choice.

Four levels of the cost attribute were selected, in common with the four severity levels of all other attributes. In the literature of DCE involving a cost attribute to determine WTP, the severity levels are not usually equal but increase exponentially (e.g., 2, 4, 8, 16), and this approach was used here to inform the selection of levels. The lowest level of £10 was selected because about a quarter of the respondents being asked the question were not willing to pay £10 to improve their health. The highest level of £600 was selected as the upper end because about one-third of the respondents were willing to pay £600 to improve their health. Levels of £75 and £200 were selected to represent the intermediate cost levels to ensure good coverage.

Selecting Profiles

Eight attributes from the classification system plus four cost levels resulted in 262,144 profiles, and many millions of possible pairs. Therefore, a subset of profiles was selected using D-optimal methods in Ngene software, distributed by ChoiceMetrics [29] to

Dimension	Level	Wording		
Mood	1	You <u>never</u> find yourself losing your temper over small things		
	2	You sometimes find yourself losing your temper over small		
		things		
	3	You <u>usually</u> find yourself losing your temper over small		
		things		
	4	You <u>always</u> find yourself losing your temper over small		
		things		
Hypoglycemic attacks	1	You never worry about going hypo		
	2	You sometimes worry about going hypo		
	3	You <u>usually</u> worry about going hypo		
	4	You <u>always</u> worry about going hypo		
Vitality	1	You are never tired		
	2	You are sometimes tired		
	3	You are usually tired		
	4	You are <u>always</u> tired		
Social Limitations	1	Your days are <u>never</u> tied to meal times		
	2	Your days are sometimes tied to meal times		
	3	Your days are <u>usually</u> tied to meal times		
	4	Your days are <u>always</u> tied to meal times		
Control	1	You feel you have a lot of control of your diabetes		
	2	You feel you have some control of your diabetes		
	3	You feel you have <u>little control</u> of your diabetes		
	4	You feel you have no control of your diabetes		
Hassle	1	You find your life with diabetes is never a hassle		
	2	You find your life with diabetes is sometimes a hassle		
	3	You find your life with diabetes is often a hassle		
	4	You find your life with diabetes is <u>always</u> a hassle		
Stress	1	You find your life with diabetes is never stressful		
	2	You find your life with diabetes is sometimes stressful		
	3	You find your life with diabetes is often stressful		
	4	You find your life with diabetes is <u>always</u> stressful		
Support (All support you	1	You feel totally supported with your diabetes		
have; from family, friends,	2	You feel you have a lot of support with your diabetes		
and health care	3	You feel you have a little support with your diabetes		
professionals)	4	You feel you have no support with your diabetes		

Fig. 1 - Health and self-management in diabetes classification system.

produce a design that enables estimation of the parameters in a prespecified regression model with precision. The design selected 120 choice sets across 10 survey versions (12 per survey) and was piloted (50 respondents) to generate priors for each attribute level of each dimension. These priors were then used to generate the design (i.e., select 120 choice sets) for the main study. The rationale for this was to improve the efficiency of the design with information regarding the magnitude of the parameter values.

Respondents

Presently, there is no consensus in the literature about whether values should be obtained from the general population or from a patient sample [30]. This article does not propose to review this debate, but given the diversity of viewpoints, values have been collected from both general population and individuals with diabetes.

General population sample

Respondents were recruited by a market research agency using an existing online panel. Respondents were selected to obtain a representative sample of the UK population in terms of age and sex. They were rewarded for their participation with a nominal amount of vouchers that could be accumulated and exchanged for goods.

Patient sample

Diabetic patients were recruited in various ways to ensure a good coverage across different groups in terms of severity, background, and setting:

 Posters in clinics and invitation letters posted with the usual clinic appointment letter to patients at Sheffield Teaching Hospitals;

- 2. Invitation letters posted to members of the Sheffield Diabetes Research Database who have previously provided consented to receive information about diabetes research projects;
- 3. Diabetes UK and DAFNE Online advertised a link to the online
- 4. Twitter and Facebook posts to diabetes charities, organizations, and the University of Sheffield staff and students and using a social media marketing company.

All patient respondents were offered optional entry into a prize draw for £50 shopping vouchers per 50 respondents.

The DCE Survey

Respondents were presented with the project information sheet and consented. They then completed sociodemographic and health questions asking about sex, age, education level, annual household income, and whether they have diabetes. After this, respondents were provided with information about what it is like to live with diabetes and were asked to complete the Health and Self-Management in Diabetes classification system for themselves if they had diabetes or imagine someone with diabetes to familiarize themselves with the classification system. Respondents were then given an introduction to the DCE questions (shown in Supplemental Materials found at http://dx.doi. org/10.1016/j.jval.2017.06.016). Respondents then answered 1 practice DCE question plus 12 DCE questions, where respondents chose between health description A and health description B, an example question of which is shown in Figure 2. Respondents were told to imagine that these health descriptions would last for 10 years and then they would die. Attributes that differed across the two health descriptions were highlighted in yellow. Finally, patients were also asked about their self-management of diabetes.

Piloting

The draft survey instrument was shown to a general patient and public involvement panel for comments and piloted with 50 members of the general population to inform the final survey design.

Analysis

Sociodemographic and self-reported health characteristics were analyzed and categorized.

The DCE data were modeled using the following model specification:

$$\mu_{ij} = \alpha_i + \beta_1 c_{ij} + \beta'_2 x_{ij} + \varepsilon_{ij}, \tag{1}$$

where c_{ij} represents cost, β_1 is the coefficient for costs, and β'_2 represents the coefficients for the 24 nonreference attribute levels of the classification system. This model produces unanchored values for scenarios or combinations of attribute levels. The monetary value (WTP) of each level of each attribute is estimated by dividing the coefficient attached to the relevant attribute level by the coefficient attached to the cost attribute (after previous checking for the linearity of the cost attribute) to generate the marginal rate of substitution, $\frac{\beta_{2ij}}{\beta_4} \frac{\beta_{2ij}}{\beta_4}$. This enables the monetary value of self-management alone, or the combined monetary value of HRQOL and self-management, to be estimated. Models were estimated separately for the general population and the diabetic patient samples. Models were estimated using the conditional logit model with robust standard errors, and confidence intervals and standard errors of the WTP estimates were determined using the delta method. Model performance was examined using sign, significance, and logical consistency of coefficients, log likelihood, and pseudo-R². The models assumed that cost was linear and a continuous variable. This was examined by modeling cost as a categorical variable and plotting the cost coefficients [31].

Robustness

Regressions were re-estimated excluding all responses to the DCE questions that were achieved within an implausibly short time period because it is likely that such quick responses indicate that these respondents are less likely to have read, understood, and considered the profiles. Regression models were also estimated to determine whether sociodemographic characteristics impacted on the results, including main effects plus interaction effects (one at a time) for the sociodemographic characteristics of age, sex, low income, high income, EQ-5D score, and whether the respondent has diabetes.

Results

The Sample

The characteristics of the general population and diabetic patient samples compared with those of the UK general population are presented in Table 1. The general population sample has a similar proportion of males and similar employment status to the UK general population, but has a lower proportion of respondents older than 65 years. The diabetic patient samples have a lower proportion of males, respondents with a degree qualification, employed respondents, and respondents in the higher income category; a larger proportion of respondents older than 65 years; and a lower EQ-5D-5L score (scored using the

	Health description A	Health description B	
	You <u>always</u> find yourself losing your temper over small things You <u>sometimes</u> worry about going hypo	You never find yourself losing your temper over small things You never worry about going hypo	
	You are <u>never</u> tired	You are <u>always</u> tired	
	Your days are <u>never</u> tied to meal times	Your days are <u>usually</u> tied to meal times	
	You feel you have some control of your diabetes You find your life with diabetes is never a hassle You find your life with diabetes is never stressful You feel totally supported with your	You feel you have <u>no</u> control of your diabetes You find your life with diabetes is <u>always</u> a hassle You find your life with diabetes is <u>always</u> stressful You feel you have a little support with	
	diabetes	your diabetes	
Which do you prefer?	The monthly cost of treatment to you is £10	The monthly cost of treatment to you is £75	

Fig. 2 – Example DCE question. DCE, discrete choice experiment.

Table 1 – Samples of respondents to the DCE survey,	by population, in	comparison with	the UK general
population.			

Characteristic	General population $(n = 1,497)$	Diabetic population $(n = 405)$	UK general population [†]
Sex, male	48.6%	41.5%	49.1%
Age (y)			
18–44	47.0%	40.0%	46.6%*
45–64	39.7%	35.3%	32.5%
>65	13.3%	24.7%	20.9%
Has a degree or equivalent professional qualification	52.9%	45.9%	
Main activity			
Employed	60.5%	45.4%	61.7%
Retired	18.0%	29.9%	13.9%
Housework	6.7%	5.7%	4.3%
Student	4.5%	3.7%	9.3%
Seeking work	1.3%	1.5%	
Unemployed	3.1%	3.0%	4.4%
Long-term sick	4.8%	9.4%	4.3%
Other	1.1%	1.5%	2.2%
EQ-5D-5L, mean \pm SD	0.79 ± 0.25	0.70 ± 0.27	
Have diabetes	13.2%	100.0%	
Have type 1 diabetes	2.5%	47.7%	
Have type 2 diabetes	10.6%	52.3%	
Annual household income (£)			
Up to 5,199	2.8%	4.7%	
5,200–10,399	6.1%	6.4%	
15,400–15,599	8.6%	10.9%	
15,600–20,799	8.5%	11.4%	
20,800–25,599	10.6%	11.4%	
26,000–31,199	10.2%	8.1%	
31,200–36,399	9.6%	5.9%	
36,400–51,999	15.5%	13.6%	
>52,000	18.5%	11.1%	
Prefer not to say	9.6%	16.5%	

DCE, discrete choice experiment; EQ-5D-5L, five-level EuroQol five-dimensional questionnaire.

cross-walk from EQ-5D-5L to EQ-5D [32]). For a single DCE task level, 37.1% of responses were completed within 10 seconds, 60.2% of responses were completed within 20 seconds, 94.1% of responses were completed within 1 minute, and 99.1% of responses were completed within 3 minutes.

Regression Analysis

Regression results for the conditional logit model with robust standard errors and the WTP values generated using the marginal rate of substitution are presented in Table 2. The specified model is acceptable for both the general population and the patient population using log likelihood and pseudo- R^2 . The WTP values reflect the amount that respondents would be willing to pay each month in pound sterling (£) to avoid decrement in the HRQOL or the self-management attributes.

The cost coefficient has the expected negative coefficient, showing that individuals are willing to pay to avoid decrements in health or self-management outcomes. Previous analyses (not reported) indicated that it was appropriate to assume cost was linear and continuous, using a plot for each sample of the cost levels and coefficients estimated using a model in which cost was included as a categorical variable.

The estimated coefficients of the HRQOL and the self-management attributes are all negative and logically consistent, with the exception of level 2 coefficients for mood, vitality, and support, although only the vitality level 2 coefficient is approaching significance (at the 10% level). Out of a possible 25 coefficients, 21 and 17 are statistically significant in the models in the general population and the diabetic patient population, respectively. All level 3 and level 4 coefficients are negative and significant with the exception of the level 3 coefficient for vitality in the model estimated for the diabetic patient population.

Table 3 presents the rank ordering of the level 4 coefficients for the general population and the diabetic patient samples, which indicate the ordering of which attributes have the largest impact on WTP at the most severe level. There are noticeable differences, particularly for vitality and mood.

Figure 3 shows the relative size of the WTP values for the general population and diabetic patient samples and shows the relative importance of each of the attributes. The size of the WTP values is similar to those of the HRQOL and the self-management attributes, implying that self-management and HRQOL are equally important. In general, the coefficients of the patient sample tend to be larger than those of the general population sample for more severe levels of each attribute, but smaller for the least severe level of each attribute. This suggests that

^{*} Age distribution is here reported as the percentage of all adults aged 18 y and older.

[†] Statistics for England in the Census 2011. The census includes persons aged 16 y and older, whereas this study surveys only persons aged 18 y and older.

Table 2 – Regression analysis of DCE survey responses, by p	population, reporting unanchored and WTP
estimates.	

Variable	General population unanchored estimates $(n=1497)$	Diabetic population unanchored estimates (n = 405)	General population WTP estimates (£)	Diabetic population WTP estimates (£)
Mood L2	0.0414	0.0033	16.52	2.79
Mood L3	-0.1226 [*]	-0.1094^{\dagger}	-48.91 [*]	-93.31 [†]
Mood L4	-0.2713 [*]	-0.3847 [*]	-108.25 [*]	-328.29 [*]
Hypoglycemia L2	-0.0413	-0.0605	-16.49	-51.60
Hypoglycemia L3	-0.1814 [*]	-0.1426^{\ddagger}	-72.37 [*]	-121.71 [‡]
Hypoglycemia L4	-0.2415 [*]	-0.2807 [*]	-96.38 [*]	-239.53 [*]
Vitality L2	0.0581 [†]	0.1075 [†]	23.18 [†]	91.71 [†]
Vitality L3	-0.3036 [*]	-0.1016	-121.14 [*]	-86.68
Vitality L4	-0.5918 [*]	-0.2770 [*]	-236.17 [*]	-236.37 [*]
Social limitations L2	-0.0205	-0.0153	-8.17	-13.06
Social limitations L3	-0.0977*	-0.1585 [*]	-38.97 [*]	-135.23 [*]
Social limitations L4	-0.1707 [*]	-0.2612 [*]	-68.12 [*]	-222.91 [*]
Control L2	-0.1427*	-0.0236	-56.94 [*]	-20.10
Control L3	-0.3010 [*]	-0.3372 [*]	-120.12 [*]	-287.72 [*]
Control L4	-0.4441 [*]	-0.5343 [*]	-177.22 [*]	-455.88 [*]
Hassle L2	-0.0971*	-0.0649	-38.73 [*]	-55.35
Hassle L3	-0.2328*	-0.2138 [*]	-92.89 [*]	-182.46*
Hassle L4	-0.4134*	-0.3563 [*]	-164.95 [*]	-304.05 [*]
Stress L2	-0.1543*	-0.0601	-61.56 [*]	-51.29
Stress L3	-0.2188*	-0.1465 [*]	-87.30 [*]	-125.03 [*]
Stress L4	-0.3715 [*]	-0.3230 [*]	-148.25 [*]	-275.61 [*]
Support L2	0.0506	0.0385	20.20	32.88
Support L3	-0.1266 [*]	-0.1859 [*]	-50.50 [*]	-158.61 [*]
Support L4	-0.2987*	-0.3438 [*]	-119.19 [*]	-293.38 [*]
Cost	-0.0025 [*]	-0.0012*		

Note. Pseudo- $R^2 = 0.160$ for general population model; pseudo- $R^2 = 0.101$ for diabetic population model. Log likelihood = -10458.4 for general population model; log likelihood = -3027.3 for diabetic population model.

DCE, discrete choice experiment; WTP, willingness to pay.

patients are willing to pay less to avoid mild health states but are willing to pay more to avoid severe health states in comparison with the general population. For example, the WTP to avoid the mild state of state 222222222 is £121.99 per month for the general population and £64.02 per month for the diabetic patient population. In contrast, the WTP to avoid the most severe state of 444444444 is £1118.53 per month for the general population and £2356.02 per month for the diabetic patient population.

Robustness

Robustness analyses excluded all responses for a given DCE task that were answered in less than 5 seconds (19% and 5% of responses for general population and patients, respectively) and in less than 10 seconds, respectively (40% and 16% of responses for general population and patients, respectively), resulting in larger WTP values at levels 3 and 4. This suggests that the

General po	pulation	Diabetic population		
Variable	WTP estimate (£)	Variable	WTP estimate (£	
Vitality L4	-236.17	Control L4	-455.88	
Control L4	-177.22	Mood L4	-328.29	
Hassle L4	-164.95	Hassle L4	-304.05	
Stress L4	-148.25	Support L4	-293.38	
Support L4	-119.19	Stress L4	-275.61	
Mood L4	-108.25	Hypoglycemia L4	-239.53	
Hypoglycemia L4	-96.38	Vitality L4	-236.37	
Social limitations L4	-68.12	Social limitations L4	-222.91	

^{*} P < 0.01.

 $^{^{\}dagger}$ P < 0.1.

 $^{^{\}ddagger}$ P < 0.05.

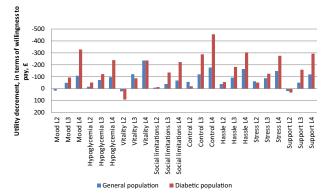


Fig. 3 – Anchored coefficients for the general population and diabetic patients.

exclusion of responses from respondents who may not have understood or engaged with the task will increase the size of the WTP estimates. Regression models including main effects plus interaction effects (one at a time) for the sociodemographic characteristics of age, sex, low income, high income, EQ-5D score, and whether the respondent has diabetes found no distinguishable pattern in terms of which coefficients were significant. This was surprising for the income variables, given that it was anticipated that WTP may increase as annual household income increases.

Discussion

This article describes the use of DCE including a cost attribute to value a classification system describing the impact of health and self-management of diabetes on quality of life. The classification system is distinct in that it provides a formal and consistent way to take account of self-management across different interventions and one that is not intervention-specific. Nevertheless, to use this information to inform health technology assessment, a means of weighting the severity levels of each attribute to produce a single score is required. This article describes a novel application of the technique of DCE involving a cost attribute to enable the valuation of a standardized classification system to provide WTP values for every state defined by the classification system. The article also provides a comparison of general population and diabetic population WTP values.

In the regression analysis, the WTP values for the HRQOL and the self-management attributes are of similar magnitude, suggesting that HRQOL and self-management have a similar relative impact in terms of their utility decrement or amount respondents are willing to pay to avoid more severe levels of the attribute. This has important implications for policy, because it suggests that the impact of changes in self-management on the lives of people with diabetes can be as important as any HRQOL improvement resulting from the change.

The largest difference in the rank ordering of the level 4 coefficients between the general population and the patient samples is the ranking of the vitality attribute, which is ranked the most important for the general population, whereas it is ranked the second least important for the patient population. This is interesting given that it is the attribute in the classification system derived from the generic short form 36 health survey, whereas the other items were all derived either from a diabetes-specific measure or from interviews with people with diabetes. Although this suggests that the general population may place a higher relative value on reductions in vitality than the diabetic patient population, the absolute value is similar for both samples.

In addition, the mood attribute is ranked the second most important for the diabetic population but the third least important for the general population. Nevertheless, both the hypoglycemic attack and the social limitations attributes are ranked among the least important for both populations.

The WTP per month to avoid state 44444444 is £1118.53 for the general population and £2356.02 for the diabetic patient population. This is significantly larger than the largest cost level included in the survey of £600 per month. An indication of how large these values are relative to annual household income can be calculated. Approximately 23.9% of the general population sample has an annual household income less than £20,800, and for these respondents this would represent at least 64.5% of their income (£13,422.36). Obviously, this is a somewhat unrealistic example because it includes the most severe state experienced by respondents with the lowest income, but indicates that the results should be interpreted with caution, because they may indicate only relative values rather than absolute values. It is possible that the values suffer from "hypothetical bias," when people state that they are willing to pay higher values than they would actually be willing to pay when using their own money in a laboratory or field experiment [33]. This is an important limitation of this type of study that is purely hypothetical.

One limitation of the study is that the patient sample is relatively small given that the survey design included 12 blocks of 10 DCE questions, meaning approximately 29 to 34 respondents answered each block of 10 questions. This may have impacted on the significance of coefficients for the regression models estimated on the patient samples.

The regression models had some insignificant and some inconsistent results. It is, however, important to note that it is not uncommon for valuation surveys of classification systems to find this; for example, the UK valuation of the SF-6D using standard gamble had both inconsistent and insignificant coefficients [4,5].

The design of the survey may have impacted on the results. The DCE tasks ask respondents to consider two health descriptions that contained 18 pieces of information (9 pieces per description). This is a large amount of information to simultaneously consider and could mean that respondents did not fully consider all information or all attributes. Other valuation surveys of classification systems have also included a large number of attributes (e.g., the survey by Norman et al. [34] included 11 attributes), and yet in the design these have fixed some attributes at the same level across both profiles in the DCE, and this is an option recommended for use in future surveys. It is possible that there may be interaction in preferences for some of the attributes, for example, mood and stress. Nevertheless, the study design selected choice sets assuming an additive model, meaning that interaction effects cannot be accurately estimated.

The use of existing general population online panels can be criticized for not being representative of the UK population. Members of online panels may differ from the general population in that they exclude the computer illiterate and those without access to the Internet. In addition, respondents have signed up to be members of a panel with a market research agency and stated that they are willing to answer surveys in return for points that can be exchanged for goods, and therefore their motivation for undertaking the survey may have an impact on their responses, potentially making their responses unrepresentative. Nevertheless, in terms of sociodemographic characteristics measured, they were found to be similar to the UK general population except that there being fewer who were older than 65 years and in contrast a larger proportion of retired respondents, although they may differ in terms of other unmeasured characteristics. The patient sample was different in ways that would be expected for those with diabetes.

The major limitation of this study is concerned with the accuracy of the elicitation of preferences for the cost attribute. The UK health care system is publicly funded, meaning that the general population receives free health care at the point of use with the exception of small charges such as prescription charges. This means that respondents are not used to paying for health care, or paying to improve their health state. This may have affected the results because it may mean that responses are not fully considered or fully informed because of inexperience in paying for health care or health improvement. It is possible that the selection of four levels rather than a higher number for the cost attribute may have also affected the results. In addition, respondents were forced to make a choice, meaning that they could not state indifference or have an "opt out" response indicating that they were not prepared to pay any cost for their health. This may have affected the results, because this does not reflect real life in which people can choose to not pay to improve their health. From a modeling perspective, however, an indifference or opt-out option is not beneficial, although it would indicate these respondents who may have chosen at random. One possibility is to include a zero cost level, and further research is encouraged to determine how this impacts on responses.

The DCE in this study was purposively designed to maximize statistical efficiency and minimize any dominance, for example, when one profile is clearly dominant or dominant with the exception of one attribute such as cost. Therefore, although we can observe within choice sets whether the low or high cost option was selected (where cost differs), we are unable to infer whether this selection was only due to cost, mainly due to cost, or despite cost. A better way to examine this question would be a binary choice survey with specifically selected scenarios to examine dominance of the cost attribute, a small think-aloud survey asking respondents the reasons for their choices directly, and/or research using eye tracking devices to determine where respondents focus their gaze when making their choices. This is an important issue for further research.

The wording of the cost attribute may have impacted on the results. In the profiles the wording was "The monthly cost of treatment to you." It is possible that respondents may not have correctly interpreted the cost as being applicable every month and may have instead interpreted it as a one-off payment. The frequency of payment as monthly rather than annually may have also had an effect as, for example, altering the wording to annual rather than monthly may not have multiplied the size of the WTP estimates by 12 times.

Conclusions

This article contributes to the methodological literature on the valuation of standardized patient-reported outcome measures using WTP for potential application in CBA. This approach could potentially add to the use of WTP in economic evaluation. The levels of the cost attribute used in the DCE survey were derived empirically, adding strength to the approach. The project also compared general population and diabetic patient values. Overall this approach shows promise, but there are some concerns with some inconsistencies in results, insignificant coefficients, and large WTP estimates observed that extend beyond the cost levels used in the survey. The novel application of DCE with a cost attribute to value a classification system provides an alternative approach to DCE with duration, an approach that has been previously used to value classification systems. Further research applying DCE with a cost attribute to value a classification system and to compare this method with DCE with duration is ongoing.

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Supplemental Materials

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REFERENCES

- Clark MD, Determann D, Petrou S, et al. Discrete choice experiments in health economics: a review of the literature. Pharmacoeconomics 2014;32:883–902.
- [2] Feher MD, Brazier J, Schaper N, et al. Patients' with type 2 diabetes willingness to pay for insulin therapy and clinical outcomes. BMJ Open Diabetes Res Care 2016;4:e000192.
- [3] Herdman M, Gudex C, Lloyd A, et al. Development and preliminary testing of the new five-level version of EQ-5D (EQ-5D-5L). Qual Life Res 2011;20:1727–36.
- [4] Brazier J, Roberts J, Deverill M. The estimation of a preference-based measure of health from the SF-36. J Health Econ 2002;21:271–92.
- [5] Brazier JE, Roberts J. The estimation of a preference-based measure of health from the SF-12. Med Care 2004;42:851–9.
- [6] Bansback N, Brazier J, Tsuchiya A, et al. Using a discrete choice experiment to estimate health state utility values. J Health Econ 2012;31:306–18.
- [7] Bansback N, Hole AR, Mulhern B, et al. Testing a discrete choice experiment including duration to value health states for large descriptive systems: addressing design and sampling issues. Soc Sci Med 2014;114:38–48.
- [8] Mulhern B, Bansback N, Brazier J, et al. Preparatory study for the revaluation of the EQ-5D tariff: methodology report. Health Technol Assess 2014;18:1–191: vii–xxvi.
- [9] Norman R, Cronin P, Viney R. A pilot discrete choice experiment to explore preferences for EQ-5D-5L health states. Appl Health Econ Health Policy 2013;11:287–98.
- [10] Norman R, Viney R, Brazier J, et al. Valuing SF-6D health states using a discrete choice experiment. Med Decis Making 2014;34:773–86.
- [11] Viney R, Norman R, Brazier J, et al. An Australian discrete choice experiment to value EQ-5D health states. Health Econ 2014;23:729–42.
- [12] National Institute for Health and Care Excellence. Guide to the Methods of Technology Appraisal. London: National Institute for Health and Care Excellence, 2013.
- [13] Pharmaceutical Benefits Advisory Committee. Guidelines for Preparing Submissions to the Pharmaceutical Benefits Advisory Committee. Canberra, Australia: Australian Department of Health, 2015.
- [14] Jendle J, Torffvit O, Ridderstrale M, et al. Willingness to pay for diabetes drug therapy in type 2 diabetes patients: based on LEAD clinical programme results. J Med Econ 2012;15(Suppl. 2):1–5.
- [15] Ratcliffe J. The use of conjoint analysis to elicit willingness-to-pay values. Proceed with caution? Int J Technol Assess Health Care 2000;16:270-5.
- [16] Ryan M. A comparison of stated preference methods for estimating monetary values. Health Econ 2004;13:291–6.
- [17] Ryan M, Watson V. Comparing welfare estimates from payment card contingent valuation and discrete choice experiments. Health Econ 2009;18:389–401.
- [18] Health and Social Care Information Centre. Prescriptions dispensed in the community, England 2005–2015. Available from: http://content. digital.nhs.uk/catalogue/PUB20664/pres-disp-com-eng-2005-15-rep.pdf. [Accessed July 26, 2017].

- [19] Diabetes UK. State of the nation 2016. Available from: www.diabetes. org.uk/Documents/Position%20statements/Diabetes%20UK%20State% 20of%20the%20Nation%202016.pdf. [Accessed August 26, 2016].
- [20] National Institute for Health and Care Excellence. NICE Guideline [NG28]: Type 2 Diabetes in Adults: Management. London: National Institute of Health and Care Excellence, 2015.
- [21] Welch GW, Jacobson AM, Polonsky WH. The problem areas in diabetes scale: an evaluation of its clinical utility. Diabetes Care 1997;20:760–6.
- [22] Carlton J, Elliott J, Rowen D, et al. Developing a questionnaire to determine the impact of self-management in diabetes: giving people with diabetes a voice. Health and Quality of Life Outcomes 2017;15:146.
- [23] Mulhern B, Labeit A, Rowen D, et al. Developing preference based measures for diabetes: DHP-3D and DHP-5D. Diabet Med [published online ahead of print May 6, 2017]. http://dx.doi.org/10.1111/dme.13377.
- [24] Meadows K, Steen N, McColl E, et al. The Diabetes Health Profile (DHP): a new instrument for assessing the psychosocial profile of insulin requiring patients—development and psychometric evaluation. Qual Life Res 1996;5:242–54.
- [25] Meadows KA, Abrams C, Sandbaek A. Adaptation of the Diabetes Health Profile (DHP-1) for use with patients with type 2 diabetes mellitus: psychometric evaluation and cross-cultural comparison. Diabet Med 2000:17:572–80.

- [26] Ware JE, Snow KK, Kolinski M, et al. SF-36 Health Survey Manual and Interpretation Guide. Boston, MA: The Health Institute, New England Medical Centre, 1993.
- [27] Mohamed AF, Zhang J, Johnson FR, et al. Avoidance of weight gain is important for oral type 2 diabetes treatments in Sweden and Germany: patient preferences. Diabetes Metab 2013;39:397–403.
- [28] Johnson FR, Mohamed AF, Ozdemir S, et al. How does cost matter in health-care discrete-choice experiments? Health Econ 2011;20:323–30.
- [29] ChoiceMetrics (2012) Ngene 1.1.1 User Manual & Reference Guide, Australia.
- [30] Brazier JE, Ratcliffe J, Salomon J, et al. Measuring and Valuing Health Benefits for Economic Evaluation. Oxford, UK: Oxford University Press. 2016.
- [31] Payne K, Fargher EA, Roberts SA, et al. Valuing pharmacogenetic testing services: a comparison of patients' and health care professionals' preferences. Value Health 2011;14:121–34.
- [32] van Hout B, Janssen MF, Feng YS, et al. Interim scoring for the EQ-5D-5L: mapping the EQ-5D-5L to EQ-5D-3L value sets. Value Health 2012;15:708-15.
- [33] Loomis J. What's to know about hypothetical bias in stated preference valuation studies? J Econ Surv 2011;25:363–70.
- [34] Norman R, Viney R, Aaronson NK, et al. Using a discrete choice experiment to value the QLU-C10D: feasibility and sensitivity to presentation format. Value Health 2016;25:637–49.