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# Social Valuation of EQ-5D Health States: The Chilean Case

Victor Zarate<sup>1,\*</sup>, Paul Kind<sup>1</sup>, Paulina Valenzuela<sup>2</sup>, Alberto Vignau<sup>2</sup>, Pedro Olivares-Tirado<sup>3</sup>, Alberto Munoz<sup>3</sup>

<sup>1</sup>Centre for Health Economics, The University of York, Heslington, York, United Kingdom; <sup>2</sup>DATAVOZ/STATCOM, Santiago, Chile; <sup>3</sup>Departamento de Estudios y Desarrollo, Superintendencia de Salud Chile, Santiago, Chile

ABSTRACT

Background: Cost-effectiveness analysis has been recommended by many national agencies around the world as a valid methodology to improve resource allocation within the health-care system. If the preferences of the society are taken into account in such a decision-making process, it is generally recommended that these values should be elicited by using a generic health-related quality-of-life instrument, such as the EuroQol five-dimensional (EQ-5D) questionnaire. Objectives: To estimate a set of social values for EQ-5D questionnaire based on the time trade-off valuation technique for use in Chile. Methods: A valuation questionnaire was applied to a probabilistic sample of 2000 individuals, aged 20 years or older, living in the Metropolitan region. The fieldwork took place during October to November 2008. Utility weights for 42 health states were calculated directly by the application of time trade-off. Several random effect and ordinary least-squares regression models were fitted

to these valuations to predict the full set of 243 health states generated by the EQ-5D system. The best model was chosen by applying criteria of parsimony, goodness of fit, and prediction capacity. **Results:** The selected regression model was robust and showed better predictive characteristics than others reported in similar studies conducted elsewhere. The chosen regression model showed a R² of 0.34, mean absolute error of 0.017, and high predictive capacity. **Conclusions:** This study provides an EQ-5D social value set for domestic use in Chile. Our results differ from those reported in other countries, justifying the need to perform local studies that adequately reflect societal health preferences.

**Keywords:** Chile, EQ-5D, health status, preference weights, time tradeoff (TTO).

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## Introduction

The valuation of EuroQol five-dimensional (EQ-5D) questionnaire health states in terms of their domestic social preferences is of primary importance for any country that intends conducting economic evaluation to inform high-level decisions regarding the allocation and use of scarce health-care resources. The problem facing many countries that lack such a domestic value set is that of identifying an appropriate alternative source. The EuroQoL Group provides some advice on this matter through its Web site, advocating that if it is not possible to identify a suitable "donor" source of EQ-5D values then the U.K. Measurement and Valuation of Health (MVH) value set should be considered as the default option. There are other alternatives of course, including the BIOMED visual analogue scale value set produced by pooling valuation data from several European countries. More recent and novel methods for estimating provisional value sets for EQ-5D have been proposed by using valuation exchange rates [1]. Within Latin America, there are two other options based on the analysis of Spanish-speaking Hispanics in the U.S. valuation survey [2] and the recently published results of an Argentine population survey [3]. Clearly, any values based on such second-best approaches are likely to be in error, but the extent of such errors remains unknown (and unknowable) until primary valuation data can be collected from a national population survey.

During 2008, a decision was made within the Chilean Superintendency of Health, a governmental institution responsible for the oversight of health insurances, to commission a valuation survey to calibrate EQ-5D for subsequent use in economic evaluation and health technology assessment applications within Chile. A specific requirement set by the Superintendency was that the study methodology should be based on the MVH protocol [4]. The timetable stipulated required speedy action to describe and document the methods, recruit and train interviewers, identify and recruit respondents, complete the fieldwork, undertake the analysis, and submit a final report.

This article deals primarily with the analysis of the valuation data generated in this Chilean survey and the selection of a model for use in estimating time trade-off (TTO) utilities for EQ-5D health states. In so doing, it raises a number of issues regarding the specification of such models and the means by which we might consider marginal differences between alternative models. These have wider significance for the EuroQoL Group, especially when faced with the future comparison of three- and five-level models, but also in examining alternative estimation models that are often based on somewhat different components.

Conflicts of interest: The authors have no conflicts of interest to report.

<sup>\*</sup> Address correspondence to: Victor Zarate, Centre for Health Economics, Alcuin "A" Block, The University of York, Heslington, York YO10 5DD LIK

E-mail: victor.zarate@york.ac.uk; E-mail: victor.zarate@york.ac.uk; vzarateb@gmail.com.

## **Methods**

#### Data and study population

Data analyzed in this research was obtained from the Chilean Valuation of the EuroQol EQ-5D Health States study that was carried out by the Research Department of the Superintendency of Health between October and November 2008. The target population was 4,627,801 millions civilians, noninstitutionalized adults, aged 20 years or more, who resided in the Metropolitan region that contains approximately 41% of the total Chilean population and also represents the geographical area where the capital of the country (Santiago) is located. Based on sociodemographic information obtained from the last available census conducted in 2002, a multistage probability sample of 2000 individuals was drawn from the target population. Sample-size estimations were based on the estimated number of respondents needed to detect a difference of 0.05 in mean TTO scores between two EQ-5D health states with a type I probability error of 5%, 80% power, and design effect of 1.2. TTO mean values and standard deviations used for the sample calculation were taken from the Spanish-speaking Hispanic respondents who participated in the U.S. Valuation of the EuroQol EQ-5D Health States study [2,5]. As in the U.S. study [5], data were collected through household interviews performed by 22 professional interviewers who were trained in two sessions by P.K. and V.Z. 1 month before the start of the fieldwork. No economic incentive was offered to the respondents before or after interviews. Subjects were excluded from the analysis if they had incomplete or inconsistent valuation data based on criteria reported elsewhere [5].

## EQ-5D

The EQ-5D is a standardized measure of health status that describes health by a classification system that comprises five dimensions (i.e., mobility, self-care, usual activities, pain/discomfort, and anxiety/depression) and three levels of severity for each dimension (i.e., no problem, moderate problem, and severe problems) [6]. A set of 243 health states is defined by this descriptive system, each one labeled by a unique five-digit code. Thus, the health state 11111 represents having no problems on any dimension and 33333 represents having severe problems in all five dimensions. To ease the valuation task for the respondents, a subset of 42 EQ-5D health states was sorted into 5 overlapping subsets of 12 health states following the same adapted MVH U.K. protocol used in the 2002 U.S. Valuation of the EuroQol EQ-5D Health States study [5]. Within these five sets, the state unconscious was removed given that it is not formally defined by the five-dimensional classification system and plays no part in any estimation model. The main instrument used in this study was the official Chilean EQ-5D questionnaire, provided by the EuroQol Group through its Web site http://www.euroqol.org. This official version was slightly modified in terms of the labeling of the "self-care" dimension in which the first level of severity was rephrased from "I have no problems with self-care" to "I have no problem washing or dressing myself." This modification was introduced to improve consistency within this category.

## TTO protocol

Respondents were asked to value one of the five subsets of EQ-5D health states selected at random by using the TTO elicitation protocol [7]. This methodology consists essentially in providing a con-

|                                      | Metropolitan region | Metropolitan $region_{wgt}$ | Chile                  |
|--------------------------------------|---------------------|-----------------------------|------------------------|
| Gender, % (n)                        |                     |                             |                        |
| Female                               | 62.0 (1239)         | 52.5 (1051)                 | 51.2 (5.8189.13)*      |
| Male                                 | 38.0 (761)          | 47.5 (949)                  | 48.8 (5.555.816)*      |
| Age, % (n)                           |                     |                             |                        |
| Mean (SE)                            | 46.83 (0.4)         | 43.46 (0.4)                 | 43.7 (16.1)*           |
| 20–44 years                          | 47.6 (953)          | 56.2 (1125)                 | 55.8 (6.345.981)*      |
| 45–64 years                          | 35.0 (699)          | 31.5 (629)                  | 31.5 (3.587.621)*      |
| 65+ years                            | 17.4 (348)          | 12.3 (246)                  | 12.7 (1.441.127)*      |
| Educational attainment, % (n)        |                     |                             |                        |
| <8 years                             | 15.8 (316)          | 13.1 (262)                  | 28.5 (2.834.392)†      |
| 8–12 years                           | 52.4 (1048)         | 52.2 (1045)                 | 47.1 (4.687.127)†      |
| 13+ years                            | 30.6 (611)          | 33.4 (669)                  | 24.4 (2.424.701)†      |
| Self-reported health problems, % (n) |                     |                             |                        |
| Mobility                             | 20.8 (416)          | 17.3 (347)                  | 15.4 <sup>††</sup>     |
| Self-care                            | 8.6 (173)           | 7.7 (153)                   | 3.9 <sup>††</sup>      |
| Usual activities                     | 16.0 (321)          | 14.0 (281)                  | 17.1 <sup>††</sup>     |
| Pain/discomfort                      | 44.2 (883)          | 39.6 (793)                  | 50.8 <sup>††</sup>     |
| Anxiety/depression                   | 33.9 (678)          | 30.9 (618)                  | 42.3 <sup>††</sup>     |
| Self-rated VAS, % (n)                |                     |                             |                        |
| Mean (SE)                            | 73.82 (0.46)        | 75.68 (0.46)                | 75.42 (0.3)††          |
| 81–100                               | 37.8 (757)          | 41.8 (837)                  | 42.3 <sup>††</sup>     |
| 61–80                                | 33.4 (667)          | 33.3 (666)                  | 32.2 <sup>††</sup>     |
| 41–60                                | 21.4 (429)          | 18.4 (368)                  | 18.2 <sup>††</sup>     |
| 21–40                                | 4.7 (94)            | 4.1 (82)                    | 6.3 <sup>††</sup>      |
| 0–20                                 | 2.6 (53)            | 2.3 (47)                    | $1.0^{\dagger\dagger}$ |

SE, Standard error; VAS, visual analogue scale.

<sup>\*</sup> Source: Population figures were estimated for June 2008, National Statistics Institute (INE).

<sup>&</sup>lt;sup>†</sup> Source: 2002 National Census, National Statistics Institute (INE).

<sup>††</sup> Source: 2005 EQ-5D health survey, Superintendency of Health.

tinuous set of two options for a given health state till the respondent reaches a point where he or she is indifferent to both alternatives. The first set of options is whether a given health state is better or worse than being dead. If the respondent thinks that the given health state is better than being dead, he or she is then asked to state whether it is preferred to live in that state for 10 years or to live in full health for x number of years (x  $\leq$  10); if the given health state is assessed as worse than dead, the two options offered are either to live in the given state for (10 - x, where x < 10) years followed by living in full health for x years or immediate death. TTO values are then bounded between -1 and 1 following a linear transformation that is reported in detail elsewhere [7].

## Statistical analysis

A number of random effect (RE) and linear regression models were used to analyze the TTO valuation data. The former approach takes into account the variability in responses within and between individuals. The latter, on the other hand, allows us to incorporate the survey design given that it is possible to apply sampling weights to correct for any imbalance in the achieved sample.

In terms of the modeling, the dependent variable was calculated as 1 minus the transformed TTO value assigned by each individual. Independent variables included a set of 10 dummy variables (i.e., M2, M3, Sc2, Sc3, Ua2, Ua3, Pd2, Pd3, Ad2, and Ad3) and a constant that represents having a problem in any of the five EQ-5D dimensions. In addition, extra independent variables were tested to account for interaction between different dimensions. These interaction terms were as follows:

- N2: whether there is any dimension on level 2;
- C2: the number of dimensions on level 2;
- C2sq: the square of the number of dimensions on level 2;

- N3: whether there is any dimension on level 3;
- C3: the number of dimensions on level 3;
- C3sq: the square of the number of dimensions on level 3;
- X2: whether there are two or more dimensions on level 2 or 3;
- X3: whether there are three or more dimensions on level 2 or 3;
- X4: whether there are four or more dimensions on level 2 or 3;
- X5: whether there are five dimensions on level 2 or 3.

Regression models with different combinations were individually tested in RE and multiply tested through a stepwise procedure in ordinary least-squares, without imposing restrictions on the number of additional interaction terms to be included. Goodness-of-fit statistics considered relevant in the analysis were Pearson's correlation coefficient between the observed and the predicted health state values (i.e.,  $\mathbb{R}^2$  overall), the mean absolute error (MAE) for predicting the 42 core EQ-5D health states, and the number of predictive errors greater than 0.025, 0.05, and 0.10. Normality of the residuals was analyzed by using scatter plots. Heteroskedasticity was explored by using the Breusch-Pagan test. Specification of the models was analyzed by using the Ramsey RESET test. Robustness of the model was assessed by randomly splitting the sample into two and using the predicted value set of one-half to estimate the observed values of the other half. Observed TTO values were compared with corresponding values from Argentina [3], Spanish-speaking Hispanics from the United States [2], Spain [8], and the United Kingdom [4]. All the statistical analyses were conducted by using Stata 10 [9].

#### Results

Completed interviews were obtained from 2000 individuals after 5008 household visits. From the total of households, 21% of the

| Table 2 – Parameter estimates and fit statistics for alternative random effect (RE) and ordinary least-squares (OLS) regression models in the Chilean valuation study. |               |               |                |                   |                 |                    |    |
|--|---------------|---------------|----------------|-------------------|-----------------|--------------------|----|
|  | Basic (RE)    | N3 (RE)       | C3sq (RE)      | C3sq +<br>X5 (RE) | C3sq (OLS)      | C3sq + X5 (OLS)    |    |
|  | Coefficient P | Coefficient P | Coefficient 1  | Coefficient       | P Coefficient P | Coefficient P      |    |
| MO2  | 0.128 (0.007) | 0.124 (0.007) | 0.114 (0.007)  | 0.108 (0.008)     | 0.121 (0.010)   | 0.115 (0.010)      |    |
| MO3  | 0.310 (0.010) | 0.300 (0.010) | 0.452 (0.012)  | 0.448 (0.012)     | 0.454 (0.017)   | 0.449 (0.017)      |    |
| SC2  | 0.130 (0.007) | 0.136 (0.007) | 0.126 (0.007)  | 0.118 (0.008)     | 0.129 (0.010)   | 0.121 (0.010)      |    |
| SC3  | 0.312 (0.010) | 0.289 (0.010) | 0.425 (0.012)  | 0.421 (0.012)     | 0.431 (0.016)   | 0.428 (0.016)      |    |
| UA2  | 0.178 (0.008) | 0.130 (0.008) | 0.135 (0.008)  | 0.126 (0.008)     | 0.129 (0.010)   | 0.119 (0.011)      |    |
| UA3  | 0.342 (0.010) | 0.250 (0.011) | 0.402 (0.010)  | 0.411 (0.010)     | 0.404 (0.014)   | 0.413 (0.014)      |    |
| PD2  | 0.107 (0.008) | 0.125 (0.008) | 0.116 (0.008)  | 0.110 (0.008)     | 0.108 (0.009)   | 0.103 (0.009)      |    |
| PD3  | 0.301 (0.009) | 0.251 (0.009) | 0.403 (0.010)  | 0.398 (0.010)     | 0.403 (0.013)   | 0.397 (0.013)      |    |
| AD2  | 0.094 (0.007) | 0.099 (0.007) | 0.105 (0.007)  | 0.100 (0.007)     | 0.114 (0.010)   | 0.108 (0.010)      |    |
| AD3  | 0.246 (0.008) | 0.196 (0.009) | 0.360 (0.010)  | 0.353 (0.010)     | 0.367 (0.013)   | 0.359 (0.014)      |    |
| Intercept  | 0.107 (0.010) | 0.070 (0.010) | 0.080 (0.010)  | 0.092 (0.012)     | 0.083 (0.008)   | 0.096 (0.008)      |    |
| N3   |               | 0.184 (0.011) |                |                   |                 |                    |    |
| C3sq   |               |               | -0.025 (0.001) | -0.027 (0.001)    | -0.026 (0.002)  | -0.028 (0.002)     |    |
| X5   |               |               |                | 0.049 (0.013)     |                 | 0.052 (0.017) 0.00 | )3 |
| R <sup>2</sup> overall   | 0.337         | 0.342         | 0.344          | 0.344             | 0.346           | 0.346              |    |
| Mean absolute error  | 0.048         | 0.036         | 0.02           | 0.017             | 0.021           | 0.016              |    |
| No. (of 42) > 0.025  | 30            | 27            | 18             | 10                | 18              | 10                 |    |
| No. (of 42) > 0.05   | 17            | 12            | 2              | 1                 | 2               | 2                  |    |
| No. (of 42) > 0.10   | 4             | 0             | 0              | 0                 | 0               | 0                  |    |

Note: All coefficients significant at P < 0.001 unless otherwise stated. Standard errors are given in parentheses. R2 overall represents the correlation between observed and predicted time trade-off values.

The official report of the Chilean valuation study included a recommendation to adopt a model in which four dummy variables were used corresponding to the number of level 3 elements in each health state (F13, F23, F33, and F43). F13 was set to 1 if there was one level 3 element, else 0. The C3 squared term captures a similar effect more efficiently. Since the results are virtually identical, the more parsimonious model is shown here.

addresses were ineligible and only 7% of the individuals did not respond to the survey, resulting in a response rate of 40%.

Sociodemographic and self-reported health characteristics of the achieved sample are described and compared with national figures in Table 1. After the application of corrective weights to the raw data, age and gender distribution of the sample (i.e., 52.5% female and 43.46 years mean age) closely resemble the characteristics of the country as a whole. The level of education as expected tends to be higher in the Metropolitan region when compared with the national figure. In terms of self-reported health characteristics, the highest prevalence of any EQ-5D health problem was in the pain/discomfort and anxiety/depression dimensions, with 39.6% and 30.9%, respectively. Higher prevalence on these two dimensions has been reported previously for Chile in a nonprobabilistic sample of the population selected by quota in 2005 [10]. Both the mean and the distribution of visual analogue scale scores appear to be extremely similar between the Metropolitan region and Chile.

On the basis of the data completeness and logical consistency criteria previously described, the final sample included 1967 respondents. Several RE and ordinary least-squares models were developed to fit the TTO valuations; models only with the highest goodness of fit are reported in Table 2. In both types of regression models, the best functional form comprised a constant, the basic 10 dummy variables for the five dimensions, plus two extra interaction terms: C3sq and X5. Only C3sq yielded a negative estimate, which produces a positive quadratic adjustment according to the number of level 3 in a given EQ-5D health state. As expected, RE models generally produced a slightly better goodness of fit in comparison with the ordinary leastsquares model. The C3sq + X5 was the best-performing RE model of all, with an R<sup>2</sup> of 0.344, MAE of 0.017, and lowest number of errors greater than 0.025 (i.e., 10 of 42) and 0.05 in absolute magnitude (i.e., 1 of 42). Directly observed and predicted values for the 42 health states are presented in Table 3.

Residuals of the C3sq + X5 (RE) model do not appear normally distributed in a normal probability plot showing a slightly

| Health state | and predicted values for 42 health states based on the C3sq + Observed |                  |                | Predicted                    |                |
|--------------|--|------------------|----------------|------------------------------|----------------|
|              | n  | Mean             | Standard error | Mean                         | Absolute error |
| 11112        | 787  | 0.765            | 0.013          | 0.808                        | 0.043          |
| 11113        | 391  | 0.616            | 0.025          | 0.582                        | 0.034          |
| 11121        | 787  | 0.809            | 0.010          | 0.798                        | 0.011          |
| 11122        | 394  | 0.689            | 0.020          | 0.698                        | 0.009          |
| 11131        | 390  | 0.570            | 0.026          | 0.537                        | 0.033          |
| 11133        | 790  | 0.266            | 0.024          | 0.265                        | 0.001          |
| 11211        | 785  | 0.790            | 0.011          | 0.782                        | 0.001          |
| 11312        | 394  | 0.438            | 0.030          | 0.424                        | 0.014          |
| 12111        | 786  | 0.775            | 0.012          | 0.790                        | 0.014          |
| 12121        | 391  | 0.773            | 0.012          | 0.680                        | 0.013          |
|              |  |                  |                |                              | 0.032          |
| 12211        | 394  | 0.654            | 0.022          | 0.664                        |                |
| 12222        | 787  | 0.467            | 0.020          | 0.454                        | 0.013          |
| 12223        | 391  | 0.250            | 0.034          | 0.228                        | 0.022          |
| 13212        | 394  | 0.270            | 0.034          | 0.288                        | 0.018          |
| 13311        | 788  | 0.190            | 0.025          | 0.184                        | 0.006          |
| 13332        | 391  | -0.195           | 0.034          | -0.179                       | 0.016          |
| 21111        | 786  | 0.799            | 0.012          | 0.800                        | 0.001          |
| 21133        | 392  | 0.107            | 0.036          | 0.157                        | 0.050          |
| 21222        | 787  | 0.457            | 0.020          | 0.464                        | 0.007          |
| 21232        | 394  | 0.226            | 0.034          | 0.203                        | 0.023          |
| 21312        | 392  | 0.328            | 0.032          | 0.316                        | 0.012          |
| 21323        | 391  | 0.062            | 0.035          | 0.034                        | 0.028          |
| 22112        | 394  | 0.589            | 0.025          | 0.582                        | 0.007          |
| 22121        | 790  | 0.593            | 0.017          | 0.572                        | 0.021          |
| 22122        | 394  | 0.471            | 0.028          | 0.472                        | 0.001          |
| 22222        | 394  | 0.276            | 0.032          | 0.297                        | 0.021          |
| 22233        | 394  | -0.146           | 0.035          | -0.136                       | 0.010          |
| 22323        | 392  | -0.127           | 0.035          | -0.133                       | 0.006          |
| 22331        | 788  | -0.026           | 0.025          | -0.019                       | 0.007          |
| 23232        | 394  | -0.174           | 0.033          | -0.186                       | 0.012          |
| 23313        | 790  | -0.143           | 0.024          | -0.142                       | 0.001          |
| 23321        | 789  | -0.065           | 0.024          | -0.034                       | 0.031          |
| 32211        | 392  | 0.234            | 0.034          | 0.243                        | 0.009          |
| 32223        | 391  | -0.228           | 0.033          | -0.188                       | 0.040          |
| 2232         | 391  | -0.232           | 0.033          | -0.223                       | 0.009          |
| 32313        | 394  | -0.177           | 0.034          | -0.179                       | 0.002          |
| 32331        | 394  | -0.177<br>-0.225 | 0.033          | -0.224                       | 0.002          |
| 33212        | 392  | -0.223<br>-0.070 | 0.036          | -0.22 <del>4</del><br>-0.079 | 0.001          |
| 33232        | 391  | -0.328           | 0.033          | -0.391                       | 0.063          |
|              |  |                  |                |                              |                |
| 33321        | 790  | -0.244           | 0.023          | -0.239                       | 0.005          |
| 33323        | 394  | -0.424           | 0.030          | -0.452                       | 0.028          |
| 33333        | 1965   | -0.494           | 0.012          | -0.497                       | 0.003          |

Table 4 – Chilean predicted preference weights for 243 EuroQol five-dimensional questionnaire health states based on the C3sq + X5 (random effect) model. Value Value Value Value State State State State State Value 11111 1 12322 0.196 21233 0.031 23221 0.17 32132 -0.0480.021 11112 0.808 12323 0.024 21311 0.416 23222 32133 -0.16611113 0.582 12331 0.089 21312 0.316 23223 -0.15132211 0.243 11121 0.798 12332 -0.01121313 0.144 23231 -0.03732212 0.143 11122 0.698 12333 -0.12921321 0.306 23232 -0.18632213 -0.0290.472 0.514 21322 0.206 23233 -0.30432221 0.133 11123 13111 11131 0.537 13112 0.414 21323 0.034 23311 0.076 32222 -0.01623312 32223 11132 0.437 13113 0.242 21331 0.099 -0.024-0.1880.265 13121 0.404 21332 -0.00123313 -0.14232231 -0.07411133 11211 0.782 13122 0.304 21333 -0.11923321 -0.03432232 -0.22311212 0.682 13123 0.132 22111 0.682 23322 -0.18332233 -0.34111213 0.456 13131 0.197 22112 0.582 23323 -0.30132311 0.039 11221 0.672 13132 0.097 22113 0.356 23331 -0.18732312 -0.061-0.17911222 0.572 13133 -0.02122121 0.572 23332 -0.33632313 11223 0.346 13211 0.388 22122 0.472 23333 -0.432321 -0.07111231 0.411 13212 0.288 22123 0.246 31111 0.487 32322 -0.2211232 0.311 13213 0.116 22131 0.311 31112 0.387 32323 -0.33811233 0.139 13221 0.278 22132 0.211 31113 0.215 32331 -0.22411311 0.524 13222 0.178 22133 0.039 31121 0.377 32332 -0.37313223 32333 11312 0.424 0.006 22211 0.556 31122 0.277 -0.43711313 0.252 13231 0.071 22212 0.456 31123 0.105 33111 0.147 11321 0.414 13232 -0.02922213 0.23 31131 0.17 33112 0.047 11322 0.314 13233 -0.14722221 0.446 31132 0.07 33113 -0.07111323 0.142 13311 0.184 22222 0.297 31133 -0.04833121 0.037 11331 0.207 13312 0.084 22223 0.071 31211 0.361 33122 -0.06311332 0.107 13313 -0.03422231 0.185 31212 0.261 33123 -0.18111333 -0.01113321 0.074 22232 0.036 31213 0.089 33131 -0.11612111 0.79 13322 -0.02622233 -0.13631221 0.251 33132 -0.21612112 0.69 13323 -0.14422311 0.298 31222 0.151 33133 -0.2812113 0.464 13331 -0.07922312 0.198 31223 -0.02133211 0.021 12121 0.68 13332 -0.17922313 0.026 31231 0.044 33212 -0.07912122 0.58 13333 -0.24322321 0.188 31232 -0.05633213 -0.19712123 0.354 21111 0.8 22322 0.039 31233 -0.17433221 -0.08912131 0.419 21112 0.7 22323 -0.13331311 0.157 33222 -0.23812132 0.319 21113 0.474 22331 -0.01931312 0.057 33223 -0.35612133 0.147 21121 0.69 22332 -0.16831313 -0.06133231 -0.24212211 0.664 21122 0.59 22333 -0.28631321 0.047 33232 -0.3910.406 12212 0.564 21123 0.364 23111 31322 -0.05333233 -0.45512213 0.338 21131 0.429 23112 0.306 31323 -0.17133311 -0.12912221 0.554 21132 0.329 23113 0.134 31331 -0.10633312 -0.22912222 0.454 21133 0.157 23121 0.296 31332 -0.20633313 -0.29312223 0.228 21211 0.674 23122 0.196 31333 -0.2733321 -0.23921212 12231 0.293 0.574 23123 0.024 32111 0.369 33322 -0.38821213 0.089 32112 33323 -0.45212232 0.1930.348 23131 0.269 12233 0.021 21221 0.564 23132 -0.01132113 0.097 33331 -0.33812311 0.406 21222 0.464 23133 -0.12932121 0.259 33332 -0.48712312 0.306 21223 0.238 23211 0.28 32122 0.159 33333 -0.49712313 0.134 21231 0.303 23212 0.18 32123 -0.013

bow-shaped pattern that indicates that residuals have excessive skewness. The Breusch–Pagan test produced a P value of <0.001 that rejects the null hypothesis that the variance of the residuals is homogeneous and therefore confirms the heteroscedasticity of residuals. The Ramsey RESET test indicated that all tested models suffer to some extent from misspecification, which is a common finding in these types of models given that no independent variables are related to particular characteristics of the respondent. The C3sq + X5 (RE) models proved to be robust given that the value set estimated on a randomly selected half of the sample closely predicted the values of the other half with an MAE of 0.028 and a high, significant correlation ( $R^2$ = 0.996). The full set of Chilean preference weights for

21232

0.203

23213

0.008

32131

12321

0.296

the 243 EQ-5D health states based on the selected C3sq + X5 (RE) model is provided in Table 4.

0.052

Observed TTO values' comparison between different countries showed that Chilean valuations tend to be lower than those previously reported in Argentina [3] and in the United States by the Spanish-speaking Hispanics [2] (Fig. 1). Nonetheless, when Chilean TTO values were compared against valuations collected in Spain [8] and the United Kingdom [4], they showed a surprisingly high level of agreement across the entire EQ-5D health spectrum (albeit for the subset of 42 states for which values had been directly elicited), being the closest to the Spanish values with a mean absolute difference of 0.088 ( $R^2$ = 0.976).

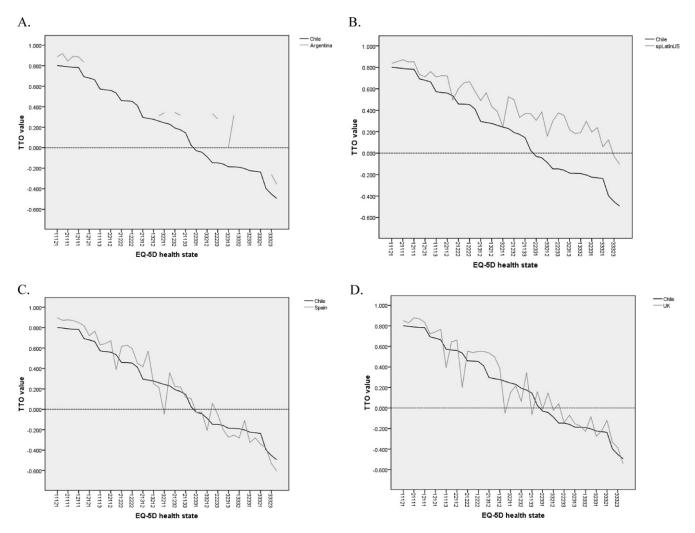


Fig. 1 – Comparison of observed time trade-off (TTO) values in Chile versus Argentina (A), Spanish-speaking Hispanics in the United States (B), Spain (C), and the United Kingdom (D). Note: Comparisons are based on 42 EuroQol five-dimensional (EQ-5D) questionnaire health states with the exemption of Argentina where only 22 health states were valued.

## Discussion

To the best of our knowledge, this study represents only the second EQ-5D valuation project to be performed in South America after the experience reported in Argentina [3]. The sample frame in our case was limited to the largest geographical area of the country by budget constrains and the explicit requirement of the Superintendency of Health. Because of this geographic restriction, we believed that it would be convenient to conduct further research to assess the impact that this constraint could have had on the overall results of this study. The Superintendency also set as a condition that the Chilean EQ-5D valuation study should follow the MVH protocol [4] in terms of the number of states selected for valuation tasks. This particular requirement left open the question of whether or not the development of a more efficient study protocol for use in Latin America is possible.

In the Chilean EQ-5D valuation study, only minor modifications to the original U.K. protocol were made: 1) Respondents valued a randomly selected fixed set of EQ-5D health states; these sets were previously used in the U.S. Valuation of EQ-5D Health States study; 2) the state unconscious was not considered in any valuation task; and 3) the official Chilean EQ-5D questionnaire was slightly modified for this study in its "self-care" dimension in which the first level of severity was rephrased from "I have no problems with self-care" to "I have no problem washing and

dressing myself." The decision of modifying the first level of severity on this dimension was based on feedback given by the interviewers during the training sessions. Although no psychometric tests were performed to support the implementation of this minor change, the research group thought that this modification would improve the consistency in the dimension's descriptions without critically compromising the structure of the questionnaire.

In terms of regression modeling, the development of the model C3sq  $\,+\,$  X5 provided the best fit for the valuation data with few prediction errors based on both REs and ordinary least-squares regressions. Given the inherent correlation structure of the valuation data, the former outperformed the latter in terms of goodness of fit. Corrective weights do not significantly alter the beta coefficients when applied to raw data; nonetheless, they did produce few logical inconsistencies in the full 243 estimated EQ-5D value set and therefore the C3sq  $\,+\,$  X5 (RE) model was selected as the preferred one.

All the regression models produced residuals that were not normally distributed; nonetheless, this condition is not required in order to obtain unbiased estimates of the regression coefficients. Heteroscedascity of the residuals was found in the selected C3sq  $\pm$  X5 (RE) model and was accordingly corrected through the estimation of robust standard errors. Splitting the sample randomly into two and using one half to predict the values of the

other half confirmed robustness of the chosen model with high correlation between estimated and predicted values and an MAE of 0.028.

Comparison of observed TTO values between countries allows the analysis of valuations without the noise introduced by modeling techniques. Chilean values differ clearly from the ones recently reported in Argentina [3] and in the United States by the Spanish-speaking Hispanics [2]. Despite the fact that only 22 EQ-5D health states were assessed in Argentina, the distribution of values looks very similar to the Hispanic community in the United States. The opposite occurs when Chilean values are graphically compared with those from Spain and the United Kingdom, which both show extremely congruent results. Similarities are closer to the Spanish TTO values with a mean absolute difference of 0.088. These findings give support to the idea that different societies value health status differently and this process is not related necessarily to geographical areas or culture but may be associated with other unobserved variables. To improve the understanding of differences in health preferences between different societies, more research based on the microlevel analysis of multinational studies is urgently needed.

## Conclusion

This study generated a preference-weighting system for the EQ-5D health states in Chile. The chosen C3sq + X5 (RE) regression model produces a value set with good fit, an MAE of 0.017, and only one prediction error exceeding 0.05 in absolute magnitude. We hope that our results will contribute to the development of cost-utility analysis in Chile and encourage other countries in the region to

perform similar studies in their societies with the aim of improving decision-making processes in Latin America.

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