Valuing health-related quality of life using a hybrid approach: Tunisian value set for the EQ-5D-3L

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Abstract
Objective To develop a value set for EQ-5D-3L based on the societal preferences of the Tunisian population.
Methods A representative sample of the Tunisian general population was obtained through multistage quota sampling involving age, gender and region. Participants (n = 327), aged above 20 years, were interviewed using the EuroQol Portable Valuation Technology in face-to-face computer-assisted interviews. Participants completed 10 composite time trade-off (cTTO) and 10 discrete choice experiments (DCE) tasks. Utility values for the EQ-5D-3L health states were estimated using regression modeling. The cTTO and DCE data were analyzed using linear and conditional logistic regression modeling, respectively. Multiple hybrid models were computed to analyze the combined data and were compared on goodness of fit measured by the Akaike information criterion (AIC).
Results A total of 300 participants with complete data that met quality criteria were included. All regression models showed both logical consistency and significance with respect to the parameter estimates. A hybrid model accounting for heteroscedasticity presented the lowest value for the AIC among the hybrid models. Hence, it was used to construct the Tunisian EQ-5D-3L valuation set with a range of predicted values from –0.796 to 1.0.
Conclusion This study provides utility values for EQ-5D-3L health states for the Tunisian population. This value set will be used in economic evaluations of health technologies and for Tunisian health policy decision-making.

Keywords EQ-5D-3L · Tunisian value set · Composite time trade-off · Discrete choice experiment · Health-related quality of life · Health measurement

Background
Health Technology Assessment (HTA) has been increasingly used to inform healthcare policy decisions [1]. Through systematic evaluation of health interventions, decision-makers can support their policies in health care resource allocation by seeking evidence of efficiency in resource use. The World Health Organization recommends the use of HTA as a tool to support reimbursement decision-making and, e.g., in drug price-setting negotiations [2]. In addition, HTA can encourage a more efficient patient-centered approach when allocating resources by considering population-specific needs in decision-making [3].

The use of health economic evaluations in low- to middle-income countries remains limited due to a lack of expertise and to poor local data [4, 5]. In Tunisia, price negotiations for new therapies are part of the drug licensing process and involve many stakeholders such as the Ministry of Health,
the Ministry of Trade, the National Health Insurance fund and the Tunisian central pharmacy. This process mainly uses external reference pricing as a benchmark. Since 1997, cost-effectiveness is one of the elements to be taken into consideration during drug licensing, price negotiations and reimbursement policy, especially for expensive and innovative drugs [6]. With the recent establishment of a national HTA agency, the National Authority for Assessment and Accreditation in Healthcare “INEAS”, Tunisia is taking a step towards implementing HTA in its health policy.

The quality-adjusted life year (QALY) is the standard outcome measure in health care economic evaluations, as recommended in current cost-utility analysis guidelines [7–9]. In this type of economic analysis, the incremental cost of a health technology is compared to the incremental health improvement expressed in QALYs, which can reflect the benefit of an intervention on both life expectancy and health-related quality of life (HRQOL). QALYs are calculated by multiplying life years by a correction factor: a health utility value attributed to each health state. The utility scale is anchored at 0 indicating a preference equal to immediate death, and 1 for preferences equal to full health. Negative values are assigned to states considered to be worse than dead (WTD). Utility weights are obtained by valuing how individuals perceive the quality of life associated with health states from a generic preference-based measure (GPBM). The most commonly used GPBM is EQ-5D [10]. This instrument was developed to measure, compare and value health status across disease areas [11]. The EQ-5D in its original version, EQ-5D-3L, defines \(3^5 = 243\) different states in its descriptive system across five dimensions—mobility (MO), self-care (SC), usual activities (UA), pain or discomfort (PD), and anxiety or depression (AD)—each with three levels of problems (none, some and extreme problems/unable to) [11]. Every health state is described using a unique digit code, (e.g., ‘11122’ refers to a health state with no problems in MO, SC and UA, but some problems in PD and AD).

Utilities can be assigned to EQ-5D-3L health states by using a value set. These value sets or tariffs are constructed by eliciting preferences from the general population for the health states using a valuation method such as the Time Trade-Off (TTO) [12]. The EuroQol Group’s current protocol for the valuation of health states, the EQ-VT, uses composite Time Trade-Off (cTTO) and discrete choice experiments (DCE) to determine utility values for EQ-5D health states [13]. Although the EQ-5D-3L descriptive system has been translated into Arabic and validated during a previous study [14], its use in economic evaluation remains limited due to the absence of a value set in Tunisia and other Arabic-speaking countries. Acknowledging the logistical complexity of 5-Level valuation studies, we aimed to develop a value set for EQ-5D-3L based on the preferences of the Tunisian population as a first step towards patient-centered research.

**Methods**

The study and the data collection were conducted in compliance with the EuroQol Group’s valuation protocol, EQ-VT [13]. Minor changes were made to the study design, as the protocol was originally designed for the EQ-5D-5L, and also in order to accommodate the local research teams’ constrained resources. As in previous EQ-5D valuation studies using the EQ-VT protocol, two preference elicitation techniques were used: cTTO and DCE. The questionnaire was administered via computer-assisted personal interviews using the EuroQol portable valuation technology v1.7 (EQ-PVT), power point-based software similar to the EQ-VT v2.1 software [13]. Cyclic data quality control (QC) was performed to ensure interviewers’ compliance to the protocol [15]. The methods and analyses in this paper comply with the CREATE guidelines for reporting valuation studies of multiattribute utility-based instruments [16].

**Sample selection and recruitment strategy**

The EuroQol protocol requires a sample size of 300 participants [17]. In order to ensure an adequate number of valid responses, a larger sample of 350 respondents was targeted: non-institutionalized individuals from the general population, aged above 20 years and able to read, comprehend and complete the interviews, were eligible to participate in the study.

In order to preserve the representativeness of the population, multistage quota sampling was performed proportional to the region of residency. The national territory was divided into six regions following the Tunisian census: northwest, northeast, center east, center west, southeast and southwest. We also used a quota sampling in each region in terms of age and gender as both these factors showed evidence of being related to health state values [18, 19]. The sample quotas were based on the latest Tunisian population census of 2014 [20]. We used a mixed recruitment strategy, through the personal contacts of interviewers and their relatives and by direct approaches in public spaces such as coffee and other shops, and universities.

**Health state selection**

Since the study combined cTTO and DCE tasks, an experimental hybrid design was developed in order to minimize the number of states and respondents required to obtain significant statistical estimates [13, 17]. The cTTO module included 28 health states, divided into 3 blocks of 10
EQ-5D-3L health states. Each block included the state ‘33333’ and 9 different states, at least one of them a mild state with only one deviation from full health. Eighteen of these health states were obtained using a 2-(3,5,2) orthogonal array distributed among the different blocks [21, 22]. Ten non-orthogonal states, including mild and intermediate states, were added according to block severity level balancing (i.e., the sum of health states severity index is equal in each block) and to minimum overlapping (i.e., reducing the number of health states with similar levels on the same attribute) [23].

The DCE module consisted of 60 pairs of states divided into 6 blocks. Health states were selected following the design developed by Stolk et al [24]. Each participant was randomly assigned to one of the DCE and TTO blocks. The order of the health states and choice pairs was randomized within the cTTO and DCE tasks. The feedback module was omitted to limit the interview time duration and reduce the risk of a participant dropping out of the interview.

Eliciting preferences

Composite TTO uses two different approaches to value health states: conventional TTO for better than dead (BTD) health states and lead-time TTO (LT-TTO) for WTD states [25]. Following the QALY scale, health utility values range on a scale anchored at 0 (death) and 1 (full health) and bound to -1 for WTD states [26]. In conventional TTO, the utility value (u) ranges from 0 to 1, and is equal to x/10 where x is the number of years in full health when the participant states the indifference point. For WTD health states, values are calculated taking into account the lead-time of 10 years, u = (x - 10)/10. Hence, u ranges from -1 (trading all years of the lead-time) to 0. During the cTTO task, an interactive visual scale aid allowed the participant to adjust to the number of years traded.

In the DCE task, participants were asked to choose their preferred alternative from two impaired EQ-5D-3L health states with no specification with respect to time duration.

Interviews started with a general presentation of the study and its purposes. Informed consent was obtained from all respondents and general background questions (i.e., age, sex and region of residency) were collected before starting the survey. Respondents were asked to report on their own health using the EQ-5D-3L questionnaire and the visual analogue scale (EQ VAS). Subsequently, they started the cTTO task and received instructions on how it works. The respondents first completed 3 wheelchair examples, in which both the BTD and the WTD tasks were shown to them. Next, the respondents completed three practice cTTO questions using EQ-5D-3L health states that were not included in the design. Finally, all respondents completed 10 cTTO tasks for health states from the block of health states they were assigned. Upon completion of the cTTO, the respondents received instructions on the DCE tasks, and subsequently completed 10 of these. Lastly, the respondents completed a socio-demographic questionnaire, after which they were thanked for their participation and debriefed. Each respondent received compensation.

Quality control process

The quality control process enacted was based on the EuroQol QC protocol [15]. During the interviews, the EQ-PVT software allows the collection of metadata similar to that collected when using the EQ-VT software. Overall, the data were assessed for protocol compliance, interviewer effects, and for face validity.

Five interviewers were recruited among Pharm.D students at the Faculty of Pharmacy of Monastir. They were trained by the principal investigators (CD and HF) during a 3-day workshop using training materials provided by the EuroQol Group. After completing the training, each interviewer performed five practice interviews for which the data were retained if all the quality criteria in data collection were met, as shown in Fig. 1. After validation of the interviewers, continuous monitoring of data quality was performed through cyclic QC (i.e., QC after every round of data collection) by the EuroQol support team (BR and FAS). This was followed by a discussion including general and individual feedback of the interviewers’ performances.

Statistical analysis

Sample characteristics

Descriptive statistics (frequencies and percentages) were used to report the sample socio-demographic characteristics (age group, region of habitat, gender, marital status, education level, employment status and health coverage) and compare them with national values. Self-reported health status on the EQ-5D-3L descriptive system and on the EQ VAS were also analyzed. To describe the participants’ responses to the preference elicitation tasks, we used the misery index score (MIS), defined as the sum of the attribute levels (e.g., for the health state ‘12321’ MIS = 1 + 2 + 3 + 2 + 1 = 9). This is a very crude measure, as there is no discrimination between the attributes, but it allowed us to compare health states by the number of deviations from full health. The responses to the cTTO tasks were analyzed: (i) by eliciting the value’s distribution for the overall responses, (ii) for the ‘pit’ state (‘33333’, MIS = 15), and (iii) by reporting the mean values observed per MIS. The DCE responses were described in terms of preference proportion by pair of health states and by MIS.
Data modeling

Regression models were estimated to determine values for all 243 health states described by EQ-5D-3L. The cTTO and DCE data were modeled separately first, and later combined into a hybrid regression model.

Multiple regression models for the cTTO data were estimated and compared in terms of prediction accuracy using the mean absolute error (MAE). The dependent variable was defined as disutility (i.e., 1 minus the cTTO utility value observed for a health state). Thus, coefficients expressed utility decrements of moving from level 1 (no problem) to upper levels. Disutility was explained by 10 independent variables associated with the coefficients representing the utility decrement of moving from level 1 to intermediate severity level (2) and extreme severity level (3) in each of the five EQ-5D dimensions. Dummy variables were named referring to the dimension and to the severity level associated with each (e.g., if the health state included extreme problems in usual activities UA2 would be equal to zero and UA3 to 1).

As the responses in the cTTO and DCE tasks were clustered within respondents, the cTTO data were analyzed using a random-intercept generalized least squares (GLS) model (Model 1). The mean and standard deviations of the observed cTTO values varied between the most severe health states and milder health states, which may have affected the data modeling. Hence, a homoscedasticity test was performed using the Breusch–Pagan Lagrange multiplier test [27]. As homoscedasticity was rejected, a model that corrected for multiplicative heteroscedasticity was estimated (Model 3) [13, 28]. In addition, a Tobit model censoring at...
− 1 (Model 2) was performed, since the cTTO task censors values below − 1 by limiting the possible time to trade to 10 years in the lead-time part of the task [29]. Finally, a Tobit model censoring at -1 and accounting for heteroscedasticity was explored (Model 4). All model equations are reported in Appendix 1.

For the DCE observations, conditional logistic regression (Model A) was performed using the same cTTO model’s dummy parameters, and the dependent variable was the stated choice for each health state pair (i.e., 0 or 1 for the health state A of each pair, as the EQ-PVT randomly assigns the order of appearance of the pairs and their configuration). The model generated values on a latent arbitrary scale that required to be rescaled to produce QALYS, and was thus anchored on the utility range (0 death, 1 full health). We assumed that the DCE model coefficients were proportional to those of the cTTO model [17, 30]. Hence, a proportional rescaling parameter θ (theta) was introduced in order to allow comparison between the dichotomous and the continuous models [29].

Finally, hybrid models were estimated, using the same assumptions as in the cTTO models. Thus, a standard hybrid model (Model I), a hybrid Tobit model censoring at -1 (Model II), a hybrid model correcting for heteroscedasticity (model III), and a hybrid Tobit model correcting for multiplicative heteroscedasticity (Model IV) were estimated. The models were compared in terms of a set of criteria: logical consistency of the parameter estimates, significance of the parameters, and goodness of fit of the model measured by the Akaike information criterion (AIC).

Although each model estimated took the form of equation 1, details of the various models estimated are reported in the Appendix.

\[ Y = \beta_0 + \beta_1 \times MO_2 + \beta_2 \times MO_3 + \beta_3 \times SC_2 + \beta_4 \times SC_3 + \beta_5 \times UA_2 + \beta_6 \times UA_3 + \beta_7 \times PD_2 + \beta_8 \times PD_3 + \beta_9 \times AD_2 + \beta_{10} \times AD_3. \] (1)

Statistical analyses were performed using STATA/MP 13.

**Results**

**Data collection and quality check procedure**

Data collection took place between June and September 2019. A total of 327 participants were interviewed nationwide. Interviewers were free not to include the data they collected if they felt that the participant clearly did not understand the task. One of the five interviewers was excluded for showing non-compliance with the interviewer’s protocol, leading to data quality issues. This did not improve over two consecutive data collection rounds. Subsequently, the data collected by this interviewer were excluded. Data monitoring indicated satisfactory data quality for the other interviewers. Six additional interviews were excluded for non-completion of the DCE or the cTTO tasks, resulting in 300 interviews to be included for data analysis.

**Sample characteristics**

The study sample was generally representative of the Tunisian population in terms of age, gender and region of habitat, with a slight over-representation of the northeast region of the country and of male youths aged between 20 and 39 years old, as reported in Table 1.

No problems in any dimension of EQ-5D-3L (i.e., state ‘11111’) was self-reported by 83 participants (27.7%), 43 of whom (52.4%) gave a score of at least 90 on the EQ VAS, with a mean score equal to 83.91 (SD = 13.4) and a median equal to 90. Overall, the mean observed EQ VAS score was equal to 72.7 and only 6% of the respondents reported a score lower than 50.

The distribution of the responses in the cTTO task is reported in Fig. 2, where 790 responses (26.3%) were negative and 369 (12.3%) of the observations equaled -1. In addition, cTTO values were clustered at the value 1, with 395 observations (13.2%). For 16 health states, observations ranged from -1 to 1, and from 0 to 1 for 5 other states. The mean cTTO value was negative for 8 health states with a lowest value for the state ‘33333’ of − 0.72.

For the DCE data, the observations show that participants chose the state with a lower MIS in 73.81% of their responses. For pairs of health states with an equal MIS, life A was chosen in 50.88% of the answers. The pair state number 21 showed a maximum in similar responses with 99.2% of the respondents preferring life B.

**Modeling**

All models derived logically consistent and significant parameter estimates (p < 0.05). For all cTTO models, the constant term was nearly zero and non-significant and was therefore suppressed. Model 1 presented the lowest AIC value; meanwhile Model 3 had the lowest
MAE (see Table 2). Although Model 1 was the best-fitting model using AIC, it was not the most precise in terms of MAE.

Coefficients estimated by the DCE model (Model A) were rescaled using the theta parameter. Model A showed the same ranking with respect to the dimensions, with a larger weight for the mobility dimension compared to the models for the cTTO data.

All the hybrid models (Table 3) showed both logical consistency and significance of the parameter estimates ($p < 0.001$ for all coefficients). Re-estimating the models with the constant term did not show any significant change in the parameter estimates as presented in the supplemental materials. Model III had the lowest AIC of the four hybrid models, equaling 6511.941. The difference with the hybrid Model I having the second lowest AIC value ($\Delta I = AIC_I - AIC_{III} = 396.511$), and calculating the likelihood of model III:

$$\text{Exp}(-1/2 \Delta I) = 7.92 \times 10^{-87}$$

showed no evidence to support Model I and that there was no probability for the latter model to minimize the data information loss [31]. Furthermore, the same model showed the lowest MAE calculated on the basis of the cTTO observations. Hence, the hybrid model corrected for heteroscedasticity (Model III) was chosen to construct the Tunisian value set (supplemental material) using the following utility equation (2):

$$U = 1 - M_0 \times 0.076 - M_3 \times 0.597 - SC_2 \times 0.165 - SC_3$$

$$\times 0.34 - UA_2 \times 0.078 - UA_3 \times 0.251 - PD_2 \times 0.057 - PD_3$$

$$\times 0.276 - AD_2 \times 0.095 - AD_3 \times 0.332.$$ (2)

These coefficients express the utility decrement associated with each health problem, thus allowing the calculation of utilities for the 243 health states described in EQ-5D-3L. For example, the utility value associated with the health state ‘11223’ is equal to

$$\text{Utility}(11223) = 1 - (0 \times 0.076) - (0 \times 0.597) - (0 \times 0.165)$$

$$- (0 \times 0.34) - (1 \times 0.078) - (0 \times 0.251) - (1 \times 0.057)$$

$$- (0 \times 0.276) - (0 \times 0.095) - (1 \times 0.332) = 0.533.$$ (3)
Discussion

In this study, a Tunisian value set for EQ-5D-3L was developed using a hybrid approach in accordance with EuroQol Group recommendations, thus (i) facilitating its utilization in international cross-country studies, and (ii) allowing comparisons between Tunisian valuations and those of other countries.

The hybrid approach assumes that a single utility function underlies a participant’s responses, and estimating optimal parameters for the combined data would require the creation of a single likelihood function by combining the likelihood functions of the continuous and dichotomous responses.

Table 2 Parameter estimates and fit statistics of the cTTO models

|               | GLS (Model 1) | Tobit GLS (Model 2) | HET (Model 3) | Tobit HET (Model 4) |
|---------------|---------------|---------------------|---------------|---------------------|
|               | β (SE)        | β (SE)              | β (SE)        | β (SE)              |
| MO2           | 0.086 (0.020) | 0.074 (0.022)       | 0.055 (0.016) | 0.088 (0.019)       |
| MO3           | 0.546 (0.020) | 0.592 (0.022)       | 0.565 (0.023) | 0.548 (0.020)       |
| SC2           | 0.161 (0.018) | 0.168 (0.020)       | 0.142 (0.017) | 0.164 (0.018)       |
| SC3           | 0.398 (0.019) | 0.439 (0.021)       | 0.396 (0.021) | 0.397 (0.019)       |
| UA2           | 0.103 (0.020) | 0.094 (0.022)       | 0.070 (0.016) | 0.103 (0.019)       |
| UA3           | 0.246 (0.019) | 0.274 (0.021)       | 0.262 (0.022) | 0.247 (0.019)       |
| PD2           | 0.054 (0.018) | 0.047 (0.020)       | 0.052 (0.016) | 0.054 (0.018)       |
| PD3           | 0.299 (0.019) | 0.322 (0.021)       | 0.308 (0.022) | 0.300 (0.019)       |
| AD2           | 0.094 (0.020) | 0.088 (0.022)       | 0.086 (0.016) | 0.092 (0.020)       |
| AD3           | 0.280 (0.019) | 0.300 (0.021)       | 0.283 (0.022) | 0.277 (0.019)       |
| Constant      | 0.000 (0.024)* | −0.009 (0.027)*     | 0.018 (0.015)* | −0.007 (0.024)*     |
| Uncensored Observations | 3000               | 2631               | 3000               | 2631               |
| Right-Censored Observations | 0                   | 369                | 0                   | 369                |
| AIC           | 3320.868       | 4193.709            | 3555.007         | 3308.948            |
| BIC           | 3398.951       | 4271.792            | 3687.147         | 3447.095            |
| MAE           | 0.341          | 0.342               | 0.335             | 0.342               |
| Predicted value for the pit state | −0.769             | −0.927             | −0.814            | −0.765             |

cTTO: composite time trade-off, β: coefficient, SE: standard error. Dimensions: MO: mobility, SC: self-care, UA: usual activities, PD: pain/discomfort, AD: anxiety/depression, GLS: general least squares, HET: model correcting for heteroscedasticity, AIC: Akaike Information Criterion, BIC: Bayesian Information Criterion, MAE: Mean absolute error.

*P value > 0.05

Fig. 2 Distribution of the observed cTTO values in percentages
(i.e., cTTO and DCE data) [20, 29]. Hence, we assumed the best-fitting continuous model could produce the best-fitting hybrid model, so we compared the cTTO models first, then the hybrid models. The assumption was refuted as model III was the best-fitting hybrid model (while Model 1 was the best-fitting cTTO model). The kernel density plots of the prediction value for the 243 health states, using the different single models (3 and A), showed high convergence between the cTTO and the DCE predictions, as shown in Fig. 3. This reflects the complementarity of using both elicitation techniques in order to produce estimates that were more consistent and to reach a closer utility function to the real one. The utility range varied from [-0.814 to 1] for Model 3, [-0.821 to 1] for Model A and [-0.796 to 1] for Model III (see Table 4). Combining the data produced a narrower

| Parameter estimates and fit statistics of the hybrid models | HYBRID (Model I) | HYBRID TOBIT (Model II) | HYBRID HET (Model III) | HYBRID TOBIT HET (Model IV) |
|---------------|-----------------|-------------------------|------------------------|-----------------------------|
| β (SE)        | β (SE)          | β (SE)                  | β (SE)                 | β (SE)                      |
| MO2           | 0.082(0.014)    | 0.077(0.016)            | 0.076(0.012)           | 0.071(0.011)                |
| MO3           | 0.587(0.015)    | 0.634(0.017)            | 0.597(0.016)           | 0.658(0.020)                |
| SC2           | 0.177(0.013)    | 0.181(0.015)            | 0.165(0.012)           | 0.164(0.013)                |
| SC3           | 0.338(0.014)    | 0.364(0.016)            | 0.340(0.015)           | 0.370(0.017)                |
| UA2           | 0.082(0.015)    | 0.081(0.016)            | 0.078(0.012)           | 0.076(0.011)                |
| UA3           | 0.244(0.014)    | 0.266(0.016)            | 0.251(0.014)           | 0.273(0.016)                |
| PD2           | 0.052(0.014)    | 0.045(0.015)            | 0.057(0.012)           | 0.055(0.012)                |
| PD3           | 0.270(0.014)    | 0.288(0.016)            | 0.276(0.014)           | 0.296(0.016)                |
| AD2           | 0.095(0.015)    | 0.089(0.016)            | 0.095(0.012)           | 0.093(0.012)                |
| AD3           | 0.329(0.014)    | 0.350(0.015)            | 0.332(0.014)           | 0.357(0.016)                |
| AIC           | 6908.452        | 7788.361                | 6511.941               | 6937.185                    |
| BIC           | 6988.650        | 7868.560                | 6658.972               | 7084.215                    |
| MAE (cTTO observations as a benchmark) | 0.342            | 0.343                   | 0.341                  | 0.344                       |
| Prediction for the pit state | -0.768          | -0.902                  | -0.796                 | -0.954                      |
| Continuous uncensored observations | 3000            | 2631                    | 3000                   | 2631                        |
| Continuous right-censored observations | 0               | 369                     | 0                      | 369                         |
| Dichotomous observations | 2903            | 2903                    | 2903                   | 2903                        |

$cTTO$ composite time trade-off, $β$ coefficient, $SE$ standard error, Dimensions: $MO$ mobility, $SC$ self-care, $UA$ usual activities, $PD$ pain/discomfort, $AD$ anxiety/depression, $GLS$ general least squares, $HET$ model correcting for heteroscedasticity, $AIC$ Akaike information criterion, $BIC$ Bayesian information criterion, $MAE$ mean absolute error

![Fig. 3](Image) The kernel density plots of the prediction value for the 243 health states using the single models and the hybrid model. Model III: Hybrid model corrected for heteroscedasticity; Model A: Conditional logistic model. Model 3: Generalized Least Square model corrected for heteroscedasticity.
In all models, MO3 had the highest weight (0.597), followed by SC3 (0.340), suggesting that mobility is by far the most important health dimension based on Tunisian preferences. This reflects how the public perceive physical disability and could be explained by the social disparities facing disabled people in Tunisia (i.e., discrimination in employment, lower access to education, and reduced quality of life) [32, 33]. The predicted value for the pit state (−0.796) was lower than the ones observed in the value sets for high-income countries such as the USA or France (−0.573 and −0.525), but was similar to other low- to middle-income countries such as Indonesia or Ethiopia (−0.865 and −0.718), possibly reflecting divergences in health perceptions between countries with different income levels [34–37].

Modeling in our study did not account for religious beliefs, as suggested by some authors who considered participants’ answers in Christian and Muslim communities to be biased by their beliefs [38, 39]. Although 43 participants out of the 300 stated that their religious beliefs affected their responses, only one non-trader was reported during the valuation study. Furthermore, the Tunisian value set had lower utility values when compared with Iran’s value set which was presumed to be influenced by religion [40]. It may be necessary to explore whether religion should be accounted for in culturally religious communities during valuation studies.

Applying the existing crosswalk methodology developed by Van Hout et al. to the Tunisian EQ-5D-3L value set allowed values to be estimated for the 3125 EQ-5D-5L health states and these are presented in the supplemental materials [41]. We observed similarities with recent international crosswalk studies, notably those of Sri Lanka and Poland [42, 43]. The Tunisian crosswalk value set has proportionally fewer health state utilities lower than zero or higher than 0.8 when compared with EQ-5D-3L tariff values. It has been suggested that this effect was due to restricting the range of index values in order to obtain an equivalent severity scope [43].

The main limitation of our study was that the sample may not have been fully representative of the population, as illiterate participants were not included in the study. Illiterate people represent 18.8% of the general population in Tunisia (22.34% for those over 20 years), but since the ability to read is required to complete the valuation tasks, it was impossible to include them in the sample [20]. Another limitation is that, although our recruitment strategy covered all the nation’s regions, the quota sampling did not account properly for rural areas. Finally, due to technical issues with EQ-PVT, 97 DCE observations were corrupted and the data concerned were lost, which corresponded to a loss of DCE data of roughly 10 interviews.

Table 4 Parameter estimates of the single models and hybrid model used to construct the value set

| Health Dimension | MO2 | MO3 | SC2 | SC3 | UA2 | UA3 | PD2 | PD3 | AD2 | AD3 | Constant |
|------------------|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|----------|
| β (SE)           | 0.055 (0.016) | 0.565 (0.023) | 0.142 (0.017) | 0.396 (0.021) | 0.070 (0.016) | 0.262 (0.022) | 0.052 (0.016) | 0.308 (0.022) | 0.086 (0.016) | 0.283 (0.023) | 0.018 (0.015)* |
| β (SE)           | 0.397 (0.078) | 2.279 (0.103) | 0.676 (0.074) | 1.151 (0.091) | 0.341 (0.082) | 0.863 (0.083) | 0.220 (0.072) | 0.942 (0.085) | 0.386 (0.086) | 1.364 (0.094) | 0.018 (0.015)* |
| Rescaled β      | 0.109 | 0.629 | 0.186 | 0.318 | 0.094 | 0.238 | 0.060 | 0.260 | 0.106 | 0.376 | 0.0000 |
| β (SE)           | 0.076 (0.012) | 0.597 (0.016) | 0.165 (0.012) | 0.340 (0.015) | 0.078 (0.012) | 0.251 (0.014) | 0.057 (0.012) | 0.276 (0.014) | 0.095 (0.012) | 0.332 (0.014) | 0.796 |

*P value > 0.05

cTTO composite Time Trade-off, DCE discrete choice experiments, β coefficient, SE standard error, Dimensions: MO mobility, SC self-care, UA usual activities, PD pain/discomfort, AD anxiety/depression, GLS general least squares, HET model correcting for heteroscedasticity.
Conclusion

This is the first Tunisian preference-based value set to be published. These EQ-5D-3L health state utilities reveal the preferences of a sample of the Tunisian general population with respect to different impaired health states. The effect of having impaired mobility on HRQoL was the largest of all 5 dimensions. The Tunisian EQ-5D-3L values differed from those derived in other countries. This EQ-5D-3L value set should be considered for utilization in HTA in order to assist health policy decision-making in Tunisia.

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Author contributions The study was conceived by MRK, AK, CD and JC following the EuroQol protocol. CD and HF coordinated the data acquisition. JC contributed in the data collection. BR and FAS performed the data quality control. JC and BR performed the data analyses which were reviewed by MK. JC wrote the first draft of the manuscript. All authors provided their intellectual inputs, reviewed and recommended revisions to the final submitted manuscript.

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Data availability The datasets generated during and/or analyzed during the current study are available from the corresponding author on reasonable request.

Compliance with ethical standards

Conflict of interest All authors declare that they have no conflict of interest.

Ethical approval This study was performed in line with the principles of the Declaration of Helsinki and was approved after examination by the medical research department of the Tunisian ministry of health as recommended by the ethics committee of the “Habib Thameur” hospital.

Consent to participate Informed consent was obtained from all individual participants included in the study.

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