



Value Set for the EQ-5D-Y-3L in Hungary

Fanni Rencz¹ · Gábor Ruzsa^{2,3} · Alex Bató^{1,4} · Zhihao Yang⁵ · Aureliano Paolo Finch⁶ · Valentin Brodzsky¹

Accepted: 4 September 2022 / Published online: 20 September 2022
© The Author(s) 2022

Abstract

Background The Hungarian health technology assessment guidelines recommend the use of the EuroQol instrument family in quality-adjusted life-year calculations. However, no national value set exists for the EQ-5D-Y-3L or any other youth-specific instrument.

Objective This study aims to develop a national value set of the EQ-5D-Y-3L for Hungary based on preferences of the general adult population.

Methods This study followed the international valuation protocol for the EQ-5D-Y-3L. Two independent samples, representative of the Hungarian general adult population in terms of age and sex were recruited to complete online discrete choice experiment (DCE) tasks and composite time trade-off (cTTO) tasks by computer-assisted personal interviews. Adults valued hypothetical EQ-5D-Y-3L health states considering the health of a 10-year-old child. DCE data were modelled using a mixed logit model with random-correlated coefficients. Latent DCE utility estimates were mapped onto mean observed cTTO utilities using ordinary least squares regression.

Results Overall, 996 and 200 respondents completed the DCE and cTTO surveys, respectively. For each domain, the value set resulted in larger utility decrements with more severe response levels. The relative importance of domains by level 3 coefficients was as follows: having pain or discomfort > feeling worried, sad or unhappy > mobility > doing usual activities > looking after myself. Overall, 12.3% of all health states had negative utilities in the value set, with the worst health state having the lowest predicted utility of -0.485 .

Conclusion This study developed a national value set of the EQ-5D-Y-3L for Hungary. The value set enables to evaluate the cost utility of health technologies for children and adolescents based on societal preferences in Hungary.

1 Introduction

Health technology assessment (HTA) plays an important role in enhancing health policy decision making about the coverage and use of health interventions [1]. HTA provides a systematic evaluation of new health technologies

✉ Fanni Rencz
fanni.rencz@uni-corvinus.hu

¹ Department of Health Policy, Corvinus University of Budapest, 8 Fővám tér, 1093 Budapest, Hungary

² Department of Statistics, Corvinus University of Budapest, Budapest, Hungary

³ Doctoral School of Psychology, Institute of Psychology, Eötvös Loránd University, Budapest, Hungary

⁴ Károly Rác Doctoral School of Clinical Medicine, Semmelweis University, Budapest, Hungary

⁵ Health Services Management Department, Guizhou Medical University, Guiyang, China

⁶ EuroQol Office, EuroQol Research Foundation, Rotterdam, The Netherlands

Key Points for Decision Makers

This study developed a national value set of the EQ-5D-Y-3L for Hungary, derived from the Hungarian adult general population.

The value set allows to estimate utilities from EQ-5D-Y-3L responses in children and adolescents and can be used for cost-utility analyses of health technologies.

Hungary is the first country in Central and Eastern Europe with value sets for the EQ-5D-3L, EQ-5D-5L and EQ-5D-Y-3L.

(e.g. treatment, diagnostic procedure) based on both clinical and economic evidence. In most countries, cost-utility analysis, resulting in cost/quality-adjusted life-year (QALY) estimates is a preferred form of economic evaluations in national HTA guidelines [2]. QALY is a summary measure of health gain that takes into account

health-related quality of life and the length of survival. To compute QALYs, health-related quality of life is mostly measured using generic preference-accompanied measures [3]. These instruments comprise domains that describe health states (i.e. descriptive system) and a set of preference weights (i.e. value set) that allows to assign utilities to health states. In Hungary, HTA guidelines developed by the Ministry of Human Capacities recommend the use of the EuroQol instrument family to obtain utilities [4–6]. Hungarian value sets are available for the two adult versions of the EQ-5D, the EQ-5D-3L and EQ-5D-5L, both having been developed according to the latest EuroQol Valuation Technology (EQ-VT) protocol [7].

The EQ-5D-3L and EQ-5D-5L, which have been developed for use in the adult population, are typically not suitable to measure health outcomes in young populations, such as adolescents and children. Adult and youth instruments may differ in a number of ways, including health domains and item content, age-appropriate language and the value sets used to assign utilities to health states [8, 9]. A growing number of generic preference-accompanied measures have been developed for young populations. Examples include the EQ-5D-Y-3L, EQ-5D-Y-5L (experimental version), Child Health Utility 9D (CHU9D), Assessment of Quality of Life-6D (AQoL-6D) Adolescent, Health Utilities Index (HUI) 2 and 3, 16D and 17D [10, 11]. Among these, the largest amount of positive psychometric evidence has been reported about the EQ-5D-Y-3L [12]. The EQ-5D-Y-3L is a three-level youth measure targeting populations aged 8–15 years [13, 14]. However, it requires a separate value set due to the differences in the wording of descriptive system and utilities elicited for children/adolescents in comparison with adults [9, 15–22].

Recently, an international protocol has been established for the valuation of EQ-5D-Y-3L, which advocates to rely upon adult general population preferences for hypothetical health states [23]. Arguments supporting this approach include coherence with the population used to generate value sets for adult measures; adults bear the costs of healthcare through health insurance; adults' capacity of decision making (both in general and about health) and legal rights to participate in certain activities (e.g. voting, signing a contract, getting married); ethical difficulties around the acceptability of valuation exercises involving the concept of death for young populations; and matters of understanding and cognitive burden associated with the task in adolescents or children [24–27]. In Hungary, no value set exists for the EQ-5D-Y-3L or any other youth-specific health status measure. This study therefore aimed to develop a national value set for the EQ-5D-Y-3L in Hungary based on preferences of the general adult population.

2 Methods

This study followed the international valuation protocol for the EQ-5D-Y-3L [23] and the Checklist for Reporting Valuation Studies of the EQ-5D (CREATE) [28]. Ethical approval was obtained from the research ethics committee of the local research team's institution (no. KRH/31/2021).

2.1 EQ-5D-Y-3L

The official Hungarian version of the EQ-5D-Y-3L was used in this study. The EQ-5D-Y-3L consists of a descriptive system that measures health across five domains and a visual analogue scale (EQ VAS) with endpoints of 0 (the worst health you can imagine) and 100 (the best health you can imagine) [13]. The five domains include mobility (walking about), looking after myself (washing or dressing), doing usual activities (going to school, hobbies, sports, playing, doing things with family or friends), having pain or discomfort, and feeling worried, sad or unhappy. In the EQ-5D-Y-3L, each domain has three response levels (level 1 = no problems/no pain or discomfort/not worried, sad or unhappy; level 2 = some problems/some pain or discomfort/a bit worried, sad or unhappy; and level 3 = a lot of problems/a lot of pain or discomfort/very worried, sad or unhappy) defining a total of 243 different health state profiles. The numbers assigned to the level of each domain can be summarised as a five-digit string, whereby 11111 (full health) indicates no problems in any domains and 33333 (worst health state) refers to the worst level of problems in all domains.

2.2 Study Design

The study design used a two-step valuation approach involving data collections through two independent surveys with different modes of administration, samples, and preference elicitation methods (online survey with discrete choice experiment [DCE] tasks and face-to-face computer-assisted personal interviews with composite time trade-off [cTTO] tasks). Both surveys comprised of the following elements: introduction page, informed consent page, introductory questions, self-reported health on the EQ-5D-Y-3L to familiarise respondents with the measure, preference elicitation tasks, three debriefing questions and background questions (e.g. employment, marital and parental status, and the presence of chronic health conditions). The face-to-face survey included three introductory questions about the respondents' age, sex and experience with severe illness, while questions about education, place of living and geographical region appeared among the background questions. By contrast, the online survey contained five introductory questions to allocate respondents to a quota group based on age, sex, education, place of living and geographical region, and an extra question about experience with severe illness.

2.3 Preference Elicitation Methods

The valuation tasks were conducted using the EQ-VT software (v2.1) [29]. Adults completed preference elicitation tasks considering the health of a 10-year-old child (exact phrasing: ‘Considering your views for a 10-year-old child’) [23]. The DCE tasks asked the participants to express their preference between two different EQ-5D-Y-3L health states (labelled as option ‘A’ and option ‘B’). The cTTO task was composed of a conventional 10-year TTO for better-than-dead states and a lead-time variant (i.e. 10 years in full health followed by 10 years in an EQ-5D-Y-3L state) for worse-than-dead states [30]. The smallest tradable unit was 6 months. Examples of the DCE and cTTO tasks are presented in Online Resource 1. At the respondent’s point of indifference (t), the utilities for the EQ-5D-Y-3L health states were calculated as follows:

$$U = t/10 \text{ for better-than-dead states}$$

$$U = (t - 10)/10 \text{ for worse-than-dead states}$$

The possible range of observed utilities was -1 to 1 .

2.4 Health State Selection

The health states valued in this study were those described in the international valuation protocol of the EQ-5D-Y-3L [23]. For the DCE design, 150 health state pairs were selected using a Bayesian efficient design allowing for the estimation of main effects and all two-way interactions. The design attempts to minimise the number of implausible health states, provides an appropriate balance across both severity levels and utilities, and uses an overlap in two domains for all pairs [18]. The 150 pairs were distributed over 10 blocks of 15 pairs. Respondents were randomly allocated to one of the 10 blocks and the order of the pairs within each block was also randomised. In addition, each respondent completed three fixed dominant pairs, whereby one health state was always unanimously better than the other (e.g. 21123 vs. 22233). The first and last pair were set at dominant pairs for all respondents, while the third dominant pair was presented at a random point.

For the cTTO design, one block of 10 health states was used, containing three mild states (11112, 11121 and 21111), two moderate states (22223 and 22232), four severe states (31133, 32223, 33233 and 33323) and the worst health state (33333). The order of the health states was randomised across respondents. According to the EQ-VT protocol, each respondent valued two wheelchair examples, three practice health states (21112, 32323, 13311) and 10 ‘real’ EQ-5D-Y-3L states [29, 31]. After the valuation, a ranking of the 10 health states (‘feedback

module’) based on their responses was displayed to the respondents who had the opportunity to remove one or more cTTO valuations (regardless of being inconsistent or not) [32].

2.5 Sampling, Recruitment, and Data Collection

The target sample size was approximately 1000 respondents for the DCE and 200 for the cTTO survey. Members of the general population aged 18 years or above who provided informed consent were included in the study. The language of both surveys was Hungarian. For the DCE survey, participants were recruited from members of a large Hungarian online panel. Respondents for the cTTO interviews were recruited by the interviewers using their existing contacts and volunteers. For both surveys, a quota sample was drawn using interlocking quotas for age and sex to achieve population representativeness according to census data reported by the Hungarian Central Statistical Office [33]. Furthermore, we used separate ‘soft’ quotas for education, place of living and geographical region. We attempted to fill in these quota brackets approaching the census data as much as possible, but we did not close the quotas when these were completed. Online panel members received survey points redeemable for gift vouchers or prize-drawing tickets as compensation, while no remuneration was provided for participation in the cTTO interviews. The online DCE survey took place in April and May, and the cTTO interviews were performed between March and September 2021. A team comprising a PhD student (A.B.) and three MSc students was used, with each person conducting 50 interviews. All but one interviewer had extensive experience with valuation studies as they had previously worked as interviewers in the adult EQ-5D-3L and EQ-5D-5L parallel valuation study in Hungary [7]. Interviewers received a standardised training on valuation methods, the EQ-VT protocol and the quality control (QC) procedure. Each interviewer conducted 10 pilot interviews before the formal data collection commenced.

2.6 Quality Control

Both the DCE and cTTO surveys included a QC process developed for valuation studies of EuroQol instruments. In the DCE survey, the following QC rules were established [34, 35]:

- (1) Speeding: respondents were required to spend at least 150 s on completing the 18 (15 ‘real’ and 3 ‘dominant’) DCE pairs.
- (2) Dominance test: respondents were expected to answer correctly to all three dominant pairs.

Respondents failing any of the above tests were blocked from proceeding to the background questions and were excluded from the data analysis based on quality grounds. In the cTTO interviews, the standard QC procedure was followed to assess interviewers' performance in terms of protocol compliance and interviewer effects [32]. Interviews were assessed against the following four QC criteria: (1) no explanation of the worse-than-dead format in the wheelchair task; (2) the interviewer spent less than 3 min on explaining the wheelchair tasks; (3) the 10 'real' health states were completed within less than 5 min; and (4) the worst health state was valued at least 5 years better than the health state valued as being the worst ('33333 inconsistency'). Data quality was primarily monitored by one of the authors (F.R.) and discussed with the EQ-VT support team at 25, 50 and 75% of the target sample size. The QC criteria for the cTTO tasks did not serve as exclusion criteria.

2.7 Data Analysis and Modelling

Respondents' background characteristics were summarised using descriptive statistics. The value set was developed using both DCE and cTTO data. Following the international valuation protocol [23], statistical modelling was conducted in two steps. First, the DCE valuation data were used to estimate the relative importance of domains and levels on a latent scale. The dependent variable was coded 1 for the chosen option (e.g. 'A') in each choice task and 0 for the other alternative (e.g. 'B'). The models included main effect dummy-coded variables for level 2 and level 3 problems in each domain, resulting in a total of 10 parameters. DCE responses were analysed in a probabilistic choice/random utility model framework ('mlogit' package in R), using both 'conditional logit' (fixed coefficients) and 'mixed logit' (correlated random coefficients) variants [36]. Mixed logit is preferred over the conditional logit model as it allows for unobserved heterogeneity in individuals' preferences over choice option characteristics [37]. Maximum likelihood estimation was implemented using the Broyden–Fletcher–Goldfarb–Shanno (BFGS) algorithm. Concerning the mixed logit model, unconditional likelihood values were approximated via 5000 pseudo-random draws from the (supposedly normal) distribution of correlated random coefficients. Multiple criteria were used in selecting between DCE models. These included theoretical considerations (e.g. accounting for preference heterogeneity), goodness of fit as measured by the log-likelihood value and the Bayesian information criterion (BIC), and prediction accuracy in terms of the mean absolute error (MAE) and mean square error (MSE) of the observed versus predicted choice probabilities for the 150 pairs. For the latter two, in both the conditional and mixed logit models, we used the following formulas:

$$\text{MAE} = \sum_s |m(p_s(A)) - Pr_s(A)|/S \text{ and } \text{MSE} = \sum_s (m(p_s(A)) - Pr_s(A))^2/S,$$

whereby $S = 150$ is the number of health state pairs involved in the DCE task, s is the index for identifying health state pairs, $Pr_s(A)$ is the proportion of individuals who were presented health state pair s and chose option A , and $m(p_s(A))$ is the mean estimated probability of choosing option A across individuals who were presented health state pair s , calculated according to each specific model version.

In the second step of modelling, the cTTO data were used for anchoring the best-performing DCE model onto the 0 (= dead) to 1 (= full health) scale. No respondents were excluded from the analysis of cTTO data. Respondents who gave the same '1' value to all health states (i.e. non-traders) were not excluded as we considered these valuations as the reflections of respondents' true preferences. However, following previous valuation studies, responses flagged by participants on the feedback module were excluded from the data analysis [38]. Two different anchoring methods were tested to link DCE and cTTO data: worst health state anchoring and mapping [39]. For worst health state anchoring, we rescaled the regression coefficients for each level of each dimension in the DCE model by the ratio of the mean observed cTTO disutility of the worst health state (33333) to the latent scale DCE value thereof. For the mapping, the mean observed cTTO disutilities of the 10 health states were regressed on the latent scale DCE values thereof using an ordinal least squares regression. We tested the model with and without constant and selected the preferred model based on parameter significance and model fit (R-squared). The estimated regression coefficient (β) was used to rescale each regression coefficient of the best-performing DCE model.

The two anchoring approaches were compared by computing MAE between predicted and mean observed cTTO utilities for the 10 health states and also separately for the three mild states. For all analyses, a p value < 0.05 was considered statistically significant. Statistical analyses were conducted in Stata 14 (StataCorp LLC, College Station, TX, USA) and R version 4.0.5 (R Core Team, Vienna, Austria).

3 Results

3.1 Respondent Characteristics

In total, 3273 individuals responded to the invitation to participate in the online DCE survey, 1251 (38.2%) of whom completed the survey. Of non-completers, 159 (7.9%) declined consent, 1518 (75.1%) belonged to a quota that was full, and 345 (17.1%) dropped out or lost the internet connection. Of the 1251 completers, 255 (20.4%) did not pass the QC criteria and thus were excluded. The majority of these respondents failed

the dominance test(s) ($n = 219$, 85.9%) [Online Resource 2]. Overall, 221 people were invited for the cTTO survey, 16 (7.2%) of whom were unwilling to participate, 3 (1.4%) were not interviewed due to conflicting schedules, and 2 (0.9%) refused to engage in face-to-face interactions due to the coronavirus disease 2019 (COVID-19) pandemic. Thus, 996 and 200 respondents from the DCE and cTTO surveys, respectively, formed the final analytical samples for this study.

The sociodemographic and health-related characteristics of the samples are reported in Table 1. The DCE and cTTO samples were similar in most characteristics. The overall study population was closely representative of the general population based on age, sex, geographical region, place of living, employment, and marital status; however, we had an overrepresentation of college/university-educated respondents.

3.2 Descriptive Statistics of the Composite Time Trade-Off (cTTO) Utilities

Two (1.0%) of the 200 cTTO interviews were flagged on the QC criteria (one on wheelchair time and another on 33333 inconsistency). There were two (1.0%) non-traders. A total of 40 (20.0%) individuals had at least one inconsistent cTTO response (18.0% related to moderate or severe health states, 1.5% involving 33333 and 0.5% related to the mild states). Fifty-eight respondents (29.0%) flagged overall 68 (3.4%) cTTO responses on the feedback module that reduced the rate of inconsistency to 8.0% (all related to the moderate or severe states). The distribution of cTTO responses (after the feedback module) is plotted in Fig. 1. Overall, 13.0% of the 1932 unflagged cTTO responses were at 1, 28.0% were worse than dead, and 2.7% were at -1 . The mean observed cTTO utilities for the 10 health states ranged between 0.948 (11112) and -0.517 (33333) [Table 2].

3.3 Modelling Discrete Choice Experiment (DCE) Data

Both the conditional and mixed logit models resulted in larger utility decrements with more severe response levels in each domain (Table 3). In both models, the relative importance of domains by level 3 coefficients was having pain or discomfort > feeling worried, sad or unhappy > mobility > doing usual activities > looking after myself. The mixed logit model demonstrated considerably better goodness-of-fit statistics and prediction accuracy, including higher (i.e. less negative) log-likelihood (-6480 vs. -5602) and lower BIC (13056 vs. 11830), MAE (0.0474 vs. 0.0348) and MSE (0.0040 vs. 0.0022). The mixed logit was selected as the final model based on its better fit indices, prediction accuracy and ability to account for preference heterogeneity.

3.4 Anchoring Onto cTTO Utilities

The ordinary least squares regression without constant was selected as the preferred mapping model to rescale the DCE model with model parameters and fit statistics as follows: $\beta = -0.09509$ (SE 0.00201, $p < 0.001$), R -squared 0.996, model F test (1,9) 2230.36 ($p < 0.001$). The two anchoring approaches (worst health state anchoring and mapping) produced very similar models in terms of parameter estimates and prediction accuracy (Table 4). The predicted utility for the pits state was slightly lower (-0.517 vs. -0.485) and the proportion of health states with negative utilities was somewhat higher with worst health state anchoring than with mapping (13.2 vs. 12.3%). The agreement between the predicted and observed utilities is displayed in Fig. 2. The mapping approach, resulting in marginally better agreement with mean observed cTTO utilities as indicated by the lower MAE values both for all 10 health states (0.0486 vs. 0.0485) and the three mild states separately (0.0311 vs. 0.0298) was selected as the final value set. With the selected value set, utilities for the 243 EQ-5D-Y-3L health states ranged between -0.485 (33333) and 1 (11111), with the utility of 0.962 for the mildest impaired state (12111). The full set of the 243 predicted EQ-5D-Y-3L utilities is provided in Online Resource 3. For example, using the Hungarian value set, the predicted utility for the EQ-5D-Y-3L health state 12321 may be calculated as follows:

$$U(12321) = 1 + MO1 + LAM2 + UA3 + PD2 + WSU \\ 1 = 1 + 0 - 0.038 - 0.252 - 0.133 + 0 = 0.577$$

4 Discussion

In this study, we developed a national value set of the EQ-5D-Y-3L for Hungary, derived from the Hungarian adult general population. This is the first value set for any generic preference-accompanied measure targeting younger populations in Hungary.

The Hungarian EQ-5D-Y-3L value set was estimated using a mixed logit model with random correlated coefficients, and mapping the resultant values onto the QALY scale. The domains with the largest utility decrements were ‘having pain or discomfort’ and ‘feeling worried, sad or unhappy’. The domain of ‘looking after myself’ was associated with the smallest utility decrements, very likely because self-care may be interpreted differently in the case of a child due to the existing adult support network even for healthy children [21]. Although a direct comparison with the Hungarian adult EQ-5D-3L value set is limited due to the differences in wording of descriptive systems, preference elicitation methods (adult: cTTO vs. youth: DCE and cTTO),

Table 1 Characteristics of the sample

Variables	Reference population ^a	Total sample [n = 1196]		cTTO included [n = 200]		DCE included [n = 996]		DCE excluded [n = 255]		
		%	n	%	n	%	n	%	n	%
Sex										
Female	53.1	630	52.7	108	54.0	522	52.4	119	46.7	
Male	46.9	566	47.3	92	46.0	474	47.6	136	53.3	
Age, years										
18–24	10.0	125	10.5	22	11.0	103	10.3	29	11.4	
25–34	15.2	188	15.7	31	15.5	157	15.8	52	20.4	
35–44	19.5	234	19.6	39	19.5	195	19.6	55	21.6	
45–54	16.0	198	16.6	31	15.5	167	16.8	45	17.6	
55–64	16.8	205	17.1	33	16.5	172	17.3	36	14.1	
65–74	13.0	159	13.3	25	12.5	134	13.5	30	11.8	
75+	9.6	87	7.3	19	9.5	68	6.8	8	3.1	
Highest level of education										
Primary school or less	45.4	276	23.1	57	28.5	219	22.0	66	25.9	
Secondary school ^b	33.3	448	37.5	82	41.0	366	36.7	120	47.1	
College/university degree	21.3	472	39.5	61	30.5	411	41.3	69	27.1	
Place of residence										
Capital	17.9	318	26.6	42	21.0	276	27.7	56	22.0	
Other town	52.6	608	50.8	96	48.0	512	51.4	130	51.0	
Village	29.5	270	22.6	62	31.0	208	20.9	69	27.1	
Geographical region										
Central Hungary	30.4	427	35.7	49	24.5	378	38.0	72	28.2	
Western Hungary	30.2	357	29.8	85	42.5	272	27.3	64	25.1	
Eastern Hungary	39.5	412	34.4	66	33.0	346	34.7	119	46.7	
Employment status^c										
Employed	53.1	691	57.8	119	59.5	572	57.4	–	–	
Retired	26.1	281	23.5	53	26.5	228	22.9	–	–	
Disability pensioner	3.1	28	2.3	6	3.0	22	2.2	–	–	
Student	3.1	91	7.6	17	8.5	74	7.4	–	–	
Unemployed	4.7	61	5.1	2	1.0	59	5.9	–	–	
Homemaker/housewife	1.0	44	3.7	3	1.5	41	4.1	–	–	
Parenting status^d										
Not parent	NA	497	41.6	78	39.0	419	42.1	–	–	
Parent	NA	699	58.4	122	61.0	577	57.9	–	–	
Has child(ren) < 18 years of age	NA	245	20.5	45	22.5	200	20.0	–	–	
Marital status										
Married	45.6	484	40.5	74	37.0	410	41.2	–	–	
Domestic partnership	13.4	236	19.7	55	27.5	181	18.2	–	–	
Single	18.5	259	21.7	37	18.5	222	22.3	–	–	
Widowed	11.4	95	7.9	21	10.5	74	7.4	–	–	
Divorced	11.1	96	8.0	8	4.0	88	8.8	–	–	
Other	–	26	2.2	5	2.5	21	2.1	–	–	
Self-perceived health status										
Excellent	NA	102	8.5	35	17.5	67	6.7	–	–	
Very good	NA	321	26.8	69	34.5	252	25.3	–	–	
Good	NA	489	40.9	57	28.5	432	43.4	–	–	
Fair	NA	244	20.4	32	16.0	212	21.3	–	–	
Poor	NA	40	3.3	7	3.5	33	3.3	–	–	

Table 1 (continued)

Variables	Reference population ^a	Total sample [n = 1196]		cTTO included [n = 200]		DCE included [n = 996]		DCE excluded [n = 255]	
	%	n	%	n	%	n	%	n	%
History of chronic illness ^{e,f}									
Yes	48.0	777	65.0	111	55.5	666	66.9	-	-
No	52.0	400	33.4	88	44.0	312	31.3	-	-

cTTO composite time trade-off, DCE discrete choice experiment, NA not available

^aHungarian Central Statistical Office (KSH) Microcensus 2016

^bWith completed final examination

^cThe sum of the general population is < 100% owing to an ‘other’ category accounting for 8.9%

^dIncluding biological, stepchildren and adopted children

^eN = 19 refused to answer

^fReference values: Hungarian Central Statistical Office (KSH), Health at a Glance 2019

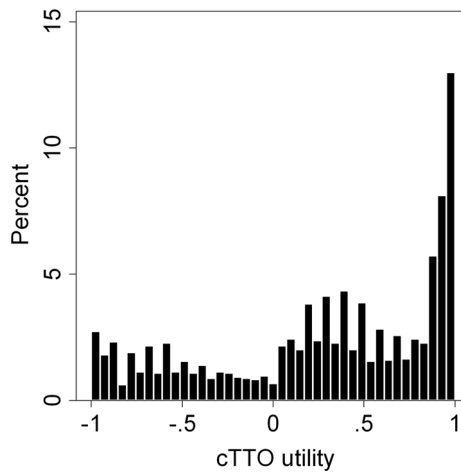


Fig. 1 Distribution of observed cTTO utilities for the 10 EQ-5D-Y-3L health states. Responses flagged by respondents on the feedback module are not included. cTTO composite time trade-off

valuation perspective (adult for self vs. adult considering the health of a 10-year-old child) and data collection period (adult: pre-COVID vs. youth: during the pandemic), it is notable that in the Hungarian adult value set, the domains of mobility and self-care have the greatest utility decrements. Our results are similar in range and domain importance order to those reported from other European countries with EQ-5D-Y-3L value sets. In Hungary, the utility of the worst health state was -0.485, compared with utilities of -0.283 in Germany [40], -0.539 in Spain [34] and -0.691 in Slovenia [41].

Following the EQ-5D-Y-3L international valuation protocol [23], we asked adults to complete the valuation tasks considering the health of a 10-year-old child. However, there may be systematic differences in preferences and value sets depending on who (e.g. adult vs. adolescent) and for whom (self vs. other, adult vs. child) values health states [42]. There is an increasing body of literature that suggests the

Table 2 Observed summary statistics for the 10 EQ-5D-Y-3L health states (cTTO)

Health state	n ^a	Mean	SD	Median	Q1	Q3	Min	Max	WTD %
11112	200	0.948	0.068	0.950	0.900	1.000	0.700	1.000	0
11121	200	0.918	0.080	0.950	0.900	1.000	0.600	1.000	0
21111	200	0.938	0.080	0.950	0.900	1.000	0.600	1.000	0
22223	189	0.449	0.316	0.500	0.350	0.650	-0.800	1.000	8
22232	190	0.361	0.311	0.400	0.250	0.550	-0.800	1.000	9
31133	186	0.013	0.449	0.125	-0.300	0.350	-0.950	1.000	44
32223	184	0.219	0.389	0.300	0.050	0.500	-0.850	1.000	22
33233	194	-0.334	0.493	-0.450	-0.750	0.150	-0.950	1.000	65
33323	191	-0.237	0.491	-0.300	-0.650	0.200	-1.000	1.000	61
33333	198	-0.517	0.504	-0.725	-1.000	0.050	-1.000	1.000	72

cTTO composite time trade-off, min minimum, max maximum, Q1 first quartile, Q3 third quartile, SD standard deviation, WTD worse than dead

^aAfter excluding responses flagged by respondents in the feedback module

Table 3 Results of DCE modelling

	Conditional logit model			Mixed logit model with random-correlated coefficients ^a		
	Estimate	SE	<i>p</i> value ^b	Estimate	SE	<i>p</i> value ^b
Parameters						
MO2	- 0.352	0.058	< 0.001	- 0.565	0.085	< 0.001
MO3	- 1.594	0.088	< 0.001	- 2.698	0.150	< 0.001
LAM2	- 0.241	0.048	< 0.001	- 0.398	0.080	< 0.001
LAM3	- 1.148	0.068	< 0.001	- 1.983	0.128	< 0.001
UA2	- 0.548	0.043	< 0.001	- 0.825	0.080	< 0.001
UA3	- 1.561	0.061	< 0.001	- 2.650	0.129	< 0.001
PD2	- 0.838	0.043	< 0.001	- 1.398	0.085	< 0.001
PD3	- 2.687	0.068	< 0.001	- 5.056	0.194	< 0.001
WSU2	- 0.503	0.044	< 0.001	- 0.872	0.080	< 0.001
WSU3	- 1.808	0.058	< 0.001	- 3.214	0.132	< 0.001
Domain importance	PD > WSU > MO > UA > LAM			PD > WSU > MO > UA > LAM		
Goodness of fit and prediction accuracy						
Log-likelihood	- 6480			- 5602		
BIC	13056			11830		
MAE	0.0474			0.0348		
MSE	0.0040			0.0022		

BFGS Broyden–Fletcher–Goldfarb–Shanno, *BIC* Bayesian information criterion, *LAM* looking after myself, *MAE* mean absolute error, *MO* mobility, *MSE* mean squared error, *PD* having pain or discomfort, *SE* standard error, *UA* doing usual activities, *WSU* feeling worried, sad or unhappy

^aThe mixed logit model with random correlated coefficients was estimated using 5000 pseudo-random draws and the BFGS optimisation algorithm

^b*p* values were estimated for the difference from the previous level

feasibility of including adolescents in valuation of the EQ-5D-Y-3L [43–45]. This is also supported by evidence from valuation studies of other instruments, such as the CHU9D and vignette-based studies [46, 47]. In DCE studies with the EQ-5D-Y-3L in the UK, Spain, Germany and Slovenia, adolescents gave less importance to ‘doing usual activities’, ‘having pain or discomfort’ and ‘feeling worried, sad or unhappy’ than adults valuing health states for a child [18, 19]. Furthermore, the perspective is also challenging as there may be differences in the person respondents imagine (e.g. themselves as a child, their own child, another child they know, or no particular child) [42]. A recent study has shown, for example, that respondents are less willing to trade life-years on behalf of others [22]. A promising new approach is to elicit preferences from a mixed sample of adults and adolescents [24].

There are numerous strengths of this study. The data collection was among the first wave of youth EQ-5D valuation studies conducted in over 10 countries, including Belgium, China, Germany, Indonesia, Japan, The Netherlands, Slovenia and Spain [34, 40, 41, 48, 49]. Furthermore, a similar EQ-VT protocol and the same software and interviewers were used in this study as for the valuation of the adult EQ-5D-3L and EQ-5D-5L in Hungary in 2018–2019, maintaining methodological consistency [7]. However, this study also has some limitations. First,

preference data were collected during the COVID-19 pandemic with varying levels of stringency and vaccine availability during the study period. It is unknown if the pandemic had an impact on how the general population valued health states, in particular whether these utilities are different from pre-COVID utilities and whether they will remain stable for the post-pandemic era. Findings from a recent study comparing EQ-5D-5L health state valuations (by using EQ VAS) before and during the pandemic in the UK suggest that COVID-19 may have affected health preferences among members of the general public [16]. Nevertheless, findings from EQVAS values may not be generalisable to cTTO utilities. Second, the online DCE survey may be subject to selection bias because digital literacy and internet access were prerequisites to participate in the study. Furthermore, 255 respondents were excluded from the DCE survey due to speeding or providing inconsistent responses which decision may be considered overly strict with fast decision makers and those susceptible for random mistakes. Third, despite the overall good representativeness and similarities in terms of sociodemographic characteristics, the DCE and cTTO samples differed in their recruitment and incentivisation strategies and this may represent a further limitation. Fourth, the cTTO task involved one block of 10 health states that somewhat limited our options for modelling. Finally, in both the DCE and cTTO tasks, health states were valued considering the health

Table 4 Anchoring DCE onto cTTO utilities

	DCE anchored at the worst health state (33333)	DCE mapped onto cTTO (value set)
Mobility		
Some problems walking about	- 0.055	- 0.054
A lot of problems walking about	- 0.262	- 0.257
Looking after myself		
Some problems washing or dressing	- 0.039	- 0.038
A lot of problems washing or dressing	- 0.193	- 0.189
Doing usual activities		
Some problems doing usual activities	- 0.080	- 0.078
A lot of problems doing usual activities	- 0.258	- 0.252
Having pain or discomfort		
Some pain or discomfort	- 0.136	- 0.133
A lot of pain or discomfort	- 0.492	- 0.481
Feeling worried, sad or unhappy		
A bit worried, sad or unhappy	- 0.085	- 0.083
Very worried, sad or unhappy	- 0.312	- 0.306
MAE (predicted vs. observed)		
All states	0.0486	0.0485
Mild states	0.0311	0.0298
Utilities		
U (mildest not 11111)	0.961	0.962
U (33333)	- 0.517	- 0.485
Mean utility (SD)	0.363 (0.303)	0.376 (0.297)
Median utility (IQR)	0.385 (0.437)	0.398 (0.427)
N (%) of health states WTD (of 243)	32 (13.2)	30 (12.3)

cTTO composite time trade-off, DCE discrete choice experiment, IQR interquartile range, MAE mean absolute error, SD standard deviation, WTD worse than dead

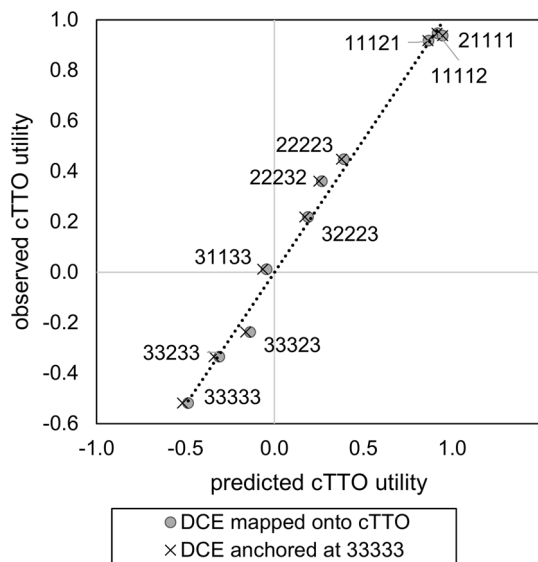


Fig. 2 Agreement between predicted and observed utilities. cTTO composite time trade-off, DCE discrete choice experiment

of a 10-year-old child. Further research is warranted if such preferences are representative for the entire age range of the instrument (8–15 years) [50, 51].

5 Conclusions

This study provided a national value set of the EQ-5D-Y-3L for Hungary based on general adult population preferences. The value set allows to estimate utilities from EQ-5D-Y-3L responses and can be used for cost-utility analyses of health technologies for young populations. Given the widespread use and acceptance of the adult EQ-5D questionnaires by researchers, physicians, analysts and policymakers in health-care in Hungary, it is hoped that the EQ-5D-Y-3L value set will soon become adopted by users.

Supplementary Information The online version contains supplementary material available at <https://doi.org/10.1007/s40273-022-01190-2>.

Acknowledgements The authors are grateful to Elly Stolk, Juan Manuel Ramos-Goñi, Bram Roudijk and Anabel Estévez-Carrillo for their support with the study design, data collection or data analysis. They would also like to thank the excellent work of the interviewers, Mercédesz Angyal, Zita Bagdi and Anna Nikl.

Funding Open access funding provided by Corvinus University of Budapest.

Declarations

Funding Data collection was supported by the Higher Education Institutional Excellence Program 2020 of the Ministry of Human Capacities in the framework of the 'Financial and Public Services' research project (TKP2020-IKA-02) at the Corvinus University of Budapest and by the EuroQol Research Foundation (192-2020VS). Fanni Rencz's work was supported by the János Bolyai Research Scholarship of the Hungarian Academy of Sciences (BO/00304/21) and the New National Excellence Program of the Ministry for Innovation and Technology in Hungary (ÚNKP-21-5-CORVINUS-134 and ÚNKP-22-5-CORVINUS-4).

Conflict of interest Fanni Rencz and Zhihao Yang are active members of the EuroQol Group. Aureliano Paolo Finch is a member of the EuroQol Group and is employed by the EuroQol Office, EuroQol Research Foundation. The views expressed in this article are those of the authors and do not necessarily reflect those of the EuroQol Research Foundation. Gábor Ruzsa, Alex Bató, and Valentin Brodzsky have no conflicts of interest to declare.

Availability of data and material All data in this study are available from the corresponding author upon reasonable request.

Code availability The code used for the statistical analysis is available from the corresponding author upon reasonable request.

Ethics approval All procedures performed in studies involving human participants were in accordance with the ethical standards of the institutional and/or national research committee and with the 1964 Helsinki declaration and its later amendments or comparable ethical standards. Ethical approval was obtained from the Research Ethics Committee of the Corvinus University of Budapest (no. KRH/31/2021).

Informed consent Informed consent was obtained from all patients included in this study.

Disclosure This article has been published in a special edition journal supplement wholly funded by the EuroQol Research Foundation.

Open Access This article is licensed under a Creative Commons Attribution-NonCommercial 4.0 International License, which permits any non-commercial use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. The images or other third party material in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit <http://creativecommons.org/licenses/by-nc/4.0/>.

References

1. Banta D. The development of health technology assessment. *Health Policy*. 2003;63(2):121–32.
2. Sharma D, Aggarwal AK, Downey LE, Prinja S. National Healthcare Economic Evaluation guidelines: a Cross-Country comparison. *Pharmacoecon Open*. 2021;5(3):349–64.
3. Finch AP, Brazier JE, Mukuria C. What is the evidence for the performance of generic preference-based measures? A systematic overview of reviews. *Eur J Health Econ*. 2018;19(4):557–70.
4. Kennedy-Martin M, Slaap B, Herdman M, van Reenen M, Kennedy-Martin T, Greiner W, et al. Which multi-attribute utility instruments are recommended for use in cost-utility analysis? A review of national health technology assessment (HTA) guidelines. *Eur J Health Econ*. 2020;21(8):1245–57.
5. Rencz F, Gulacsi L, Drummond M, Golicki D, Prevotnik Rupel V, Simon J, et al. EQ-5D in Central and Eastern Europe: 2000–2015. *Qual Life Res*. 2016;25(11):2693–710.
6. Guideline on the conduct and appraisal of health economic evaluations by the Ministry of Human Capacities in Hungary. [Az Emberi Erőforrások Minisztériuma egészségügyi szakmai irányelve az egészség-gazdaságtani elemzések készítéséhez és értékeléséhez.] *Egészségügyi Közlöny* 2021;71(21):2178–200.
7. Rencz F, Brodzsky V, Gulacsi L, Golicki D, Ruzsa G, Pickard AS, et al. Parallel valuation of the EQ-5D-3L and EQ-5D-5L in Hungary. *Value Health*. 2020;23(9):1235–45.
8. Matza LS, Patrick DL, Riley AW, Alexander JJ, Rajmil L, Pleil AM, et al. Pediatric patient-reported outcome instruments for research to support medical product labeling: report of the ISPOR PRO good research practices for the assessment of children and adolescents task force. *Value Health*. 2013;16(4):461–79.
9. Kreimeier S, Oppe M, Ramos-Goñi JM, Cole A, Devlin N, Herdman M, et al. Valuation of EuroQol Five-Dimensional Questionnaire, Youth Version (EQ-5D-Y) and EuroQol Five-Dimensional Questionnaire, Three-Level Version (EQ-5D-3L) health states: the impact of wording and perspective. *Value Health*. 2018;21(11):1291–8.
10. Rowen D, Rivero-Arias O, Devlin N, Ratcliffe J. Review of valuation methods of preference-based measures of health for economic evaluation in child and adolescent populations: where are we now and where are we going? *Pharmacoeconomics*. 2020;38(4):325–40.
11. Kwon J, Freijser L, Huynh E, Howell M, Chen G, Khan K, et al. Systematic review of conceptual, age, measurement and valuation considerations for generic multidimensional childhood patient-reported outcome measures. *Pharmacoeconomics*. 2022;40(4):379–431.
12. Rowen D, Keetharuth AD, Poku E, Wong R, Pennington B, Wailoo A. A review of the psychometric performance of selected child and adolescent preference-based measures used to produce utilities for child and adolescent health. *Value Health*. 2021;24(3):443–60.
13. Wille N, Badia X, Bonsel G, Burström K, Cavrini G, Devlin N, et al. Development of the EQ-5D-Y: a child-friendly version of the EQ-5D. *Qual Life Res*. 2010;19(6):875–86.
14. Ravens-Sieberer U, Wille N, Badia X, Bonsel G, Burström K, Cavrini G, et al. Feasibility, reliability, and validity of the EQ-5D-Y: results from a multinational study. *Qual Life Res*. 2010;19(6):887–97.
15. Kind P, Klose K, Gusi N, Olivares PR, Greiner W. Can adult weights be used to value child health states? Testing the influence of perspective in valuing EQ-5D-Y. *Qual Life Res*. 2015;24(10):2519–39.

16. Webb EJD, Kind P, Meads D, Martin A. Does a health crisis change how we value health? *Health Econ.* 2021;30(10):2547–60.
17. Shah KK, Ramos-Goñi JM, Kreimeier S, Devlin NJ. An exploration of methods for obtaining 0 = dead anchors for latent scale EQ-5D-Y values. *Eur J Health Econ.* 2020;21(7):1091–103.
18. Mott DJ, Shah KK, Ramos-Goñi JM, Devlin NJ, Rivero-Arias O. Valuing EQ-5D-Y-3L health states using a discrete choice experiment: do adult and adolescent preferences differ? *Med Decis Mak.* 2021;41(5):584–96.
19. Prevolnik Rupel V, Ramos-Goñi JM, Ogorevc M, Kreimeier S, Ludwig K, Greiner W. Comparison of adult and adolescent preferences toward EQ-5D-Y-3L health states. *Value Health.* 2021;24(9):1350–9.
20. Dewilde S, Janssen MF, Lloyd AJ, Shah K. Exploration of the reasons why health state valuation differs for children compared with adults: a mixed methods approach. *Value Health.* 2022;25(7):1185–95.
21. Powell PA, Rowen D, Rivero-Arias O, Tsuchiya A, Brazier JE. Valuing child and adolescent health: a qualitative study on different perspectives and priorities taken by the adult general public. *Health Qual Life Outcomes.* 2021;19(1):222.
22. Lipman SA, Reckers-Droog VT, Karimi M, Jakubczyk M, Attema AE. Self vs. other, child vs. adult. An experimental comparison of valuation perspectives for valuation of EQ-5D-Y-3L health states. *Eur J Health Econ.* 2021;22(9):1507–18.
23. Ramos-Goñi JM, Oppe M, Stolk E, Shah K, Kreimeier S, Rivero-Arias O, et al. International Valuation Protocol for the EQ-5D-Y-3L. *Pharmacoeconomics.* 2020;38(7):653–63.
24. Rowen D, Mukuria C, Powell PA, Wailoo A. Exploring the issues of valuing child and adolescent health states using a mixed sample of adolescents and adults. *Pharmacoeconomics.* 2022;40(5):479–88.
25. Versteegh MM, Brouwer WBF. Patient and general public preferences for health states: a call to reconsider current guidelines. *Soc Sci Med.* 2016;165:66–74.
26. Jonker MF, Donkers B, de Bekker-Grob E, Stolk EA. Attribute level overlap (and color coding) can reduce task complexity, improve choice consistency, and decrease the dropout rate in discrete choice experiments. *Health Econ.* 2019;28(3):350–63.
27. Ratcliffe J, Chen G, Stevens K, Bradley S, Couzner L, Brazier J, et al. Valuing Child Health Utility 9D health states with young adults: insights from a time trade off study. *Appl Health Econ Health Policy.* 2015;13(5):485–92.
28. Xie F, Gaebel K, Perampaladas K, Doble B, Pullenayegum E. Comparing EQ-5D valuation studies: a systematic review and methodological reporting checklist. *Med Decis Making.* 2014;34(1):8–20.
29. Stolk E, Ludwig K, Rand K, van Hout B, Ramos-Goñi JM. Overview, update, and lessons learned from the international EQ-5D-5L valuation work: version 2 of the EQ-5D-5L Valuation Protocol. *Value Health.* 2019;22(1):23–30.
30. Janssen BM, Oppe M, Versteegh MM, Stolk EA. Introducing the composite time trade-off: a test of feasibility and face validity. *Eur J Health Econ.* 2013;14(Suppl 1):S5–13.
31. Oppe M, Devlin NJ, van Hout B, Krabbe PF, de Charro F. A program of methodological research to arrive at the new international EQ-5D-5L valuation protocol. *Value Health.* 2014;17(4):445–53.
32. Ramos-Goñi JM, Oppe M, Slaap B, Busschbach JJ, Stolk E. Quality control process for EQ-5D-5L valuation studies. *Value Health.* 2017;20(3):466–73.
33. Hungarian Central Statistical Office, Microcensus 2016. ISBN 978-963-235-494-1ö. Budapest, Hungary. Available at: http://www.ksh.hu/docs/eng/xftp/idoszaki/microcensus2016/microcensus_2016_3.pdf. Accessed 24 Jan 2022.
34. Ramos-Goñi JM, Oppe M, Estévez-Carrillo A, Rivero-Arias O. Accounting for unobservable preference heterogeneity and evaluating alternative anchoring approaches to estimate country-specific EQ-5D-Y value sets: a case study using Spanish preference data. *Value Health.* 2022;25(5):835–43.
35. Tervonen T, Schmidt-Ott T, Marsh K, Bridges JFP, Quaipe M, Janssen E. Assessing rationality in discrete choice experiments in health: an investigation into the use of dominance tests. *Value Health.* 2018;21(10):1192–7.
36. Croissant Y. Estimation of multinomial logit models in R: The mlogit Packages. (2012) R package version 0.2-2. Available at: <http://cran.r-project.org/web/packages/mlogit/vignettes/mlogit.pdf> Accessed 4 Apr 2022.
37. McFadden D, Train K. Mixed MNL models for discrete response. *J Appl Economet.* 2000;15(5):447–70.
38. Roudijk B, Ludwig K, Devlin N. EQ-5D-5L value set summaries. Value sets for EQ-5D-5L. Cham: Springer; 2022. p. 55–212.
39. Rowen D, Brazier J, Van Hout B. A comparison of methods for converting DCE values onto the full health-dead QALY scale. *Med Decis Mak.* 2015;35(3):328–40.
40. Kreimeier S, Mott D, Ludwig K, Greiner W. EQ-5D-Y value set for Germany. *Pharmacoeconomics.* <https://doi.org/10.1007/s40273-022-01143-9>. (Epub 23 May 2022).
41. Prevolnik Rupel V, Ogorevc M. EQ-5D-Y value set for Slovenia. *Pharmacoeconomics.* 2021;39(4):463–71.
42. Lipman SA, Reckers-Droog VT, Kreimeier S. Think of the children: a discussion of the rationale for and implications of the perspective used for EQ-5D-Y health state valuation. *Value Health.* 2021;24(7):976–82.
43. Åström M, Rolfson O, Burström K. Exploring EQ-5D-Y-3L experience-based VAS values derived among adolescents. *Appl Health Econ Health Policy.* 2022;20(3):383–93.
44. Dalziel K, Catchpool M, García-Lorenzo B, Gorostiza I, Norman R, Rivero-Arias O. Feasibility, validity and differences in adolescent and adult EQ-5D-Y health state valuation in Australia and Spain: an application of best-worst scaling. *Pharmacoeconomics.* 2020;38(5):499–513.
45. Åström M, Conte H, Berg J, Burström K. “Like holding the axe on who should live or not”: adolescents’ and adults’ perceptions of valuing children’s health states using a standardised valuation protocol for the EQ-5D-Y-3L. *Qual Life Res.* 2022;31(7):2133–42.
46. Ratcliffe J, Huynh E, Stevens K, Brazier J, Sawyer M, Flynn T. Nothing about us without us? A comparison of adolescent and adult health-state values for the child health utility-9D using profile case best-worst scaling. *Health Econ.* 2016;25(4):486–96.
47. Saigal S, Stoskopf BL, Feeny D, Furlong W, Burrows E, Rosenbaum PL, et al. Differences in preferences for neonatal outcomes among health care professionals, parents, and adolescents. *JAMA.* 1999;281(21):1991–7.
48. Devlin N, Pan T, Kreimeier S, Verstraete J, Stolk E, Rand K, et al. Valuing EQ-5D-Y: the current state of play. *Health Qual Life Outcomes.* 2022;20(1):105.
49. Shiroiwa T, Ikeda S, Noto S, Fukuda T, Stolk E. Valuation survey of EQ-5D-Y based on the International Common Protocol: development of a value set in Japan. *Med Decis Mak.* 2021;41(5):597–606.
50. Reckers-Droog V, Karimi M, Lipman S, Verstraete J. Why do adults value EQ-5D-Y-3L health states differently for themselves than for children and adolescents: a think-aloud study. *Value Health.* 2022;25(7):1174–84.
51. Ramos-Goñi JM, Estévez Carrillo A, Rivero-Arias O, Rowen D, Mott D, Shah K, et al. Does changing the age of a child to be considered in 3-level version of EQ-5D-Y discrete choice experiment-based valuation studies affect health preferences? *Value Health.* 2022;25(7):1196–204.