A Systematic Review of Health State Utility Values for Influenza and Influenza-like Illness

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INTRODUCTION



Influenza imposes a substantial clinical and economic burden on health care facilities. It causes acute lower respiratory infection and hospitalizations especially among the elderly.

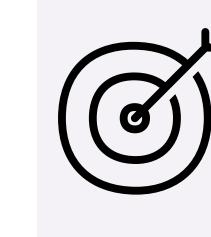


Valid and reliable information on the impact of influenza and influenza-like illness (ILI) in terms of health state utility values (HSUVs) is an essential input for health economic evaluation (e.g., cost-effectiveness or cost-utility analysis (CEA/CUA)) assessing health care interventions against influenza and ILI - such as the use of vaccines.

Table 1: Study characteristics and results

Study	Assessed groups	Method/instrument	Valued influenza/ILI health state and utility weight
Author-based assessm	nent		
Burls et al. (2002)	Adults	EQ-5D-3L	Influenza: 0.516

OBJECTIVE



To give a comprehensive overview of international studies across all populations that estimated the effects of influenza and ILI on health-related quality of life (HRQoL) expressed as HSUVs.

METHODS

We conducted a systematic literature search in MEDLINE and Embase to identify articles published from 1990/01/01 to 2023/08/15 in English, German, Portuguese, or Spanish

R	ESULTS										
Figu	re 1: PRISMA flowcha										
Identification	database search (n = 5,203) MEDLINE: 1,602	through manual search									
	Records (n = 5,262)	Duplicates removed (n = 1,137)									

to 2023/08/15 in English, Germa language. Articles reporting data		Mauskopf et al. (2000)	Patients (>12 years); applied in CEA	QWB	Influenza/ILI: 0.	5579		
with influenza or ILI were include	ed. A quality assessment	Muennig & Khan (2001)	Otherwise healthy adults (15–65 years); applied in CEA	QWB	ILI: 0.61			
was performed based on 10 pre- Data extraction encompassed:	·	Rothberg et al. (2005)		a) EQ-5D, and b) Assumption	 a) Influenza, not hospitalized: 0.32 b) Influenza, hospitalized: 0.20^a 			
Main study characteristics: stu		Expert-based assessment						
for HSUV, sample size, and as assessment and timepoint.	s instrument for	Griffin et al. (2001)	Hypothetical high-risk patients	EQ-5D	Influenza in a hypothetical high-risk patient:-0.263			
 Findings related to baseline an 		Stouthard et al. (1997)	Dutch population	Person trade-off and VAS, and interpolation	Influenza (episode of 2 weeks): 0.84			
with respective mean utility we health states.	eights and duration of				a) Influenza ('moderate to severe respiratory illness with bed rest'):			
		Stratton et al. (2000)	Children and adults (all ages)	HUI2	c) Influenza, ICI	spitalized (acute care): 0.65 J: 0.52 ecuperation following bed re		
RESULTS		Self-assessment			,		•	
Figure 1: PRISMA flowchart	Additional records identified through manual search (n = 59)	Bilcke et al. (2014)	Children (0-17 years) and adults (≥18 years)	SF-12v2 Health Survey	 a) ILI, no medical care: 0.7 b) ILI, outpatient: 0.68 c) ILI, inpatient: 0.61 d) Clinically diagnosed influenza, outpatient: 0.68 e) Clinically diagnosed influenza, inpatient: 0.62 			
Records identified through database search (n = 5,203) MEDLINE: 1,602 Embase: 3,601		Brady et al. (2001)	Not reported ^b	HUI3	a) Influenza, outpatient: 0.636 b) Influenza, inpatient: 0.35			
MEDLINE: 1,602 Embase: 3,601		Camacho et al. (2013) ^c	Children (0-17 years) and adults (≥18					
		Chung et al. (2022)	years) Adults (≥16 years)	used as quality weights ^c	Laboratory-confirmed influenza a) Within 3 days if illness onset; VAS: 48.5 b) 4-7 days after illness onset; VAS: 52.2			
Records (n = 5,262)	→ Duplicates removed (n = 1,137)	Fragaszy et al. (2018)	Children and adults (all ages)	EQ-5D-3L	a) ILI: 0.48 b) Influenza A (H1N1pdm09 PCR positive): 0.44 c) Influenza B (PCR positive): 0.36			
		Griffin et al. (2001)	Adults (≥18 years)	EQ-5D	Confirmed influenza illness a) Whole influenza illness: -0.066 b) Worst day of influenza: -0.342			
tion	Articles excluded on basis of	Hollmann et al. (2013)	Children (8-17 years) and adults	EQ-5D-3L	a) Confirmed influenza H1N1, outpatient: 0.50 b) Confirmed influenza H1N1, inpatient: 0.23			
Articles screened (n = 4,125)	Articles screened (n = 4,125) Articles meeting exclusion		Adults (≥60 years)	EQ-5D-5L	Laboratory-confirmed influenza, outpatient a) 0.54 ^d (week 0) b) 0.664 (week 1 after symptom onset) c) 0.75 ^d (week 2 after symptom onset) d) 0.77 ^d (week 3 after symptom onset) e) 0.81 ^d (week 4 after symptom onset)			
Full-text articles included for screening (n = 364)	criteria (n = 339) No influenza/ILI: 16 Methods/data source unclear: 6 No primary HSUV source: 148 No HSUVs reported: 94 Study type: 24 Language: 6 Publication type: 40 Duplicates: 5	O'Brien et al. (2003)	Otherwise healthy adults (16-64 years)	VAS ^e	 a) Confirmed influenza within 48 hours of symptom onset, day 1-7: 0.40 - 0.79 b) ILI within 48 hours of symptom onset, day 1-7: 0.40 - 0.78 			
		Osborne et al. (2000)	Otherwise healthy adults	AQoL	Symptoms of in	fluenza, day 0-7: 0.77 - 0.8	5	
Articles included in review		Pradas Velasco et al. (2009)	Otherwise healthy working-age patients	EQ-5D-3L	Confirmed influenza Social tariff based on VAS: 0.437 Social tariff based on TTO: 0.294			
Articles included in review (n = 25)		Rombach et al (2022)	At risk children (6 months to 12 years)	EQ-5D-Y and EQ-5D-3L	ILI EQ-5D-Y Proxy, day 0-28: 0.568 - 0.923 EQ-5D-Y Child, day 0-28: 0.531 - 0.941			
Characteristics of included stud	lies:	Rothberg et al. (2003)	Working-age patients (<65 years) and healthcare workers	HUI-3	a) Influenza, not hospitalized: 0.25 b) Influenza, hospitalized: 0.20 ^f			
14 studies were conducted in E and 2 each in Asia and Australia	• •	Thorrington et al. (2017)	Children in primary schools	EQ-5D	ILI: 0.2			
Many of the included utility stud		Tsuzuki & Yoshihara (2020)	Individuals (all ages)	SF-12v2 Standard	a) Influenza: 0.66 b) ILI: 0.66			
part of model-based economic of Population sizes in the studies in subjects.		Turner et al. (2003)	Otherwise healthy adults (12-65 years) and elderly/high-risk patients (≥65 years)	Likert scale (0 to 10), Likert scores were converted TTO equivalent scores	 a) Confirmed Influenza in healthy adults, day 1-21: 0.067543 - 0.86484 b) Confirmed Influenza in elderly and high-risk, day 1-21: 0.117313 - 0.668569 			
Quality of included studies:	lity: only 6 studios mot all	Van Hoek et al. (2011)	Children and adults	EQ-5D	a) Influenza A/H1N1v (PCR positive): 0.29 b) ILI (PCR negative): 0.34			
We observed varying study qua pre-defined and applicable qual Results of included studies: Utility values showed great hete	ity criteria.	Vindt Holm et al. (2004)	Otherwise healthy adolescents and adults (13-64 years)	11-point VAS Likert scales for health, sleep and usual activities and conversion into TTO equivalent scores	Influenza and ILI, day 1-21: 0.112936 - 0.86484			
the worst day of infection was -(laboratory-confirmed influenza t Total range was -0.342 for wors to 0.941 for ILI in at risk childrer	0.342 in patients with to 0.48 in patients with ILI. It day of influenza infection	Yang et al. (2017)	Children (0-15 years) ^g and adults (≥16 years)	EQ-5D-3L; telephone interview and EQ-5D proxy	Age (years) 0-4 5-15 16-59 ≥60	a) Influenza outpatient 0.6286 0.6216 0.5939 0.5733	b) Influenza inpatient 0.5900 0.6132 0.4913 0.4128	

- Only one study reported age-specific values.
- Observed differences between values potentially reflect the variety of study populations and timepoints of assessment.

Abbr. AQoL Assessment of Quality of Life; CEA cost-effectiveness analysis; CUA cost-utility analysis; EQ-5D EuroQol five-dimensional questionnaire; HUI Health Utility Index; ILI influenza like illness; PCR polymerase chain reaction; QoL Quality of Life; QWB Quality of Well-Being scale; RCT randomized controlled trial; SF-12v2 Short-Form 12 Items (version 2); TTO time trade-off; UK United Kingdom; VAS visual analogue scale

^aAuthor assumption ^bHUI3 assessment by 11 adults, but it is unclear from the original source whether they had influenza themselves. ^cTo obtain utility weights, the scores were divided by 100. ^dValues were reported graphically only and approximated using WebPlotDigitizer. ^eVAS scores were normalized into quality weights (0 to 1). ^fEstimate ^gFor children younger than 18 months the mobility and selfcare dimensions were not scored (n=91).

CONCLUSIONS

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- This systematic review revealed varying utility weights for influenza and ILI across published studies.
- Since methods used to estimate utility weights also differed considerably, researchers should take care when selecting HSUVs for further analysis, such as CEA/CUA.

CONFLICTS OF INTEREST, FUNDING & Disclosure

RH, FS, JT, and TS declare that there are no conflicts of interest regarding the research, authorship and/or publication of this abstract. They are employees of WIG2 GmbH. WIG2 GmbH received funding from Sanofi-Aventis Deutschland GmbH, Berlin, Germany for the conduct of the study. AP, OD, FA are employees of Sanofi and may hold shares and/or stock options in the company.



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