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# Systematic Review

# Evaluation of quality of life related to nutritional status

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The way in which the quality of life related to health (HRQoL) is affected by the nutritional status of the patient is a subject of constant interest and permanent debate. The purpose of the present paper is to review those studies that relate HRQoL to nutritional status and examine the tools (questionnaires) that they use to investigate this relationship. A critical review of published studies was carried out via an investigation of the following databases: MEDLINE (via PubMed); EMBASE; The Cochrane Library; Cumulative Index to Nursing and Allied Health Literature (CINAHL); Institute for Scientific Information (ISI) Web of Science; Latin American and Caribbean Health Sciences Literature (LILACS); Spanish Health Sciences Bibliographic Index (IBECS). The search was carried out from the earliest date possible until July 2007. The medical subject heading terms used were 'quality of life', 'nutritional status' and 'questionnaires'. The articles had to contain at least one questionnaire that evaluated quality of life. Twenty-eight documents fulfilling the inclusion criteria were accepted, although none of them used a specific questionnaire to evaluate HRQoL related to nutritional status. However, some of them used a combination of generic questionnaires with the intention of evaluating the same. Only three studies selectively addressed the relationship between nutritional status and quality of life, this evaluation being performed not by means of specific questionnaires but by statistical analysis of data obtained via validated questionnaires.

Quality of life: Nutritional status: Nutritional sciences: Questionnaires: Healthcare evaluation mechanisms

The concept of quality of life related to health (HRQoL) is defined with regard to the way in which illness (as a source of pain, physical dysfunction and discomfort) imposes limitations or alterations on everyday behaviour, social activities and psychological wellbeing, as well as in other aspects of personal daily life<sup>(1)</sup>.

The measurement of quality of life brings a holistic dimension to the burden of a clinical state or to the response to an operation. The relationship between quality of life and nutritional status is not well studied. Furthermore, measuring HRQoL is a complex process, being, as it is, a subjective, multifactor construct responsive to individual expectations in different facets of life. The way in which HRQoL is affected by the nutritional status of the patient is a subject of constant interest and permanent debate. It is all too well known that an

impoverishment of nutritional status leads to a decrease in physiological function, increasing the risk of complications and septic death<sup>(2,3)</sup>, that there is a significant correlation between nutrition and alterations in muscular, immune and cognitive functions<sup>(4)</sup> and therefore that an improvement in nutritional status is an influencing factor in the improvement of physiological function<sup>(5,6)</sup>.

The necessity and importance of the measurement of HRQoL, both general and specific, tied to a definite concept, can be justified on the basis of studies which show that perceived health is independently associated with medium-term mortality<sup>(7,8)</sup>. These specific instruments, designed to relate a patient's HRQoL to a specific pathology, have grown in importance in recent years. They also provide a subset of relevant data which point to a positive causality<sup>(9)</sup>.

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Consequently, the purpose and objective of the present study is to bring together those studies that relate HRQoL with nutritional status and examine the tools (questionnaires) that they use to investigate this relationship.

### Methods

Bibliographic search

Given the hierarchical structure of medical subject heading (MeSH) terms, the terms 'quality of life', 'nutritional status' and 'questionnaires' were chosen and used in conjunction with the Boolean link 'AND'.

The search was carried out from the earliest date possible (according to each database) until July 2007, the latest date considered in the present study.

In the only databases that permitted it, MEDLINE and EMBASE, the major (Majr) topic terms were used. These represent the most important concepts of an article and help to eliminate less relevant studies from the results, thereby increasing the sensitivity of the search ('quality of life' [Majr] AND 'nutritional status' [Majr] AND 'questionnaires' [MeSH]).

'Humans' was used in all databases as a search limit.

Additionally, as a secondary search, the bibliographies of the selected articles were reviewed in order to identify studies not found by the primary search.

The databases MEDLINE (via PubMed), EMBASE, The Cochrane Library, Cumulative Index to Nursing and Allied Health Literature (CINAHL), Institute for Scientific Information (ISI) Web of Science, Latin American and Caribbean Health Sciences Literature (LILACS) and Spanish Health Sciences Bibliographic Index (IBECS) were consulted.

Selection of the articles

The articles were selected via inclusion and exclusion criteria previously defined in a written protocol (10,11).

Inclusion criteria were:

- (1) Use of at least one questionnaire that evaluates quality of life;
- (2) Nutritional status of the studied individuals is, by whatever means, taken into account;
- (3) Original articles published in peer-reviewed journals.

Excluded were studies that measured HRQoL using only clinical indicators.

Validity check

The studies, with no indication of the authors, journal or database origin, were checked for relevance by the three experts in nutrition (C. W.-B., J. M. Culebras and J. Alvarez) using a yes/no checklist<sup>(11)</sup>.

Concordance analysis between the experts in nutrition (gold standard) and the obtained results gave the following results: observed agreement 90·00 (95 % CI 80·70, 99·30) %;  $\kappa$  75 (95 % CI 52, 98) %; significance test 4·74 (P<0·001); sensitivity 93·10 (95 % CI 83·88, 100) %; specificity 81·82 (95 % CI 59·03, 100) %. The silent percentage (relevant articles not found) and the noisy percentage (non-relevant articles found) were 5 (95 % CI 0, 11·75) % in both cases.

Special characteristics of the study

Although it is preferable to base a systematic review on prospective studies or studies with adequate follow-up periods, it was decided to include cross-sectional studies or studies with short follow-up periods if HRQoL had been studied using a valid questionnaire and the nutritional status of the observed patients had been taken into account. This limitation will be discussed later.

### Results

Twenty-nine papers from MEDLINE, twenty-one from EMBASE, six from the Cochrane Library and thirteen from CINAHL were obtained. All the papers found in the bibliographic database ISI Web of Science had been previously collected. No articles were found in the databases LILACS or IBECS. After eliminating redundant papers, forty documents were obtained.

Agreement between the scientific documentation experts (J. S.-V., V. Juan-Quilis and R Ballester Añon; applying the most sensitive search formula) and the experts in nutrition reduced the number of studies to thirty-one<sup>(11)</sup>.

A further study was discarded for measuring user satisfaction with nutrition services, rather than quality of life, and for not using a questionnaire that evaluated quality of life. Finally, twenty-eight documents on quality of life related to nutritional status were accepted<sup>(12–39)</sup> (Table 1).

It is worth noting how recent the studies are; the average age is 3.85 (95 % CI 2.62, 5.02) years, and the average obsolescence gives a value of 3 years and a Price index of 75 % (percentage of articles 5 years old or less).

The designs of the studied articles were: eight (28·57%) clinical trials; eleven (39·29%) prospective; seven (25·00%) cross-sectional; two (7·14%) retrospective. The disparity in design can be seen in the wide diversity of follow-up periods. The number of patients also varied widely, from a minimum of twelve to a maximum of 367.

Quality of life related to health and nutritional status

Although there are a considerable number of published studies on HRQoL, those that truly evaluate quality of life related to nutritional status are scarce. However, it is worth noting that of the articles relating HRQoL to nutritional status, eleven (39·27%) had cancer as a pathological base (13,16,17,21,27,29,30,33,35,37,38).

The review found no specific questionnaire that determined a direct link between HRQoL and nutritional status. However, three papers (10·71%) detailed a significant correlation between nutritional status and HRQoL using a valid method for measuring quality of life<sup>(19,23,29)</sup>. Another article (3·57%) referred to a possible relationship between HRQoL and nutrition<sup>(25)</sup>, but drew attention to other important factors, such as the risk of depression. A different study (3·57%) mentioned how the ingestion of foodstuffs affects HRQoL<sup>(15)</sup>, although a further paper<sup>(33)</sup> (3·57%) found no significant effect between the results obtained using The Short Form-36 Health Survey (SF-36) questionnaire and nutritional intervention. In another, an association between a deteriorating HRQoL and severe malnutrition was seen<sup>(34)</sup>.

Table 1. Summary of the twenty-eight reviewed studies in chronological order according to year of publication

Reference	Study design	QoL questionnaire	Patients included	Follow-up	Relationship between QoL and nutritional status in study results
O'Keefe <i>et al.</i> (2007) <sup>(12)</sup>	Prospective	Subjective assessment of QoL (transplantation)	Group 1: transplant patients, <i>n</i> 46, M/W 26/21, age range 22–66 years Group 2: total parenteral nutrition patients, <i>n</i> 13, M/W 10/3, age range 31–80 years	Mean 21 (range 12-36) months	QoL not evaluated in respect of nutritional status
Oates <i>et al.</i> (2007) <sup>(13)</sup>	Prospective	EORTC QLQ-C-30 and EORTC QLQ-H&N35	Fourteen patients with nasopharyngeal carcinoma Age range 27-71 years	2 years	These results emphasise the need for early nutritional intervention before commencing chemoradiotherapy
Kalaitzakis <i>et al.</i> (2006) <sup>(14)</sup>	Prospective	SF-36	128 adult patients with cirrhosis Control group: 299 age- and sex-matched reference sample from general population	6 months	Prospective follow-up studies are needed to fully clarify the role of gastrointestinal symptoms in HRQoL and nutritional status in liver cirrhosis
Trabal <i>et al.</i> (2006) <sup>(15)</sup>	Cross-sectional	EORTC QLQ-C-30	Fifty cancer patients M/W 28/22, mean age 61 (sp 14) years	3 d	The results point out that poor food intakes can affect QoL by themselves
Murawa <i>et al.</i> (2006) <sup>(16)</sup>	Prospective	Troidl	Thirty-one patients with stomach cancer-related total gastrectomy	49–127 months after surgery (mean 79-61	QoL was independent of the time elapsed from the surgery
Izutsu <i>et al.</i> (2006) <sup>(17)</sup>	Cross-sectional	WHOQOL-BREF	M/W 20/11, mean age 56·9 (sp 10·05) years 187 boys and 137 girls from urban non-slum area 157 boys and 121 girls from urban slum area Age range 11–18 years	(SD 23·41) months)	QoL was not evaluated in respect of nutritional status Living conditions, mental health, nutritional status and QoL of slum adolescents were worse than their non-slum counterparts QoL was not evaluated in respect of nutritional status
Gramignano <i>et al.</i> (2006) <sup>(18)</sup>	Prospective	QoL-OS and EuroQoL-5D	Twelve patients who had advanced solid tumours and reported fatigue M/W 2/10, age range 42–73 years	4 weeks	Improvement of symptoms with respect to fati gue and QoL in relation to oxidative stress may be explained mainly by an increase in lean body mass QoL was not evaluated in respect of nutritional status
Eriksson <i>et al.</i> (2005) <sup>(19)</sup>	Cross-sectional	SF-36	128 non-institutionalised individuals M/W 40/88, age range 70-75 years, mean age 72.9 (sp 1.5) years		Parts of the MNA can be interpreted as measurements of HRQoL Low values of SF-36 could also be used as predictors of risk of malnutrition
Allen (2005) <sup>(20)</sup>	Retrospective	OHIP-EDENT	Thirty-five edentulous adults who requested new complete dentures	Past 3 months	There was no association between diet and oral-related QoL
Ravasco <i>et al.</i> (2005) <sup>(21)</sup>	Clinical trial	EORTC QLQ-C-30	M/W 12/23, age range 52–77 years 111 colorectal cancer out-patients referred for radiotherapy M/W 66/45, age range 32–88 years, mean age 58 (sp 15) years Group 1 (n 37): individualised dietary counselling based on regular foods Group 2 (n 37): high-protein liquid supplement in addition to their usual diet Group 3 (n 37): the control group, patients were instructed to maintain their ad libitum intake	3 months	QoL was not evaluated in respect of nutritional status Dietary counselling was of similar or higher benefit, whereas even 3 months after radio therapy, it was the only method to sustain a significant impact on patient outcomes The prevalence of malnutrition at baseline was similar between the three study groups There was a linear positive association with the improvement in the patients' nutritional status
Scott <i>et al.</i> (2005) <sup>(22)</sup>	Clinical trial	SF-36	112 adult patients were recruited Intervention group: <i>n</i> 55, mean age 67-4 (sp 17) years Control group: <i>n</i> 57; mean age 68-6 (sp 17) years	12 months after PEG	QoL was not evaluated in respect of nutritional status

Table 1. Continued

Reference	Study design	QoL questionnaire	Patients included	Follow-up	Relationship between QoL and nutritional status in study results
Keller (2004) <sup>(23)</sup>	Cross-sectional	Non-validated author design	367 frail seniors (73·6 % women) Age range 54–100 years, mean age 79·3 (SD 7·9) years		Nutritional risk appears to be a significant and important factor associated with HRQoL Nutritional risk as measured by SCREEN appears to be a significant covariate in explaining differences in HRQoL among frail older adults
Gollub & Weddle (2004) <sup>(24)</sup>	Retrospective	Vailas <i>et al.</i>	Two groups Breakfast group: received a home-delivered breakfast and lunch, 5 d per week, <i>n</i> 167, age range 63–100 years, mean age 79·8 (sp 8·1) years Comparison group: received a home-delivered lunch 5 d per week, <i>n</i> 214, age range 60–100 years, mean age 77·7 (sp 9·1) years	Past 6 months	Both study groups rated global QoL and lone liness at average or moderate levels, with no group differences  QoL was not evaluated in respect of nutritional status
Hickson & Frost (2004) <sup>(25)</sup>	Cross-sectional	EuroQoL-5D	233 patients: M/F 104/129, age interquartile range 75–86 years. Two age groups Group 1: 65–74 years Group 2: 75 years and older		Study data suggest that a link exists between QoL and nutrition, but it may not be direct, and is influenced by other factors, especially a high risk of depression
Johansen <i>et al.</i> (2004) <sup>(26)</sup>	Prospective	SF-36	212 patients identified as being nutritionally at risk  Group 1: intervention, n 108, M/W 54/54, mean age 62-0 (sp 1-6) years  Group 2: control, n 104, M/W 48/56, mean age 62-4 (sp 1-7) years	> 4 d	The SF-36 QoL questionnaire did not reveal any convincing significant effect of nutritional intervention
Kennedy <i>et al.</i> (2004) <sup>(27)</sup>	Prospective	POQOLS	103 children and adolescents with acute lymphoblastic leukaemia  Age range 1–18 years, mean age 6-7 years  Group 1 (standard risk protocol): <i>n</i> 68, M/W  41/27, mean age 4-7 (sp 2-6) years  Group 2 (high-risk protocol): <i>n</i> 35, M/W  19/16, mean age 10-5 (sp 4-6) years	6 months after diagnosis	QoL was not evaluated in respect of nutritional status
Steptoe <i>et al.</i> (2004) <sup>(28)</sup>	Prospective	SF-36	271 adults patients Group 1 (behavioural counselling); n 136, M/W 54/82, mean age 43·3 (sp 13·8) years Group 2 (nutritional education counselling): n 135, M/W 51/84, mean age 43·2 (sp 14·0) years	12 months	Physical health status, mental health status and self-rated health all improved over the course of the study Few changes were observed in any of the QoL domains
Isenring <i>et al.</i> (2003) <sup>(29)</sup>	Prospective	EORTC QLQ-C-30	Sixty cancer patients M/W 51/9, age range 24–85 years, mean age 61.9 (sp 14.0) years	4 weeks	There was a significant correlation between PG-SGA score and global QoL A significant correlation was also observed between the change in PG-SGA score and change in global QoL after 4 weeks of radiotherapy
Tomíska <i>et al.</i> (2003) <sup>(30)</sup>	Clinical trial	EORTC QLQ-C-30 and VAS	Nineteen patients with far-advanced cancer suffering from anorexia and more than 5 % weight loss M/W 15/4, age range 44-78 years, mean	2 months	Significant benefit in appetite was found by VAS and QLQ-C30 questionnaire QoL was not evaluated in respect of nutritional status

age 59 years

Table 1. Continued		

Reference	Study design	QoL questionnaire	Patients included	Follow-up	Relationship between QoL and nutritional status in study results
Tidermark (2003) <sup>(31)</sup>	Clinical trial	EuroQoL-5D, SF-36 and Nottingham Health Profile	Patients with femoral neck fracture Studies I and II: $n$ 90, $>$ 65 years, M/W 24/66, mean age 89 (sp 7) years Studies III and IV; $n$ 110, $\ge$ 70 years, M/W 13/87, mean age 80 (sp 6) years Studies V and VI: $n$ 24, $\ge$ 70 years, M/W 0/24, mean age 83 (sp 5) years	Study I: 12 months Study II: minimum 24 months Studies III and IV: 24 months Study V: in hospital Study VI: 12 months	QoL was not evaluated in respect of nutritional status
Ohtsuka <i>et al.</i> (2002) <sup>(32)</sup>	Prospective	Kurihara (modified)	Thirty-one Japanese patients who underwent pylorus-preserving pancreatoduodenectomy M/W 15/16, age range 39–85 years, mean age 62 years Group 1 (Imanaga): <i>n</i> 18, M/W 11/7, mean age 62·2 (sp 2·7) years Group 2 (Traverso): <i>n</i> 13, M/W 4/9, mean age 60·8 (sp 2·7) years	1 year after surgery	Prospective QoL and nutritional status were not different between Imanaga or Traverso reconstructions  The postoperative change was similar in the objective nutritional factors and physical QoL scores
Ribaudo <i>et al.</i> (2000) <sup>(33)</sup>	Clinical trial	FAACT and BACRI	213 patients Group 1: <i>n</i> 155 cancer patients, M/W 87/68, mean age 64-1 (sp 12-3) years Group 2: <i>n</i> 58, HIV infected, M/W 57/1, mean age 39-2 (sp 8-7) years	12 weeks	QoL was not evaluated in respect of nutritional status
Laws <i>et al.</i> (2000) <sup>(34)</sup>	Cross-sectional	Not validated	64 patients Group 1 (well nourished): n 41, M/W 21/20, age range 40–85 years, mean age 64-5 years Group 2 (malnourished): n 15, M/W 6/9, age range 37–73 years, mean age 61-8 years Group 3 (severely malnourished): n 8, M/W 4/4, age range 66–84 years, mean age 73-1 years	In haemodialysis for less than 1 month	Malnutrition is associated with poorer QoL when the degree of malnutrition becomes severe
Van Bokhorst-de Van der Schuer et al. (2000) <sup>(35)</sup>	Clinical trial	EORTC QLQ-C-30 and COOP-WONCA	Forty-nine malnourished (weight loss > 10 %) head and neck cancer patients (thirty-one patients filled in both questionnaires)  Group 1 (no preoperative nutritional support): n 11, M/W 7/4, mean age 56·6 years, age range 42–76 years  Group 2 (standard enteral nutrition) n 10, M/W 3/7, mean age 58·6 years, age range 43–69 years  Group 3 (isonitrogenous enteral nutrition): n 10, M/W 6/4, mean age 61·4 years, age range 43–83 years	6 months after surgery	Enteral nutrition improves QoL of severely malnourished head and neck cancer patients in the period preceding surgery  No benefit of preoperative enteral feeding on QoL could be demonstrated 6 months after surgery
Callahan <i>et al.</i> (2000) <sup>(36)</sup>	Prospective	Quality of Well Being Scale	150 patients receiving PEG M/W 66/84, age range 60–98 years, mean age 78-9 (sp 8-1) years	Over 14-month period	QoL was not evaluated in respect of nutritional status
Bruera <i>et al.</i> (1998) <sup>(37)</sup>	Clinical trial	FLIC and VAS	Patients with advanced cancer M/W 47/37, mean age 62 (sp 11) years Group 1 (Megestrol group): <i>n</i> 62 Group 2 (placebo): <i>n</i> 60	21 d	No significant difference was observed in any of the values assessed before and after Megestrol or placebo  QoL was not evaluated in respect of nutritional status

Fable 1. Continued

Reference	Study design	QoL questionnaire	Patients included	Follow-up	Relationship between QoL and nutritional status in study results
Beller <i>et al.</i> (1997) <sup>(38)</sup>	Olinical trial	LASA (using six linear analogue self-assessment scales)	240 cancer patients over 18 years Group 1 (placebo): n 79, MW 54/25 Group 2 (low dose): n 80, MW 52/28 Group 3 (low dose): n 81, MW 53/28	12 weeks	Improvement of appetite, mood and overall QoL in these patients, although not through a direct effect on nutritional status
McLeod <i>et al</i> (1995) <sup>(39)</sup>	Cross-sectional	Time Trade-off Technique Direct Questioning of Objectives Gastrointestinal Quality of Life Sickness Impact Profile Physician Global Assessment Visick scale	Group 1 (Whipple group): n 50, MW 13/12, mean age 57.7 (so 14.6) years Group 2 (control group): n 50, MW 13/12, mean age 58.0 (so 11.1) years		QoL and nutritional status are excellent in patients following a Whipple procedure

ity of Life – Head and Neck; SF-36, Short Form-36 Health Survey; HRQoL, health-related quality of life, WHOQOL-BREF; WHO Quality of Life-BREF; QoL-OS, Quality of Life focused on symptoms of oxidative stress; EuroQoL-SD, Euro Quality of Life 5 Dimensions; MNA Mini Nutritional Assessment; OHIP-EDENT, Oral Health Impact Profile – EDENT; PEG, percutaneous endoscopic gastrostomy; SCREEN, Seniors in the Community. Risk Evaluation for European Organisation for Research and Treatment of Cancer Qual-Care Cooperative Information Project World Organization of National Colleges, Academies, and Academic Associations of General Practi EORTC QLQ-H&N35, Quality of Life; European Organisation for Research and Treatment of Cancer QoL, quality of life; M/W, men

Among the reviewed papers, ten  $(35.71\,\%)$  found no type of relationship between nutritional status, or any type of nutrition, and HRQoL $^{(12,16-18,20,22,24,27,31,33)}$ .

Nine articles (32·14%) recommended, or considered necessary, future prospective studies in order to completely clarify the correlation between HRQoL and nutritional status<sup>(14,15,19,20,23,26,27,33,34)</sup>.

It is important to emphasise the study of Ravasco *et al.*  $^{(21)}$ , where the existence of a linear association (P < 0.05) between an increase in HRQoL and an improvement in nutritional status was demonstrated. The research of Isenring *et al.*  $^{(29)}$  determined that 26% (P < 0.001) of the appreciated variation in HRQoL is explained by changes observed in nutritional status measured with the 'Patient-Generated Subjective Global Assessment' (PG-SGA). By means of multivariate analysis Keller ( $^{(23)}$ ) showed that the association between nutritional risk and HRQoL is consistent, explaining the 44% variation Hickson & Frost ( $^{(25)}$ ) describe, concluding with the necessity for a tool that shows high sensibility to alterations of HRQoL and their relationship with nutritional status.

### Questionnaire description and use

The questionnaires that were used in more than one article are: the European Organisation for Research and Treatment of Cancer Quality of Life questionnaire (EORTC-QLQ-C-30) and SF-36 on six occasions each; the Euro Quality of Life 5 Dimensions (EuroQoL-5D) on three occasions; linear analogue scale assessment (LASA) or visual analogue scales (VAS) on three occasions. In two studies, non-validated instruments were used to evaluate quality of life. The rest of the questionnaires were only used once.

It was observed that in one article (3.57%) six different questionnaires were used to measure quality of life<sup>(39)</sup>, in another (3.57%) three questionnaires<sup>(31)</sup>, in five articles (17.86%) two were used<sup>(18,22,30,35,37)</sup> and in the rest only one.

Most of the questionnaires described in the studies measured quality of life in a generic way (SF-36; EuroQoL-5D; Dartmouth Primary Care Cooperative Information Project-World Organization of National Colleges, Academies, and Academic Associations of General Practitioners/Family Physicians (COOP-WONCA); LASA or VAS; Nottingham Health Profile; Physician Global Assessment (MD global); Quality of Well Being Scale; Sickness Impact Profile; Time Trade-off Technique; Vailas; WHO Quality of Life-BREF (WHOQOL-BREF)). Several were specific for cancer (EORTC QLQ-C-30; EORTC Head and Neck questionnaire (QLQ-H&N35); Functional Living Index-Cancer; Kurihara; Quality of Life focused on symptoms of oxidative stress) or for gastrointestinal pathology (Subjective Assessment of Quality of Life (Transplantation); Troidl; Visick scale).

Only three questionnaires that can be related to quality of life could be retrieved, two of them specific for anorexia and cachexia (Bristol-Myers Anorexia Cachexia Recovery Instrument; Functional Assessment of Anorexia/Cachexia Therapy) and one specific for patients with permanent home parenteral nutrition (Direct Questioning of Objectives) (Table 2).

### Discussion

In the documentary study the validity of the articles must be emphasised. The validity was confirmed both by the good

Table 2. Questionnaires used in reviewed articles

Questionnaire	Abbreviation	Times used	Design	Domains
European Organisation for Research and Treatment of Cancer Quality of Life	EORTC QLQ-C-30	6	Cancer	Global quality of life scale  Five functional scales: physical, role, emotional, cognitive, social  Nine symptom scales: fatique, pain, nausea/vomiting, dyspnoea, insomnia, appetite loss,
				constipation, diarrhoea, financial difficulties
Short Form-36 Health Survey	SF-36	6	Generic	Thirty-six items organised in eight domains: physical functioning, role limitations caused by physical health problems, bodily pain, general health perceptions, vitality, social functioning, role limitations caused by emotional problems, mental health
Euro Quality of Life 5 Dimensions	EuroQoL-5D	3	Generic	Visual analogue scale (quality of life perceived) Change of health perceived Five dimensions: mobility, self-care, usual activity, pain or discomfort, anxiety or depression
Linear analogue scale assessment or visual analogue scale	LASA or VAS	3	Generic	Lines of a standard length (usually 10 cm), with the extremes of a variable. The patient marks the point that corresponds to their perceived status. Sometimes status is images
Non-validated questionnaire		2	Generic	Author design
Bristol-Myers Anorexia Cachexia Recovery Instrument	BACRI	1	Anorexia and cachexia	An eight-item questionnaire: BACRI 7 (seven items), subjective recovery from symptoms of anorexia or cachexia; BACRI 1 (one item), patient perception of benefit
Dartmouth Primary Care Cooperative Information Project World Organization of National Colleges, Academies, and Academic Associations of General Practitioners/Family Physicians	COOP-WONCA	1	Generic	Six dimensions: physical fitness, mental health, daily activities, social activities, change in health, overall health
Direct Questioning of Objectives	DQO	1	Home parenteral nutrition	A category scale used for three life objectives: working full time, enjoying recreation, travel
European Organisation for Research and Treatment of Cancer Quality of Life – Head and Neck	EORTC QLQ-H&N35	1	Head and neck cancer	Designed to be used together with EORTC QLQ-C-30 Six symptom scales: pain, swallowing, senses (taste/smell), speech, social eating, social contacts Seven single items: sexuality, teeth problems, problems opening mouth, dry mouth, sticky saliva, cough, feeling ill
Functional Assessment of Anorexia/Cachexia Therapy	FAACT	1	Anorexia and cachexia	Four subscales: physical wellbeing, social/family wellbeing, emotional wellbeing, functional wellbeing Also twelve items (additional concerns)
Functional Living Index-Cancer	FLIC	1	Cancer	Five domains (linear analogue scale): physical wellbeing and ability, emotional state, sociability, family situation, nausea
Gastrointestinal Quality of Life Index	GiQLI	1	Gastrointestinal	Five domains: symptoms, physical dysfunction, emotional dysfunction, social dysfunction, effects of the medical treatment carried out
Kurihara questionnaire	Kurihara	1	Cancer	Twenty-three items categorised into two domains: physical and psychosocial aspects
Nottingham Health Profile	NHP	1	Generic	Part 1: energy level, pain, emotional reaction, sleep disturbance, social isolation, physical abilities Part 2: seven items about life affected areas
Oral Health Impact Profile - EDENT	OHIP-EDENT	1	Bucodental health	Nineteen items grouped into seven domains: functional limitation, pain, psychological discomfort, physical disability, psychological disability, social disability, handicap
Physician Global Assessment	MD global	1	Generic	Assessment of wellbeing using 10 cm visual analogue scale
Quality of Life focused on symptoms of oxidative stress	QoL-OS	1	Cancer	Five subscales: functional, physical, emotional, social and family, fatigue
Quality of Well Being Scale	QWB	1	Generic	Three scales of functioning with a measure of symptoms and problems. The model separates aspects of health status and life quality into distinct components. These are life expectancy (mortality), functioning and symptoms (morbidity), preference for observed functional states (utility) and duration of stay in health states (prognosis)
Sickness Impact Profile	SIP	1	Generic	Two overall domains (physical and psychosocial). Twelve categories (sleep and rest, eating, work, home management, recreation and pastimes, ambulation, mobility, body care and movement, social interaction, alertness behaviour, emotional behaviour, communication)

Questionnaire	Abbreviation	Times	Design	Domains
Subjective assessment of Quality of Life (transplantation)		-	Liver or small-bowel transplant	Twenty-six domains: anxiety, depression, alcohol use, drug use, cognitive emotional style, mental status, physical mobility, appearance, pain and discomfort, stress experience, coping, financial, parenting, marital relationship, sexuality, digestive, urinary, sleep, energy, optimism, control impulsiveness, medical satisfaction, quality of relationship, quality of
Time Trade-off Technique	011	-	Generic	social relations, leisure recreation, vocational  This assumes an individual's reference position is to have an initial endowment of time t and to be in 'less than full health' and gets the respondent to compare this to a shorter period in a state of higher quality of life.
Troid		-	After gastrectomy	Dimensions: health state, mobility, self-care, usual activities, pain/discomfort, anxiety/depression Fourteen items divided into two groups: specific disease symptoms (eight items) and
Vailas <i>et al.</i> Visick scale			Generic After gastrectomy	Social-personal companits (six items) Six domains: global quality of life perceived, health, loneliness, food enjoyment, food security, depression Four grades: Visiok ( excellent), no gastrointestinal complaints; Visiok II (good), mild gastrointestinal
WHO Quality of Life-BREF	WHOQOL-BREF	-	Generic	companies, vision in (rail,) significant gastrontesurial companies, vision in (poor), rationed Perception of the quality of life Health status Four domains: physical, psychological, social relationship, environment

results given by measuring the obsolescence, and by the excellent result of the Price index. The excellence and the current importance of the research articles are complementary, but nevertheless important factors, in those studies referring to the health sciences.

It must be underlined that the evaluation of HRQoL is circumscribed specifically within the scope of the investigation. Its use in common medical practice would help to obtain validated information about the impact of the illness or the treatment of the patient in daily life, both of which could be useful in decision making<sup>(40)</sup>. Knowing HRQoL does not substitute the symptomatic, analytic and morphological evaluations, but complements them, introducing something as important as the patient's point of view about their perception of their own health<sup>(41)</sup>. Quality of life assessment measuring the patient's experiences of the impact of disease and therapy, expectations and satisfaction should be the 'gold standard' as an independent end point in clinical trials<sup>(21,42)</sup>.

The undertaking of prospective studies of HRQoL in clinics improves the information about the patient, which, along with the diagnosis, provides important information about the patient's perception of the effect of treatment<sup>(13,43)</sup>. Neither must it be forgotten that the objective is also to prioritise resources. Mathematical methods are applied to try to quantify the quality of life in relation to its usefulness (quality-adjusted life years) as a self-profit concept<sup>(44)</sup>.

The review proves that HRQoL has been studied as a variable in the health-illness process<sup>(45,46)</sup>, and not only, as is frequently the case, as a covariable in pathological<sup>(14,47-51)</sup> or surgical<sup>(52-54)</sup> process studies, in pharmacological follow-ups<sup>(55)</sup>, in relation to somatic<sup>(56,57)</sup> or social<sup>(58,59)</sup> aspects, or to support future recommendations<sup>(60-62)</sup>.

### Limitations to the identified studies

The present review exposes the lack of homogeneity of the studies found, produced by certain limitations, namely the different questionnaires used, the diversity of pathologies, sample sizes, methodology and variation in follow-up, all of which do not permit meaningful meta-analysis, thus making direct comparison awkward, especially those studies that apply non-validated HRQoL questionnaires. Furthermore, two studies are retrospective in design and are susceptible to bias.

It is fundamental when designing these studies that possibly confusing variables are controlled, that interaction effects are recognised and that HRQoL is evaluated at different points in the evolution of the illness, the period in which patients are having the treatment or that these are matched up with a control group<sup>(63)</sup>.

### Limitations due to the questionnaires

No mention of the patient's acceptance of the HRQoL questionnaires used has been found in the studies reviewed. The complexity of these tools or their use can be the cause of disinterestedness, partial fulfilment or desertion on the part of the participants. This conformity is a crucial methodological requisite for avoiding skewed results<sup>(64)</sup>. It is possible that the structure and appearance of the questionnaires about HRQoL are considered less important than the final results, but if this circumstance is not properly managed, it will never be known

if the results are influenced by the tool's design. On the other hand, it is convenient to limit the number of questionnaires used; some studies recommend not using more than three, if possible, or up to five in extreme cases<sup>(65)</sup>.

The use of validated and reliable measurements of HRQoL is essential. Ideally, any generic measurement of HRQoL should be replaced with a specific measurement that reflects the sensibility to the changes produced by the illness or by the influences related to the treatment. These questionnaires should not only have to be sensitive to the changes produced in the desired variable, but should also be acceptable to the patients<sup>(64)</sup>.

### Quality of life and nutritional status

The relationship existing between nutritional status and HRQoL is becoming an important question not only in the study of oncological patients<sup>(66,67)</sup>, but also in other pathologies<sup>(68)</sup> and interventions<sup>(69,70)</sup>. The improvement of this correlation, as a consequence of an appropriate nutritional intervention, enables the reduction of the number of post-surgical complications<sup>(70,71)</sup>, shortens the recovery time and the length of hospital stay, improves tolerance to the treatment<sup>(72,73)</sup> and even increases the rate of survival<sup>(74–77)</sup>, and with it a general decrease in morbidity<sup>(78,79)</sup>.

On the other hand, as has been seen in the reviewed studies, the advice and nutritional follow-up given by professionals is related directly to the improvement in nutritional status, which will be related to the improvement in HRQoL<sup>(25,80)</sup>. It has been demonstrated, in head and neck neoplasm, that nutritional advice enables improvements in quality of life greater than those obtained by nutritional supplementation without advice<sup>(81)</sup>.

Now, the efficacy of nutritional advice as a positive influence on HRQoL depends on the possibility of adapting intervention to the specific need of each type of patient. Therefore, nutritional advice should be given by dedicated, specialised groups<sup>(75,82)</sup>. Of special importance is the need for future studies that clarify the relationship between nutritional status and quality of life. This importance is recognised by studies included in the present review<sup>(14,15,19,20,23,26,27,33,34)</sup> and also in other publications that highlight the need to explore the relationship in greater detail.

Hence, the measurement of HRQoL with generic tools requires large sample sizes in order to demonstrate statistically significant differences and, in the majority of cases, these types of questionnaires are affected by uncontrolled external factors (42,83–86). Ultimately, valid HRQoL measurement tools are dependent upon patient perception, the impact of the illness, the treatment, expectations and wellbeing. There should be an independent gold standard for all research projects and everyday medical practice.

A specific tool is needed: one that is sensitive to the measurement of HRQoL and can be self-administered quickly and easily on a regular basis. Nevertheless, it must be recognised that the development of a tool to detect, evaluate and monitor the influence of the pathological base is not an easy task.

### Conclusion

Only three studies<sup>(19,23,29)</sup> selectively focused on the relationship between nutritional status and quality of life, this

evaluation being performed not by means of specific questionnaires but by statistical analysis of data obtained via validated questionnaires.

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