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Illness Perceptions, Psychological Distress and Coping Strategies as Predictors of Quality of Life After Radiotherapy in Head and Neck Cancer Patients



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ABSTRACT

Objective: To investigate the predictive value of certain psychological factors on the quality of life (QoL) of patients with head and neck cancer (HNC) after radiotherapy.

Methods: Depressive and anxiety symptoms (HADS), illness perceptions (IPQ-R), coping strategies (Brief COPE) and QoL (EORTC-QLQ-C30 and EORTC-QLQ-H&N35) were assessed in fifty patients before radiotherapy (T1) and 3 months afterwards (T2).

Results: Hierarchical regression analysis revealed that depression (HADS-D) at T1 was associated with impairment in Physical, Role, and Social Functioning (P = 0.025, P = 0.019 and P = 0.019, respectively) at T2. Additionally, Behavioral Disengagement (Brief COPE) at T1 was the coping strategy most closely associated with lower scores on Physical (P = 0.013), Emotional (P = 0.003), Cognitive (P = 0.042) and Social Functioning (P = 0.040). Also, regarding IPQ-R dimensions, stronger beliefs about Emotional Representations and weaker beliefs about Illness Coherence at T1 were associated with higher H&N35 sum score at T2 (P = 0.024 and P = 0.032, respectively). Finally, stronger Perceived Control (IPQ-R) at T1 was associated with higher Social Functioning at T2 (P = 0.011). **Conclusion:** Depressive symptoms are predictors of impaired QoL in patients with HNC after radiotherapy. Additionally, Behavioral Disengagement, as avoidance coping mechanism, and strong beliefs that HNC will affect their emotional well-being aggravate certain domains of the QoL, while belief that their illness is amenable to treatment is associated with better QoL following radiotherapy. Psychological intervention aimed at improving the QoL of patients with HNC undergoing radiotherapy should be based on identification of depression in these patients, and their coping strategies and illness perceptions.

INTRODUCTION

Head and neck cancer (HNC) is a heterogeneous group of malignancies that accounts for 5% of malignant diseases. It shows a higher prevalence in men, being the fourth most frequent cancer in the male population; its incidence increases with age (>60 years) and is associated with abuse of tobacco and alcohol. Radiotherapy (RT) is the main form of treatment, either alone or combined with surgery and/or chemotherapy, and is reported to promote the quality of life (QoL) of patients by alleviating their symptoms. The adverse effects of HNC and its treatment are related to vital functions, and include difficulties in swallowing, breathing and speech, but are also associated with such emotional dimensions as negative self-image, psychological distress, and social isolation. Many patients experience increased levels of psychological distress as they face stressful medical interventions, repetitive and different types of treatments and side effects, and go through the different stages of the disease. As a result, they often reflect a type of "burnout", i.e., a psychological deterioration as a consequence of chronic stress [1]. These effects force individuals with HNC to adapt to their new situation at the social and existential level, searching for coping strategies to maximize the adaptation to the disease and a meaning in their disease experience [2].

Individuals process the situation and, finally, adapt to their illness in various different ways, irrespective of their clinical and socio-demographic characteristics. In addition, their coping strategies are determined by a variety of factors, such as personality, religiousness and received support. According to the Common Sense Model [3], in order to adapt to a new situation the individual engages in a dynamic process of reconstruction of ideas, feelings and behavior, which can influence various dimensions of their QoL [4].

Factors that are reported to influence the QoL of patients with HNC include clinical and sociodemographic characteristics [5], but also a variety of psychological factors, including depression, anxiety, alcohol abuse, illness perceptions and coping strategies [4,6]. Evaluation of factors affecting QoL in patients with HNC continues to be of interest, as a recent 10-year follow-up study documented that QoL in the first year after diagnosis can predict survival [7,8].

Most studies investigating the association between illness perceptions, emotional distress, coping strategies and QoL in head and neck cancer patients have been cross-sectional, based on a single assessment, either immediately after diagnosis [4,9] or at follow-up [10]. As pointed out by a recent systematic review [1], the directionality of the correlations between psychological distress and coping strategies in head and neck cancer patients is not easily understood. Due to the insufficient data, it is necessary to carry out prospective design studies which will focus, mainly, on the definition of temporal patterns of psychosocial functioning that may affect the associations between the aforementioned variables and the quality of life in head and neck cancer patients. Additionally, among studies there are conflicting findings about the association between active coping style aimed at direct change and adjustment to the illness and mental distress experienced by patients with head and neck cancer [1,11,12]. The findings that appeared to be most consistent are those suggesting that avoidance and disengagement mechanisms are associated with increased mental distress in patients with head and neck cancer. To the best of our knowledge there are few prospective studies that have assessed the psychological distress, illness perceptions and coping strategies and their impact on quality of life in patients with head and neck cancer and even fewer in patients with head and neck cancer undergoing radiotherapy [13]. For all the aforementioned reasons the aim of the present study was to prospectively assess the psychological distress, illness perceptions and coping strategies adopted by patients with HNC undergoing radiotherapy at two time points, in a pre- and a post radiation period and to investigate their predictive value on QoL in a short-term after RT.

MATERIAL AND METHODS

Participants

The sample consisted of the first 50 consecutive patients diagnosed with primary HNC who started radiotherapy during the study period and agreed to participate. All patients were recruited from the same Radiation Oncology Department of a University Hospital. Exclusion criteria were age >75 years, metastatic disease, co-administration of chemotherapy, a diagnosis of dementia, abuse of substances, mental illness, previous participation in psycho-educational intervention, and inability to complete the questionnaires for any reason. From the patients who fulfilled the criteria and were invited to participate (N = 60), 6 patients were excluded due to the missing data, 4 declined to participate and 50 agreed (response rate 83.3%).

Procedure and Study Instruments

The study was conducted in accordance with the Ethical Standards as formulated in the World Medical Association Helsinki Declaration (2002) and its later amendments, and an IRB statement that IRB approval was unnecessary, was received. The participants were informed about the study objectives and data confidentiality and signed a consent form according to the aforementioned principles.

Assessment was made at two time points, the first prior to the simulation appointment and the onset of radiotherapy (T1) and the second at the first follow-up, 3 months after completion of radiotherapy (T2). The choice of the follow-up point time at 3 months after the completion of radiotherapy was based on the following elements: a. it was the first scheduled follow-up visit at the specific Radiation Oncology Department and b. it was considered as a sufficient period for the remission of the acute adverse effects of RT and, consequently, for the reduction of their effects on the quality of life of the patients. All the medical information was collected by one radiation oncologist and all the psychological assessments were conducted by one clinical psychologist. The study was prospective, questionnaire based, with repeated measures, and the patients were asked to complete the following research tools.

The Hospital Anxiety and Depression Scale (HADS)

HADS consists of 14 items and aims to measure symptoms of anxiety (HADS-A) and depression (HADS-D) [14]. The answers are given on a Likert scale of 0-3 and the score of each of the two sub-scales ranges from 0 to 21, with the higher value corresponding to more frequent and severe symptoms. Value categorization is as follows: 0-7: normal, 8-10: mild, 11-14: moderate, 15-21: severe. The questionnaire has been translated and validated in the Greek language [15].

The Revised Illness Perception Questionnaire (IPQ-R)

IPQ-R consists of 38 items and provides a quantitative assessment of the nature and strength of perceptions in nine dimensions [16]: illness identity (perception of symptoms related to the illness), acute/chronic timeline (perceived duration of the illness), timeline cyclical, consequences, personal control (self-efficacy beliefs), treatment control (treatment expectations), illness coherence (the degree of overall coherent understanding of the illness), emotional representations (symptoms of anxiety and depression experienced as result of the illness) and causal attributions of the illness: emotional (e.g., stress), behavioural (e.g., smoking) and external causes (e.g., chance) and risk factors (e.g., immune system). Responses are given on a 5-point Likert scale (ranging from "Absolutely disagree" to "Absolutely agree"). Higher values are indicative of a stronger perception. The item on the perceived cause of the illness is open-ended, requesting the participants to rank the three most important causal factors. The version used was the form of the questionnaire validated in the Greek language [17].

The Brief COPE Inventory

The Brief COPE Inventory for assessing coping strategies is the modified version of the original COPE Inventory [18], and it has been validated in Greek [19]. It includes 28 items corresponding to 14 scales: self-distraction, active coping, denial, substance use, use of emotional support, instrumental support, behavioural disengagement, venting, positive reframing, planning, humour, acceptance, religion, self-blame. The instructions were modified in order for participants to report what they did when they specifically experienced their disease (cancer) or treatment side-effects. The answers range from 1 "I do not do it at all", to 4 "I do it a lot". Higher scores correspond to more frequent use of each strategy.

EORTC-QLQ-C30 & EORTC-QLQ-H&N35

The European Organization for Research and Treatment of Cancer Quality of Life Questionnaire (EORTC-QLQ-C30) and the disease-specific questionnaire of EORTC for HNC (EORTC-QLQ-H&N35) were also administered [20,21]. The Greek versions of both questionnaires have been provided by the official website of EORTC study tools (https://gol.eortc.org). The EORTC-QLQ-C30 consists of 30 items classified into 5 domains: Physical, Role, Emotional, Cognitive and Social functioning, and the Global Health Status (GHS), while there are certain single items of symptoms. EORTC-QLQ-H&N35 consists of 35 items that correspond to seven domains: pain, swallowing, senses, speech, social eating, social contact, and sexuality. In addition, 11 independent items refer to specific physical symptoms (e.g., teeth problems, dry mouth). In both questionnaires, responses are given on Likert 4 and 7-point scales, and the scores ranged from 0 to 100. On the symptom items, the highest score corresponds to a stronger intensity, while on the functional scales, the highest score corresponds to a better quality of life. The symptoms sum score provides a generalizing score of QoL (H&N sum score).

Statistical Analysis

Mean and standard deviation (SD) were used to describe the scores on the scales used in the study. Frequencies and percentages were used for categorical variables such as demographic data and stage of cancer. A general linear model of repeated measures was applied to detect statistically significant differences between the scores at the two study periods, T1 and T2 on the various scales of EORTC-QLQ-C30, H&N35, HADS, IPQ-R and Brief COPE. In order to investigate predictive factors of QoL, the effects of the HADS, IPQ-R and Brief COPE scores at T1 and the demographic and clinical characteristics were examined in each regression model hierarchically, after examining the univariate effect of each one separately using correlation analyses or mean comparison statistics. Specifically, a blockwise forward method was adopted, entering the dimensions of the IPQ-R, HADS and Brief COPE in this specific order. The model with statistically significant outcomes was then tested for changes after controlling for the effect of demographic characteristics such as age, gender, and educational level, and for smoking status, alcohol consumption, disease stage, previous chemotherapy, and surgery. The forward blockwise model was adopted separately for each of the domains of the EORTC-QLQ-C30 and the H&N35 sum score in order to examine the outcomes in a hierarchical manner in terms of interest, but also to assure the validity, as the small number of participants would lead to unreliable models if the simultaneous enter approach had been chosen. Statistical significance was set at 0.05 for all tests. Analysis was conducted with the use of SPSS v22.0.

Variable	Value
Age (Mean ± SD) (Range)	62.9 ± 8.3 (38-74)
Gender (N, %)	
Female	7 (14)
Male	43 (86)
Family status (N, %)	
Single	3 (6)
Married/domestic partnership	39 (78)
Divorced/separated	3 (6)
Widowed	5 (10)
Educational level (N, %)	
Primary school	31 (62)
High school	11 (22)
University degree	8 (16)
Disease duration (Months, Mean ± SD)	13.42 ± 7.74
AJCC Stage of cancer (N, %)	
1	13 (26)
II	11 (22)
III	11 (22)
IV	15 (30)
Cancer site (N, %)	
Buccal mucosa	3 (6)
Maxillary	6 (12)
Lingual	3 (6)
Mouth floor	7 (14)
Oropharynx	9 (18)
Larynx	20 (40)
Salivary Gland	2 (4)
Surgery (N, %)	
Yes	29 (58)
No	21 (42)
Chemotherapy (N, %)	
Yes	23 (46)
No	27 (54)
Type of radiotherapy (N, %)	
Postoperative	21 (42)
Radical	29 (58)
Smoking (N, %)	
Yes	37 (74)
No	13 (26)
No. of cigarettes per day	32.2 ± 20.9
Years of smoking (Mean±SD) (Range)	25.59 ± 14.20 (1-55)
Alcohol consumption (N, %)	
Yes	35 (70)
No	15 (30)

 Table 1. Sociodemographic and Clinical Characteristics of the Study Patients With

Table 2. Scores on Quality of Life and Anxiety and Depression Questionnaires of Patients With Head and Neck Cancer (N = 50), Before (T1) and 3 Months After (T2) Radiotherapy

Questionnaire	T1 (Mean ± SD)	T2 (Mean ± SD)
EORTC-QLQ-C30		
Global Health Status (GHS)	54.42±25.06	57.48±24.13
Physical Functioning	76.27 ±17.72	76.87±23.96
Role Functioning	68.00±33.64	68.03±29.82
Emotional Functioning	69.67±25.52	73.30±23.45
Cognitive Functioning	83.67±22.21	86.39±20.03
Social Functioning	80.00±26.94	81.29±22.21
Fatigue	32.00±26.24	32.88±25.05
Nausea/vomiting	5.67±20.37	8.50±16.00
Pain	17.00±20.62	21.77±23.13
Dyspnoea	32.67±31.22	30.61±33.22
Insomnia	26.67±32.99	21.77±26.83
Appetite loss	15.33±24.48	25.17±27.66
Constipation	15.33±27.11	10.20±15.52
Diarrhoea	6.00±12.94	2.04±8.07
Financial problems	46.00 ±33.6	42.86± 34.69
EORTC QLQ H&N35		
Pain	19.67±19.33	22.28±21.41
Swallowing	18.39±22.01	22.45±23.95
Senses	27.33±32.60	26.53±20.39
Speech	30.89±32.16	24.04±26.09
Social eating	17.17±22.74	24.83±23.90
Social contact	15.07±23.20	16.73±23.46
Sexuality	40.48±39.09	42.01±35.89
Teeth	24.67±36.15	25.17±25.94
Opening mouth	20.00±27.77	17.69±26.44
Dry mouth	26.67±26.08	41.5±28.49*
Sticky saliva	27.33±31.36	38.78±30.69
Coughing	22.00±27.45	17.01±26.46
Feeling ill	18.00±24.48	27.21±28.60
H&N 35 Sum Score	22.52 ±13.85	24.45 ±14.46
HADS-A	6.44 ± 5.29	5.44 ± 4.28
HADS-D	6.56 ± 4.85	5.62 ± 4.29

EORTC QLQ-C30, European Organization for Research and Treatment of Cancer Quality of Life Questionnaire; EORTC QLQ H&N35, disease-specific questionnaire for head and neck cancer; HADS: Hospital Anxiety and Depression Scale (A: anxiety, D: depression); SD, standard deviation. *P <0.05

AJCC, American Joint Committee on Cancer; SD, standard deviation.

 Table 3. Scores on the Questionnaires IPQ-R and Brief Cope of Patients With Head

 and Neck Cancer (N = 50) Before (T1) and 3 Months After (T2) Radiotherapy

Questionnaire	T1 (Mean ± SD)	T2 (Mean ± SD)
IPQ-R		
Identity	10.21±1.74	11.03±1.16
Timeline acute/chronic	20.80 ± 2.44	21.13 ± 2.76
Timeline cyclical	13.61 ± 2.03	14.10 ± 2.14
Consequences	20.49 ± 2.50	19.78 ±1.96
Personal control	18.67 ± 2.38	18.57 ± 2.18
Treatment control	16.29 ± 2.41	15.80 ± 2.29
Illness coherence	14.02 ± 2.21	13.84 ± 1.66
Emotional representations	18.76 ± 1.86	18.92 ± 1.81
Causes		
Emotional cause	2.78 ± 0.79	2.73 ± 0.69
Behavioural cause	3.00 ± 0.69	3.45 ± 0.45***
Risk factors	3.08 ± 0.67	3.49 ± 0.52**
External	3.24 ± 1.18	2.90 ± 1.31
Brief COPE		
Self-distraction	2.21 ±1.04	2.04 ± 1.36
Active coping	2.30 ± 0.88	2.51 ± 1.08
Denial	1.97 ± 0.97	2.17 ± 1.03
Substance use	1.32 ± 0.71	1.36 ± 0.70
Emotional support	2.32 ±1.00	2.36 ±0.96
Instrumental support	2.15 ± 1.03	2.25 ± 1.11
Behavioural disengagement	1.55 ± 0.85	1.58 ± 0.88
Venting	2.15± 0.92	2.09± 0.99
Positive reframing	2.42 ± 0.83	2.23 ± 0.91
Planning	2.48 ± 0.93	2.49 ± 0.91
Humour	1.64 ±0.90	1.62 ±0.95
Acceptance	2.86 ± 0.96	2.88 ± 1.00
Religion	2.05 ± 0.99	2.08 ± 0.94

IPQ-R, Revised Illness Perception Questionnaire; SD, standard deviation. **P <0.05: ***P <0.001

RESULTS

Characteristics of the Sample

The characteristics of the study sample are shown in Table 1. The mean age of the participants was 62.9 ± 8.3 years (range 38-74), with the majority being of men (86%), married (78%), of low educational level (62% were primary school graduates), smokers (74%) and alcohol consumers (70%). Regarding the clinical features, 58% of the sample had undergone surgery and the majority underwent radical radiotherapy (58%) (Table1).

EORTC-QLQ-C30 and H&N35

On comparison of the scores on EORTC-QLQ-C30 and H&N35 at T1 and T2 no significant difference was found in either the main domains or in the symptoms, apart from the symptom of dry mouth, which was aggravated after radiotherapy (P = 0.015). The mean scores on some of the items (e.g., sticky saliva, a general sense of "feeling ill", and difficulties with social eating) were higher at T2 than T1, but not to a statistically significant degree (Table 2).

HADS

In general, low levels of anxiety and depression were elicited (HADS<8) at both time points, with no significant difference between the T1 and the T2 measurements (Table 2).

IPQ-R and Brief COPE

Regarding IPQ-R, the strongest beliefs at both T1 and T2 concerned the chronic timeline of the illness, its negative consequences and the emotional representations. Comparison of the mean scores on IPQ-R at T1 and T2 to explore possible changes in the patients' perceptions showed change only regarding the perceived causes of their illness. Specifically, at T2 there was a significant increase in the belief that the illness was due to suppression of the immune system (p<0.001) and to behavioural factors (e.g., smoking, alcohol) (p<0.001). Concerning the Brief COPE, no significant difference was demonstrated between the scores at T1 and T2 (Table 3).

Quality of Life Associations

No significant correlations were shown between the socio-demographic and clinical characteristics of the patients and the domains of the EORTC-QLQ-C30 and the H&N35. Significant correlations were observed between HADS subscales, Brief COPE and IPQ-R, as assessed at T1, and the domains of EO-RTC-QLQ-C30 and H&N sum score as assessed at T2, as shown in detail in Table 4.

The main aim of this study was to identify the variables that, when measured at T1, could predict QoL outcomes at T2. On hierarchical regression analysis, it became evident that depression at T1 was a significant predictor of Physical, Role and Social Functioning (EORTC-QLQ-C30). A higher HADS-D at T1 was associated with lower Physical (beta = -1.533, P = 0.025), Role (beta = -2.054, P = 0.019) and Social Functioning (beta = -1.419, P = 0.019) at T2. In addition, a higher score on Behavioural disengagement (Brief COPE) was associated with lower scores on Physical (beta = -9.572, P = 0.013), Emotional (beta = -11.341, P = 0.003), Cognitive (beta = -6.442, P = 0.042) and Social Functioning (beta = -6.791, P = 0.040). GHS at T2 was closely associated with beliefs about Treatment controllability (IPQ-R) (beta = 3.832, P = 0.007) and Substance use (Brief COPE) (beta = -11.523, P = 0.015) at T1. Two dimensions of the IPQ-R were found to be associated with H&N sum score at follow-up. Specifically, stronger Emotional representations and weaker beliefs about Illness coherence at T1 were associated with higher H&N35 sum score at T2 (beta = 2.401, P = 0.024 and beta = -1.922, P = 0.032, respectively). Finally, stronger perceived control (IPQ-R) at T1 was associated with higher Social functioning at T2 (beta = -2.931, P = 0.011) (Table 5). In analysis, all outcomes were adjusted for age, gender, education, smoking status, alcohol consumption, disease stage, surgery and previous chemotherapy.

DISCUSSION

The present study assessed psychological distress, illness perceptions, coping strategies and QoL in patients with H&N cancer before and after radiotherapy, and investigated the predictive role of psychological distress, illness perceptions and coping strategies on various dimensions of QoL three months after radiotherapy.

With regard to QoL, little change was detected between the measurements before and 3 months after radiotherapy, except in dry mouth, a symptom that was significantly worse after radiotherapy. A well-known side-effect of irradiation is mucosal dryness, along with muscle atrophy and erythema. Patients experience pain, swallowing disorders and voice problems soon after radiotherapy, with a resulting marked decrease in overall QoL [22]. In addition, it was found that certain symptoms were slightly more intense after radiotherapy, but a literature search to identify minimally important differences (MID) in QoL scores [23] revealed no reports on patients with HNC, and no further interpretation of the changes in mean scores, in terms of clinical meaningfulness, is possible.

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Table 4. Zero Order Correlations Between Scores on HADS, IPQ-R and Brief COPE Questionnaires Before Radiotherapy (T1) and EORTC-QLQ-C30 and H&N 35 (Sum Score), Three Months After Radiotherapy (T2) In Patients With Head and Neck Cancer (N = 50)

uestionnaire	GHS	PF	RF	EF	CD	SF	H&N 35
ADS							
HADS-A	-0.219	-0.174	-0.198	-0.354*	-0.176	-0.453**	0.346*
HADS-D	-0.365**	-0.443**	-0.335*	-0.347*	-0.328*	-0.496**	0.317*
'Q-R							
Identity	0.053	0.104	0.132	0.140	0.090	0.204	-0.119
Timeline (acute/chronic)	-0.206	-0.127	-0.02	-0.2	-0.342*	-0.04	0.151
Timeline cyclical	-0.194	0.003	-0.071	0.001	-0.14	-0079	0.19
Consequences	0.016	-0.113	-0.028	-0.298*	-0.134	-0.119	0.171
Personal control	0.126	0.098	0.086	0.268	0.153	0.446**	-0.303*
Treatment control	0.293*	0.246	0.245	0.141	0.34*	0.264	-0.218
Illness coherence	0.238	0.156	0.126	0.315*	0.21	0.326*	-0.347*
Emotional representations	-0.174	-0.167	-0.175	-0.323*	-0.193	-0.205	0.36*
rief COPE							
Self distraction	0.168	0.147	0.097	-0.068	-0.087	0.01	-0.014
Active coping	0.352*	0.364**	0.286*	0.15	0.075	0.093	-0.197
Denial	-0.255	-0.353*	-0.347*	-0.423**	-0.317*	-0.428**	0.403**
Substance use	-0.247	-0.196	-0.254	-0.421**	-0.058	-0.281	0.275
Emotional support	0.089	-0.101	0.026	-0.053	0.13	-0.032	0.167
Instrumental support	0.337*	0.085	0.173	-0.075	0.153	0.133	0.104
Behavioural disengagement	-0.223	-0.463**	-0.26	-0.473**	-0.291*	-0.439**	0.209
Venting	-0.014	-0.182	0.018	-0.415**	-0.126	-0.204	0.315*
Positive reframing	0.18	0.176	0.025	0.177	0.005	0.139	-0.038
Planning	0.208	0.208	0.081	0.038	-0.015	-0.052	0.026
Humour	-0.12	0.106	-0.071	-0.158	-0.053	-0.173	0.058
Acceptance	0.105	-0.052	-0.094	-0.173	-0.075	-0.094	0.19
Religion	-0.092	-0.041	-0.112	-0.233	-0.084	-0.058	0.251
Self blame	-0.073	-0.07	-0.182	-0.215	-0.237	-0.269	0.229

CF, Cognitive Functioning; EORTC QLQ-C30, European Organization for Research and Treatment of Cancer Quality of Life questionnaire; EF, Emotional Functioning; GHS, Global Health Status; HADS, Hospital Anxiety and Depression Scale (A: anxiety, D: depression); H&N35, sum score of EORTC QLQ H&N35 disease-specific questionnaire for head and neck cancer; IPQ-R, Revised Illness Perceptions Questionnaire; PF, Physical Functioning; RF, Role Functioning; SF, Social Functioning. *P <0.05; **P <0.01; ***P <0.001

Table 5. Statistically Significant Effects of Factors Measured Before Radiotherapy (T1)On Quality of Life According to EORTC QLQ-C30 and H&N 35 Three Months AfterRadiotherapy (T2) In Patients With Head and Neck Cancer (N = 50)

	Global health status		
R ² = 0.163	Beta	t	Significance
Treatment Control (IPQ-R)	3.832	2.804	0.007
Substance Use (Brief COPE)	-11.523	-2.533	0.015
	Physical functioning		
R ² = 0.266	Beta	t	Significance
HADS-D	-1.533	-2.232	0.025
Behavioural Disengagement (Brief COPE)	-9.572	-2.572	0.013
	Role functioning		
R ² = 0.093	Beta	t	Significance
HADS-D	-2.054	-2.434	0.019
	Emotional functioning		
R ² = 0.234	Beta	t	Significance
Behavioral Disengagement (Brief COPE)	-11.341	-3.185	0.003
	Cognitive functioning		
R ² = 0.157	Beta	Т	Significance
Treatment Control (IPQ-R)	2.726	2.478	0.017
Behavioural Disengagement (Brief COPE)	-6.442	-2.089	0.042
	Social functioning		
R ² = 0.370	Beta	Т	Significance
Personal Control (IPQ-R)	2.931	2.635	0.011
HADS-D	-1.419	-2.442	0.019
Behavioural Disengagement (Brief COPE)	-6.791	-2.116	0.040
	H & N 35 sum score		
R ² = 0.179	Beta	Т	Significance
Emotional Representations (IPQ-R)	2.401	2.333	0.024
Illness Coherence (IPQ-R)	-1.922	-2.216	0.032

Beta coefficient and P values for the quality-of-life outcomes after Hierarchical Regression Analysis. Results are adjusted for age, gender, educational level, smoking status, alcohol consumption, disease stage, surgery, and previous chemotherapy. Brief COPE, Brief COPE Inventory; EORTC QLQ–C30, European Organization for Research and Treatment of Cancer Quality of Life questionnaire; HADS, Hospital Anxiety and Depression Scale (A: anxiety, D: depression); H&N 35, EO-RTC disease specific questionnaire for head and neck cancer; IPQ-R, Revised Illness Perceptions Questionnaire.

Regarding psychological distress, the study findings are in line with previous reports [24,25] as low mean values in anxiety and depression levels (<8 on HADS-A and HADS-D) were found. Studying the association between HADS levels at T1 and QoL outcomes at T2, it became evident that HADS-D at T1 affected Physical, Role and Social Functioning at T2. Specifically, a higher HADS-D score before radiotherapy was associated with lower scores on Physical, Role and Social Functioning three months after radiotherapy. Depression has already been identified as an important predictor of the survival and the QoL of patients with HNC [26,27]. Patients with HNC who show high levels of anxiety and depression, more often react with a sense of hopelessness/ helplessness and constant preoccupation with the illness, resulting in a deterioration of QoL [28,29]. Reported rates of depression and anxiety in patients with HNC range from 20% to 40%, and both cross-sectional and long-term studies have shown that psychological distress is predictor, not only of QoL but also of survival [30].

With regard to the illness perceptions (IPQ-R) and coping strategies (Brief COPE), no significant differences were detected between the measurements before and after radiotherapy. This may be due to the fact that certain strategies had already been adopted before the initial study assessment, i.e., during

the interim period between the diagnosis and the start of radiotherapy. Longitudinal studies [31,32] and a systematic review [33] have concluded that, without specific intervention, illness perceptions and coping strategies show little modification over time even when they are assessed shortly after the diagnosis of a life-threatening illness.

In this study, Behavioural Disengagement (Brief COPE) at T1, as a coping strategy characterized by a passive reaction and a tendency to give up the management of external demands, appeared to affect the domains of Physical, Emotional, Cognitive and Social Functioning at T2. A large body of evidence indicates that denial, behavioural disengagement and self-blame at the initial post-diagnosis assessment are prognostic factors of poor QoL [33,34]. In addition, Substance use as a coping strategy at T1 appeared to be correlated with impairment of the overall health status (GHS) at T2. Unfortunately, one quarter of patients with HNC have been observed to continue to drink or smoke after the diagnosis of HNC [35]. Several factors may contribute to continued use, including psychological distress, lack of motivation and individual perceptions of risk and causal attributions. Several studies have reported that patients with HNC are more likely to rely on avoidant or other maladaptive strategies that are significantly associated with poorer QoL [36,37] and to adopt different strategies in different phases of the treatment [12,38]. Specifically, patients under treatment or in the less-than-6-month post-treatment phase, more frequently adopt such strategies as denial, behavioural disengagement and suppression, than patients assessed either before the start of treatment or at follow-up longer than 6 months after completion of the treatment. When people are defending themselves against a difficult situation, they tend to avoid experiences and unpleasant feelings. In the case of long-term illnesses, patients with strong perceptions of chronic timeline and multiple consequences may adopt denial/avoidance coping mechanisms, according to which they operate as if they did not have the problem; this mechanism may become more intense if they believe that certain behavioural factors have contributed to the onset of the disease [39]. Age, gender, low educational level and previous alcohol and tobacco use have been proposed to explain the adoption of these strategies [40].

In addition, it has been argued that the coping strategies adopted by the patients with HNC vary according to their levels of anxiety and depression [36]. A recent systematic review [1] highlighted the moderate to high correlation between emotional disengagement and psychological distress observed in a total of 12 relevant studies. Strategies that minimize threatening thoughts can be useful in reducing stress at critical stages of the disease, and be associated with positive adjustment [41].

Regarding illness perceptions (IPQ-R), it was found that a stronger perception of controllability by treatment at T1 was associated with better overall health status (GHS) at T2. In addition, perceived personal controllability at T1 was correlated positively with Social functioning and negatively with the H&N35 sum score at T2. Previous studies concluded that efficacy beliefs and perceived curability should be taken into account, as they appear to have a significant impact on the psychological well-being and QoL of patients with malignancy [42,43].

Moreover, Emotional Representations (IPQ-R) at T1, i.e., the extent to which the illness is believed to result in psychological distress symptoms, was positively correlated with the H&N35 sum score at T2. This means that the more strongly the patient believes that the illness will affect their emotional well-being, the more severe are the symptoms, and, consequently, the poorer is their QoL. The H&N35 sum score at T2 was also correlated with the Illness coherence (IPQ-R), which reflects the individual's overall coherent understanding of the illness. Thus, when patients fail to understand the illness in a coherent way, their experience of symptoms is more intense. It had already been shown that various illness perceptions are correlated with certain dimensions of QoL, and that those patients who focus their attention on the symptoms, experience intense fear of relapse, exhibit emotional responses to their illness and self-blame, and have poorer QoL. In particular, Emotional Representations and perceived Treatment Controllability are consistently reported to be correlated with the outcome of the illness, as demonstrated by a recent

meta-analysis [44]. Another meta-analysis, on the relationship between illness perceptions and emotional distress of patients in a variety of physical illnesses [33], concluded that patients experiencing a strong sense of hopelessness were more likely to have higher levels of anxiety and depression, in contrast to patients with positive perceptions of the effects of the illness and an increased sense of external control. However, a high level of heterogeneity is observed in the association of illness perceptions with health outcomes in cancer patients. This is probably due to the multiple moderators related to the diagnosis and treatment, and further investigation is recommended on the causal effects of illness perceptions on psychological distress, coping and QoL [45].

Suggestions and Limitations

Regular psychological assessment and early inclusion in a psychotherapy program is required for patients to achieve better adaptation to illness, with cognitive restructuring, adoption of effective coping strategies and decrease of the psychological burden [43]. Interventions focusing on the enhancement of beliefs about controllability appear to be more effective in improving emotional well-being, preventing chronic psychopathology and improving QoL [31,46]. It is of note that, although there are concerns about the participation rate of patients with HNC in psychological interventions because of low socio-economic level, they are reported to be well-involved in psycho-educational programs and cognitive behavioural therapy (CBT) [47].

The present study has some limitations and cautious interpretations of the results are needed. Firstly, the sample was small, as, given the population of the district where the study was conducted and the incidence of H&N cancer, it was possible only 50 participants to be included during the study period. Secondly, in the present study, only patients undergoing radiotherapy participated while patients with a complete removal of tumor via surgical intervention or other type of treatment were not included, a fact that limits the possibility of generalizing the results. Thirdly, the time that had elapsed between the diagnosis and the initial assessment of patients is an additional limitation, as it was not possible to evaluate prior modifications of illness perceptions and coping strategies, at what time these may had been occurred and what factors contributed. Finally, the heterogeneity of treatment which the patients had undergone prior to joining the study is a factor that restricts the generalizability of the results.

Despite the aforementioned limitations, some strengths of the study are its prospective design, the use of the extended version of the IPQ, i.e., the IPQ-R, which assesses more fully the basic dimensions of the questionnaire, and the focussing on the role of certain psychological factors on the QoL of HNC patients undergoing radiotherapy, a subject that has not been adequately studied.

It would be important for future research to be conducted with a larger sample, starting assessment immediately after diagnosis, and to conduct a long-term comparative study of the impact of psycho-educational intervention and the enhancement of effective coping strategies on the QoL of patients with HNC undergoing radiation therapy.

CONCLUSIONS

In conclusion, depressive symptoms before the start of radiotherapy are predictors of impairment of the QoL of patients with HNC three months after the completion of treatment. Behavioral Disengagement as a coping strategy and strong beliefs about emotional representations of the illness may also predict impairment in certain domains of the QoL of patients with HNC at follow-up, while belief in treatment controllability may enhance QoL. The identification and the understanding of the depressive symptoms of patients, their beliefs about their illness and their avoidance coping strategies can provide the basis for timely implementation of appropriate intervention aimed at improving the QoL of patients with HNC undergoing radiation therapy.

ARTICLE INFORMATION

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