

Nutritional status of patients treated with radiotherapy as determined by subjective global assessment

Woong Sub Koom, MD¹, Seung Do Ahn, MD², Si Yeol Song, MD², Chang Geol Lee, MD³,
Sung Ho Moon, MD⁴, Eui Kyu Chie, MD⁵, Hong Seok Jang, MD⁶, Young-Taek Oh, MD⁷,
Ho Sun Lee, RD⁸, Ki Chang Keum, MD¹

Department of Radiation Oncology, ¹Yonsei University College of Medicine, Seoul, ²Asan Medical Center, Seoul, ³Gangnam Severance Hospital, Yonsei University College of Medicine, Seoul, ⁴Research Institute and Hospital, National Cancer Center, Goyang, ⁵Seoul National University College of Medicine, Seoul, ⁶Seoul St. Mary's Hospital, The Catholic University of Korea College of Medicine, Seoul, and ⁷Ajou University School of Medicine, Suwon; ⁸Department of Nutrition and Dietetics, Severance Hospital, Yonsei University Health System, Seoul, Korea

Purpose: The purpose of this prospective multi-institutional study was to evaluate the nutritional status of patients undergoing radiotherapy (RT) for treatment of head and neck, lung, or gastrointestinal cancer.

Materials and Methods: A total of 1,000 patients were enrolled in this study at seven different hospitals in Seoul, Korea between October 2009 and May 2010. The nutritional status of patients after receiving 3 weeks of RT was evaluated using subjective global assessment (SGA). The nutritional status of each patient was rated as well nourished (A), moderately malnourished (B), or severely malnourished (C).

Results: The mean age of patients in this study was 59.4 ± 11.9 years, and the male to female ratio was 7:3. According to the SGA results, 60.8%, 34.5%, and 4.7% of patients were classified as A, B, or C, respectively. The following criteria were significantly associated with malnutrition (SGA B or C; $p < 0.001$): loss of subcutaneous fat or muscle wasting (odds ratio [OR], 11.473); increased metabolic demand/stress (OR, 8.688); ankle, sacral edema, or ascites (OR, 3.234); and weight loss ≥5% (OR, 2.299).

Conclusion: SGA was applied successfully to assess the nutritional status of most patients. The prevalence of malnutrition in a radiation oncology department was 39.2%. The results of this study serve as a basis for implementation of nutrition intervention to patients being treated at radiation oncology departments.

Keywords: Malnutrition, Nutrition assessment, Subjective global assessment, Radiotherapy

Introduction

Malnutrition is a potentially serious condition often comorbid with cancer and its treatment. The incidence of malnutrition in cancer patients has been reported to range from about 10% to 80% and malnutrition itself was one of the reasons for death

in up to 20% of cancer patients [1-3]. Malnutrition contributes to an increased risk of toxicity, infection, and healthcare costs, as well as decreased treatment response, compliance, quality of life, and ultimately patient survival [4-6]. Given the importance of nutritional status, screening patients at risk for malnutrition and providing a nutrition support program is of great concern

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Correspondence: Ki Chang Keum, MD, Department of Radiation Oncology, Yonsei University College of Medicine, 50 Yonsei-ro, Seodaemun-gu, Seoul 120-752, Korea. Tel: +82-2-2228-8112, Fax: +82-2-312-9033, E-mail: kckeum@yuhs.ac

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for patients with cancer [7]. Subjective global assessment (SGA) is a validated clinical tool for assessing nutritional status and is based on the features of a medical history and physical examination [8]. First described more than two decades ago, SGA is still being successfully used as a standardized method of assessing nutritional status in various patient populations, including those with cancer [7-12]. Furthermore, nutritional status using SGA was associated with prognosis and quality of life in cancer patients [13-15].

The acute reaction of the aerodigestive tract as a result

of undergoing radiotherapy (RT) is associated with diverse gastrointestinal symptoms and decreased food intake, resulting in deterioration of the patient's nutritional status [16]. Several prospective, randomized trials have reported the effectiveness of early nutritional intervention on the quality of life in cancer patients undergoing RT [17-20]. Despite the frequent occurrence of nutritionally related side effects, a nutritional assessment of patients with cancer receiving RT is not routine practice in the clinical setting. There has been no study as of yet investigating the prevalence of malnutrition during

A. History

1. Weight change
 Overall loss in past 6 months: _____ kg Percent loss _____
 Change in past 2 weeks: _____ increase _____ no change _____ decrease

2. Dietary intake change relative to normal
 No change _____
 Change: duration _____ weeks _____ months
 Type:
 sub-optimal solid diet _____ full liquid diet _____
 hypocaloric liquid diet _____ starvation _____

3. Gastrointestinal symptoms (persisting more than 2 weeks)
 None _____ Nausea _____ Vomiting _____ Diarrhea _____ Anorexia _____

4. Functional capacity
 No dysfunction _____
 Dysfunction: duration _____ weeks _____ months
 Type: working sub-optimally _____ ambulatory _____ bedridden _____

5. Disease and its relationship to nutritional requirements
 Primary diagnosis: _____
 Metabolic demand/Stress: none _____ low _____ moderate _____ high _____

B. Physical Examination (for each specify: 0 = normal, 1+ = mild, 2+ = moderate, 3+ = severe)
 Loss of subcutaneous fat (triceps, chest) _____
 Muscle wasting (quadriceps, deltoids) _____
 Ankle edema _____ Sacral edema _____ Ascites _____

C. Subjective Global Assessment Rating
 Well nourished A _____
 Moderately (or suspected of being) malnourished B _____
 Severely malnourished C _____

Fig. 1. Feature of subjective global assessment by Detsky et al. [8].

Table 1. Patients and nutritional characteristics of all subjects

Characteristic		Value
Age (yr)		59.4 ± 11.9
Sex	Male	703 (70.3)
	Female	297 (29.7)
Tumor site	Head and neck	286 (28.6)
	Gastrointestinal tract	444 (44.4)
Chemotherapy	Lung	270 (27.0)
	No	528 (52.8)
	Yes	472 (47.2)
Weight loss in past 6 mo (kg)		2.1 ± 3.9
Percent weight loss in past 6 mo (%)		3.9 ± 4.8
Weight change in previous 2 wk	Increase	110 (11.0)
	No change	590 (59.0)
	Decrease	275 (27.5)
	Unknown	25 (2.5)
Dietary intake change	No	667 (66.7)
	Yes	331 (33.1)
Duration of diet change (wk)		3.8 ± 7.3
Type of diet change	Suboptimal solid diet	174 (17.4)
	Liquid diet	66 (6.6)
	Starvation	6 (0.6)
	Unknown	85 (8.5)
	No	701 (70.1)
Nausea	Yes	299 (29.9)
	No	909 (90.9)
Vomiting	Yes	91 (9.1)
	No	912 (91.2)
Diarrhea	Yes	88 (8.8)
	No	494 (49.4)
Anorexia	Yes	506 (50.6)
	No	619 (61.9)
Functional capacity	Yes	380 (38.0)
	No	9.2 ± 26.3
Duration of dysfunction (wk)		9.2 ± 26.3
Type of dysfunction	Suboptimal	222 (22.2)
	Ambulatory	134 (13.4)
	Bedridden	14 (1.4)
	Unknown	10 (1.0)
Metabolic demand/stress	No	123 (12.3)
	Low	661 (66.1)
	Moderate	174 (17.4)
	High	32 (3.2)
	Unknown	10 (1.0)
Loss of subcutaneous fat	Normal	631 (63.1)
	Mild	254 (25.4)
	Moderate	104 (10.4)
	Severe	11 (1.1)

Table 1. Continued

Characteristic		Value
Muscle wasting	Normal	675 (67.5)
	Mild	221 (22.1)
	Moderate	91 (9.1)
	Severe	12 (1.2)
Ankle edema	Unknown	1 (0.1)
	Normal	901 (90.1)
	Mild	62 (6.2)
	Moderate	33 (3.3)
Sacral edema	Severe	4 (0.4)
	Normal	915 (91.5)
	Mild	48 (4.8)
Ascites	Moderate	33 (3.3)
	Severe	4 (0.4)
	Normal	902 (90.2)
	Mild	44 (4.4)
Subjective global assessment	Moderate	34 (3.4)
	Severe	4 (0.4)
	Unknown	16 (1.6)
	Well nourished	608 (60.8)
Subjective global assessment	Moderately malnourished	345 (34.5)
	Severely malnourished	47 (4.7)
	Severely malnourished	

Values are presented as mean ± standard deviation or number (%).

radiotherapy in Korea. This prospective multi-institutional study was performed to evaluate the nutritional status using SGA in ambulatory patients undergoing RT to the head, neck, thorax, abdomen, and pelvic area.

Materials and Methods

1. Participants

This prospective study was designed to investigate the nutritional status of patients with cancer receiving RT. Between October 2009 and May 2010, consecutive 1,000 patients were enrolled in seven different hospitals in Seoul, Korea. Eligibility criteria included: 1) having primary cancer of the head and neck, lung, or gastrointestinal tract; 2) receiving RT over a period of 3 weeks to the primary tumor site, regardless of RT intent (primary, adjuvant to surgery, combined with chemotherapy, or palliation); and 3) willing and able to give written informed consent. We concerned the acute radiation response of the aerodigestive tract. So, the patients treated with RT over 3 weeks to the head and neck, lung, or gastrointestinal tract were enrolled. Patients who were

Table 2. Patient and nutritional characteristics of subjects according to subjective global assessment category

Characteristic	Status	Well nourished (A)	Malnourished (B and C)	p-value
Age (yr)	<60	293 (48.2)	169 (43.1)	0.116
	≥60	315 (51.8)	223 (56.9)	
Sex	Male	419 (68.9)	284 (72.4)	0.232
	Female	189 (31.1)	108 (27.6)	
Tumor site	Head and neck	172 (28.3)	114 (29.1)	0.799
	Gastrointestinal tract	275 (45.2)	169 (43.1)	
	Lung	161 (26.5)	109 (27.8)	
Chemotherapy	No	342 (56.3)	186 (47.4)	0.006
	Yes	266 (43.8)	206 (52.6)	
Percent weight loss in past 6 mo (%)	<5	496 (81.6)	167 (42.6)	<0.001
	≥5	108 (17.8)	223 (56.9)	
Weight change in previous 2 wk	Increase or no change	496 (81.6)	204 (52.0)	<0.001
	Decrease	100 (16.4)	175 (44.6)	
Dietary intake change	No	478 (78.6)	189 (48.2)	<0.001
	Yes	130 (21.4)	201 (51.3)	
Nausea	No	464 (76.3)	237 (60.5)	<0.001
	Yes	144 (23.7)	155 (39.5)	
Vomiting	No	588 (96.7)	321 (81.9)	<0.001
	Yes	20 (3.3)	71 (18.1)	
Diarrhea	No	565 (92.9)	347 (88.5)	0.016
	Yes	43 (7.1)	45 (11.5)	
Anorexia	No	373 (61.3)	121 (30.9)	<0.001
	Yes	235 (38.7)	271 (69.1)	
Functional capacity	No	423 (69.6)	196 (50.0)	<0.001
	Yes	185 (30.4)	195 (49.7)	
Metabolic demand/stress	No or low	570 (93.8)	214 (54.6)	<0.001
	Moderate or high	36 (5.9)	170 (43.4)	
Loss of subcutaneous fat	No	528 (86.8)	103 (26.3)	<0.001
	Yes	80 (13.2)	289 (73.7)	
Muscle wasting	No	525 (86.3)	150 (38.3)	<0.001
	Yes	83 (13.7)	241 (61.5)	
Ankle edema	No	576 (94.7)	325 (82.9)	<0.001
	Yes	32 (5.3)	67 (17.1)	

Table 2. Continued

Characteristic	Status	Well nourished (A)	Malnourished (B and C)	p-value
Sacral edema	No	578 (95.1)	337 (86.0)	<0.001
	Yes	30 (4.9)	55 (14.0)	
Ascites	No	563 (92.6)	339 (86.5)	<0.001
	Yes	30 (4.9)	52 (13.3)	

Values are presented as number (%).

obviously moribund, not willing to participate, or not able to respond during interview were excluded. The ethics committee of each study hospital approved the study protocol.

p < 0.05. Data were analyzed using SPSS ver. 18.0 (SPSS Inc., Chicago, IL, USA).

2. Nutritional assessment and data collection

The nutritional status of each patient was assessed 3 weeks after the initiation of RT, and was determined using SGA tool [8]. The 3-week period between RT commencement and nutritional assessment was chosen because RT-induced acute toxicity, such as mucositis, develops during this period. The SGA tool is based on a medical history (weight loss; dietary intake change; gastrointestinal symptoms including nausea, vomiting, diarrhea and anorexia; and changes in functional capacity) and physical examination (loss of subcutaneous fat; muscle wasting; ankle edema, sacral edema, and ascites). Each patient was classified as either well nourished (SGA A), moderately or suspected of being malnourished (SGA B), or severely malnourished (SGA C). This classification was assigned on the basis of subjective weighting. The details of SGA assessment was described by Detsky et al. [8] (Fig. 1). For the purpose of this analysis, malnutrition was defined as either SGA B or SGA C. A single trained rater (physician or dietitian) in each hospital assessed nutritional status. In order to minimize inter-rater variation, all raters were educated by the principle dietician of the coordinating study center. In addition to recording nutritional information, the following information was collected from the medical record within 4 weeks before or during RT: age, sex, diagnosis, and chemotherapy.

3. Statistical analysis

All continuous variables and categorical variables were transformed into two categorical variables. Univariate analysis was conducted by performing chi-square tests to compare the association between SGA criteria and malnutrition. To identify independent risk factors for malnutrition, binary logistic regression analysis was performed. Significance was defined as

Results

1. Characteristics and nutritional status of participants

Baseline characteristics of the participants are shown in Table 1. The mean age of enrolled subjects was 59.4 ± 11.9 years. Of the total 1,000 patients, 703 (70.3%) were men and 293 (29.3%) were women. Gastrointestinal cancer, including esophageal cancer, was the most the common parameter and was present in 444 of the patients. Head and neck cancer and lung cancer affected 286 patients and 270 patients, respectively. Chemotherapy was performed either during RT or within the 4 weeks before RT in 472 patients. According to SGA, 608 (60.8%), 345 (34.5%), and 47 (4.7%) patients were SGA A, SGA B, and SGA C, respectively.

2. Impact of SGA parameters and clinical factors on malnutrition

Table 2 shows the results of univariate analysis of clinical factors and SGA parameters for each SGA classification. For this analysis, SGA B and C were grouped together in the malnourished category. Additionally, all continuous variables were transformed into categorical variables. Except for tumor site, all categorical variables were re-coded into two groups. There was no significant association between malnutrition (SGA B and C) and age (p = 0.116), sex (p = 0.232), or tumor site (p = 0.799). Chemotherapy in addition to all SGA parameters were significantly associated with malnutrition (SGA B and C). In a multivariate analysis, similar SGA parameters were coded as one (Table 3). Loss of subcutaneous fat or muscle wasting strongly was associated with the development of malnutrition (odds ratio [OR], 11.473; p < 0.001). Metabolic demand/stress was the next most strongly contributing factor (OR, 8.688; p < 0.001). Patients with ankle, sacral edema, or ascites upon

Table 3. Independent risk factors for malnourishment (subjective global assessment B and C)

	OR (95% CI)	p-value
Chemotherapy	1.180 (0.821-1.697)	0.370
Percent weight loss in past 6 mo	2.299 (1.567-3.373)	<0.001
Dietary intake change	1.407 (0.911-2.172)	0.124
Nausea/vomiting	1.080 (0.703-1.661)	0.724
Diarrhea	1.412 (0.723-2.756)	0.312
Anorexia	2.018 (1.356-3.004)	0.001
Functional capacity	1.105 (0.736-1.658)	0.632
Metabolic demand/stress	8.688 (5.319-14.192)	<0.001
Loss of subcutaneous fat or muscle wasting	11.473 (7.588-17.347)	<0.001
Ankle, sacral edema or ascites	3.234 (1.716-6.094)	<0.001

OR, odds ratio; CI, confidence interval.

physical examination were also at a higher risk of being malnourished (OR, 3.234; $p < 0.001$). Additionally, patients with $\geq 5\%$ weight loss in the previous 6 months were also more likely than other patients to be malnourished (OR, 2.299; $p < 0.001$).

Discussion and Conclusion

Three weeks after initiation of RT, we used SGA to evaluate the nutritional status of ambulatory patients with cancer whose site of treatment varied. SGA was applied successfully to assess the nutritional status of most patients. The prevalence of patients who were malnourished or at risk of becoming malnourished while being treated at one of the seven hospitals in the study population was high at 39.2%.

Malnutrition is common in cancer patients, and has been reported to occur in up to 80% of patients treated with RT. Furthermore, malnutrition is associated with increased morbidity and mortality [2,6]. It has been suggested that malnutrition increases the risk of infections caused by immune dysfunction, as well as decreasing quality of life and survival [4]. Moreover, malnutrition adversely affects patients' response to cancer treatment and increases the incidence of treatment-related toxicities. Oncologic treatment exacerbates acute and chronic symptoms due to issues with poor food intake and poor nutritional status [21]. Most of the RT-related toxicities are closely associated with nutritional problems [16]. Given the deteriorating side effect of RT, several prospective, randomized

trials have been conducted to evaluate the effectiveness of nutritional intervention on the quality of life as well as nutritional status in patients with cancer undergoing RT [17-20,22,23]. Consistently, the nutritional intervention positively influenced weight, nutritional status, and quality of life compared to usual care in patients receiving RT to the gastrointestinal or head and neck area [17,19,20,22]. Therefore, early identification of malnutrition in cancer patients and nutritional intervention may increase tolerance to cancer treatment and improve quality of life and prognosis [3,4,24,25].

Many nutrition screening tools have been proposed that combine multiple components, including dietary and medical history, weight loss, biochemical indicators of body protein, and anthropometry [26]. SGA for determining nutritional status was developed by Detsky et al. [8] in 1987. This assessment tool is simple, non-invasive, and inexpensive, consisting of a medical history (weight change, dietary intake change, gastrointestinal symptoms, and changes in functional capacity) and a physical examination (loss of subcutaneous fat, muscle wasting, ankle edema, sacral edema, and ascites). SGA categorizes patients into three nutritional status groups: well nourished (SGA A), moderately malnourished (SGA B), and severely malnourished (SGA C).

SGA is a reliable tool that enables correlation of a number of objective nutritional assessment indicators, such as morbidity, mortality, and quality of life (QoL) measures, and has been used universally in diverse clinical settings, various regions, and multicenter studies [7,9-12,27,28]. A limitation of using SGA is that it only classifies subjects into three general groups, and it does not reflect subtle changes in nutritional status. Ottery [7] developed a patient-generated SGA tool using a scoring system for patients with cancer that allowed data to be expressed as a continuous measurement. Using this tool, investigators can detect subtle changes of nutritional status. For more than two decades, however, the original SGA has been used to assess patient malnutrition in several clinical settings and is the standard method [27]. In this study, we have successfully conducted nutritional surveys using SGA in multiple institutions, using rater education to support optimal consistency. According to SGA, the prevalence of malnutrition in this study population was 39.2%. This finding is similar to the 35% of patients who were malnourished in the Australian Radiation Oncology Facilities as determined by patient-generated SGA [28]. We demonstrated the malnutrition in large number of ambulatory patients undergoing RT. This is first step to raise concern about nutrition assessment at

radiation oncology departments.

The group of patients enrolled this study is very heterogeneous and includes various levels of tumor burden and RT protocol across multiple institutions. In addition, RT parameters such as dose and volume were variable. RT-related toxicities associated with nutritional problems were not evaluated. This study did not investigate inter-rater reliability or reproducibility. To minimize observer bias, SGA assessment was performed after a training period that took place over a few days. A well-trained observer can distinguish between malnourished (SGA categories B and C) and normal patients, although they had more difficulty discriminating between moderate and severe malnutrition [29]. This study applied the SGA only at one time point and is observational because no nutritional intervention was performed. Nevertheless, our study analyzed a large population of patients with cancer receiving RT and adds the clinically useful information that malnutrition is an important problem in cancer patients undergoing RT. Further studies are needed to evaluate the effectiveness of nutritional intervention at several time points in a homogenous group of patients to determine the benefit of nutritional support on clinical outcomes in cancer patients treated with RT. In conclusion, this study identified the prevalence of malnutrition using the SGA tool in patients treated with RT to the head and neck, thorax, abdomen and pelvic area. Thirty-nine point two percent of patients were moderately or suspected of being malnourished or severely malnourished. The results of this study serve as a basis for implementation of nutrition intervention to patients being treated at radiation oncology departments.

Conflict of Interest

No potential conflict of interest relevant to this article was reported.

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