

Frequency of Symptoms in Patients With Cancer Receiving Chemotherapy, According to Gender and the Primary Site of the Cancer in Kashan, Iran, 2017

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DOI: 10.30699/acadpub.mci.3.1.17

Submitted: 22 September 2018

Revised: 19 October 2018

Accepted: 27 November 2018

e-Published: 1 January 2019

Keywords:

Neoplasms

Drug Therapy

Symptom Assessment

Abstract

Introduction: Patients with cancer under chemotherapy usually manifest a wide range of symptoms. The frequency of these symptoms is reported differently. The current study aimed at addressing the common symptoms of cancer, according to gender and primary site of cancer in patients receiving chemotherapy in Kashan, Iran, in 2017.

Methods: The current cross-sectional study was conducted on 186 patients with cancer. The frequency of 14 common symptoms in cancer was evaluated with interviewing the patients. The data were analyzed with SPSS version 13 using descriptive statistics, chi-square, the Mann-Whitney U, and the Kruskal-Wallis tests.

Results: About half of the subjects were female, and 58.1% aged above 50 years. The common primary sites of cancer were breast (23.1%), colon (21%), bone (11.3%), and lung (7%). The most common symptoms were fatigue (86.6%), followed by dry mouth, insomnia, anorexia, and pain. The subjects reported 6.672.63± symptoms in average. The patients with the history of surgery reported significantly more symptoms. The symptoms of insomnia, nausea, and diarrhea were significantly higher in female patients, while anorexia was significantly higher in male ones. Dry mouth, diarrhea, and total number of symptoms were significantly associated with the primary site of the cancer.

Conclusions: Patients with cancer experience various symptoms that are different based on the primary site of the cancer and the gender of patients. Any effective actions in palliative care should be based on the accurate assessment of the symptoms.

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INTRODUCTION

Cancer is the second cause of death in the world [1], and the third in Iran [2]. The five-year global prevalence for all cancers was estimated 28.8 million [3]. The incidence of cancer is 20% higher in males compare with females [4]. Stomach cancer in males and breast cancer in females are

the most common cancers in Iran [5]. Cancer and its treatments such as chemotherapy induce different complications. Cancer is associated with malnutrition following metabolic changes, and proteolysis and lipolysis effects of cytotoxic chemotherapies are an example of interrelated

symptoms [6]. Cancer and chemotherapy-induced nausea and vomiting are another example [7]. The symptoms such as lack of energy, dry mouth, and weight-loss can indistinguishably occur due to cancer or chemotherapy [8]. Patients with cancer experience 11 to 13 symptoms in average [9]. Fatigue, pain, nausea, vomiting, constipation, and depression are the common ones [10]. Symptoms of cancer change over the course of the disease. Progression of disease is accompanied by the development of new symptoms or the exacerbation of chronic symptoms [11].

The patients reporting more symptoms tend to have a poorer quality of life [12]. Uncontrolled symptoms are associated with increased emotional distress, decreased physical and social function, reduction in the compatibility, and even death of patients [11, 13, 14]. Studies show that caring needs of patients with cancer are considerably neglected [15-17]. Chan et al., reported nausea as the most common symptom of cancer [18]. While in another study, fatigue was the most common problem in such patients [19]. In a systematic review, the prevalence of fatigue was reported 23%-100%, and the nausea was 2%-78% [20]. These wide ranges of reported symptoms make the data unreliable for clinicians.

The way people experience symptoms has impacts on their healthcare seeking behaviors [21], and give helpful information to health personnel. Gender and primary site of cancer are two variables that might influence the perception of cancer symptoms. Vainio et al., found that the pain was more prevalent in females, and the primary site of the cancer was related to the reported symptoms [22]. An accurate assessment of the symptoms is necessary before any actions [23]. After extensive literature review, the studies on the symptoms of cancer according to the gender of patients and the primary site of the cancer in Iran were not found; therefore, the current study aimed at addressing the issue in Kashan in 2017.

METHODS

Study Design and Sampling

The current cross sectional study was conducted in health care centers of Kashan, Iran from May to November 2017. All the patients diagnosed with cancer by an oncologist and receiving chemotherapy were included in the study; sampling was continued sequentially for six months and 186 patients were enrolled in the study. The inclusion

criteria were definite diagnosis of cancer, age above 18, and having psychological, mental, and physical readiness to respond to the questions.

Instrument

The assessment tool had two parts: The first part included questions about demographic and clinical characteristics, i.e., age, gender, health insurance, disease awareness, current and previous treatments, marital status, and the site of primary cancer. The second part consisted of 14 common symptoms in cancer derived from Edmonton symptom assessment system and the Assessment Symptoms Palliative Elderly [23, 24]. The presence of symptoms in previous week was specified with the Yes or No option. Pain, dry mouth, mouth ulcer, dyspnea, cough, nausea, vomiting, constipation, diarrhea, urinary problems, fatigue, insomnia, anorexia, helplessness, and depression were the 14 recorded symptoms. To determine the scientific validity of the instrument, the questionnaire was distributed among 11 nursing faculty members of Kashan University of Medical Sciences, and content validity ratio (CVR) and content validity index (CVI) were calculated 0.63 and 0.81, respectively. The test-retest reliability was 0.87.

Data Gathering

First author attended the chemotherapy centers and questionnaires were completed through interview in the waiting rooms before starting the chemotherapy sessions.

Ethical Considerations

The current study was approved by the Ethics Committee of Kashan University of Medical Sciences (ethical code: IR.KAUMS.NUHEPM.REC.1396.02). The objectives of the study were explained to the patients, and written consent was obtained. The patients were assured that their personal information would remain confidential. The study was conducted according to the Declaration of Helsinki.

Data Analysis

Data were analyzed with SPSS version 13. Descriptive statistics were used to express data. The chi-square test was used to analyze the symptoms according to the site of primary cancer, and gender of the patients. The sum of the symptoms was considered as a variable and the Kolmogorov-

Smirnov test showed non-normal distribution of variables. The Mann-Whitney U and the Kruskal-Wallis tests were used to examine the relationship between the number of symptoms and gender, and the primary site of cancer.

RESULTS

The current study was conducted on 186 patients with cancer that 51.1% were female, and 58.1% aged above 50. The demographic and clinical characteristics of patients are provided in Table 1. The frequency of primary site of cancer was as follows: breast (n=43; 23.1%), colon (n=39; 21%), bone (n=21; 11.3%), lung (n=13; 7%), prostate (n=7; 3.8%), cervix (n=7; 3.8%), liver (n=6; 3.2%), lymphoma (n=5; 2.7%) and others or not-specified cases (n=45; 24.2%). The most common symptom was fatigue (86.6%), followed by dry mouth, insomnia, anorexia, and pain (Table 2). The subjects reported 6.67 ± 2.63 symptoms in average. The patients with the history of surgery reported

significantly more symptoms (Table 1).

The most prevalent symptoms among male patients were fatigue (83.5%), anorexia (64.8%), and dry mouth (59.3%), while in female patients there were fatigue (89.5%), insomnia (68.4%), and pain (66.3%). The symptoms of insomnia, nausea, and diarrhea were significantly higher in female patients, while anorexia was significantly higher in male ones (Table 2).

Dry mouth, diarrhea, and total number of symptoms were significantly associated with the primary site of the cancer (Table 3).

DISCUSSION

In the current study, breast and colon were the most frequent primary sites of cancer. Common symptoms were fatigue, dry mouth, insomnia, anorexia, and pain. Insomnia, nausea, and diarrhea were higher in female patients and anorexia was higher in male ones. Symptoms were significantly associated with the primary site of the cancer. Fatigue was the most

Table 1: Demographic and Clinical Information of Patients and Their Relationship With Symptoms

| | Patients, No. (%) | Number of Symptoms, mean \pm SD | P Value |
|------------------------------------|-------------------|-----------------------------------|---------|
| Gender | | | 0.12 |
| Male | 91 (48.9) | 6.36 \pm 2.7 | |
| Female | 95 (51.1) | 7 \pm 2.5 | |
| Age, y | | | 0.36 |
| <50 | 78 (41.9) | 6.4 \pm 2.7 | |
| >50 | 108 (58.1) | 6.8 \pm 2.5 | |
| Healthcare Insurance | | | 0.99 |
| Yes | 177 (95.2) | 6.6 \pm 2.6 | |
| No | 9 (4.8) | 7 \pm 2.8 | |
| Aware of Illness | | | 0.76 |
| Yes | 140 (75.3) | 6.6 \pm 2.7 | |
| No | 46 (24.7) | 6.6 \pm 2.1 | |
| Chemotherapy History | | | 0.90 |
| Yes | 72 (38.7) | 6.7 \pm 2.5 | |
| No | 114 (61.3) | 6.6 \pm 2.7 | |
| Surgical History | | | 0.007 |
| Yes | 106 (57) | 7 \pm 2.6 | |
| No | 80 (43) | 6.2 \pm 2.5 | |
| Radiotherapy History | | | 0.22 |
| Yes | 71 (38.2) | 7 \pm 2.5 | |
| No | 115 (61.8) | 6.5 \pm 2.7 | |
| The Session of Chemotherapy | | | 0.11 |
| <5 | 75 (40.4) | 6.1 \pm 2.4 | |
| 5-10 | 65 (34.9) | 6.9 \pm 2.9 | |
| >10 | 46 (24.7) | 7.1 \pm 2.3 | |

Table 2: The Frequency of Symptoms by Gender in Patients With Cancer

| | Frequency | Male | Female | P Value |
|--|--------------------|------------------|------------------|-------------|
| Fatigue, No. (%) | 161 (86.6) | 76 (83.5) | 85 (89.5) | 0.16 |
| Dry Mouth, No. (%) | 117 (62.9) | 54 (59.3) | 63 (66.3) | 0.20 |
| Insomnia, No. (%) | 116 (62.4) | 51 (56) | 65 (68.4) | 0.05 |
| Anorexia, No. (%) | 115 (61.8) | 59 (64.8) | 56 (58.9) | 0.05 |
| Pain, No. (%) | 114 (61.3) | 51 (56) | 63 (66.3) | 0.09 |
| Constipation, No. (%) | 99 (53.32) | 52 (57.1) | 47 (49.5) | 0.18 |
| Hopelessness and Depression, No. (%) | 98 (52.7) | 50 (51) | 48 (50.5) | 0.32 |
| Nausea, No. (%) | 87 (46.8) | 33 (36.3) | 54 (56.8) | 0.004 |
| Urinary Problems, No. (%) | 66 (35.5) | 37 (40.7) | 29 (30.5) | 0.09 |
| Oral Ulcer, No. (%) | 61 (32.8) | 27 (29.7) | 34 (35.8) | 0.23 |
| Dyspnea, No. (%) | 60 (32.3) | 26 (28.6) | 34 (35.8) | 0.18 |
| Cough, No. (%) | 52 (28) | 24 (26.4) | 28 (29.5) | 0.38 |
| Diarrhea, No. (%) | 51 (27.4) | 19 (20.9) | 32 (33.7) | 0.03 |
| Vomiting, No. (%) | 45 (24.2) | 20 (22) | 25 (26.3) | 0.30 |
| The Total Number of Symptoms, mean ± SD | 6.67 ± 2.63 | 6.3 ± 2.7 | 6.9 ± 2.5 | 0.11 |

common reported symptom in both genders. In a meta-analysis by Van Lancker et al., fatigue was reported as the most common symptom in patients with cancer [25]. Riechelmann et al., also reported fatigue as the most common symptom [26]. Lack of energy and weakness can lead to various problems such as frustration, anxiety, and depression [27].

Dry mouth was the second prevalent symptom. Dry mouth can reduce nutrition, and quality of life. In the study by Mercadent et al., the frequency of dry mouth was 40.4% [28] that was less than that of the current study. In Jordanian patients with cancer, dry mouth was 82.9% [12]. Dry mouth was also more common in patients with breast and liver cancers. The least common symptom in current study was vomiting that was consistent with the findings of Van Lancker et al. [25], but the frequency of vomiting was more in the study by Okuyama [17]. This discrepancy could be due to differences in chemotherapy regimen, tolerance, and type of anti-nausea drugs. Nausea was more common in the current study. Studies by Ware showed that more than half of the patients undergoing chemotherapy experience nausea or vomiting, either due to cancer or due to its complications and treatments [29]. This rate was comparable with that of the current study. The nausea was significantly higher in female patients.

Oral ulcer was reported by 61 (32.8%) patients. According to the study by Naidu et al., oral ulcers are adverse effects of radiation therapy and chemotherapy, which changes the nutritional status

of the patients. This symptom was experienced by 40% of patients [30] that was somewhat consistent with that of the current study. Insomnia was the third reported problem in the studied patients. In the study by Moustian et al., the incidence of insomnia was up to 90%. Insomnia symptoms include daily sleepiness, difficulty in falling asleep, difficulty in sleeping, and early waking trouble, which can increase mortality and morbidity in such patients [31]. Insomnia was significantly higher in females compare with males in present study. A meta-analysis suggested a female predisposition to insomnia. The risk ratio of insomnia was 1.41 (95% confidence interval (CI): 1.28-1.55) for female versus male patients [32].

Pain was the fifth prevalent symptom in the current study. Pain was higher in other studies [12, 22]. But the finding of the current study was comparable with that of the meta-analysis by Van Lancker [25]. Pain was more common in females than males, although the difference was insignificant. A meta-analysis showed that the prevalence of pain in the patients under anticancer therapies was 59%, [33]. Pain is a common and disturbing problem and despite the provided suggestions to control it, pain still is a major problem in such patients. Anorexia with the frequency of 115 (61.8%) was significantly higher in male patients. Anorexia, cachexia, and loss of adipose and muscle tissue are reported in half of all patients with cancer. Cachexia can have a profound impact on quality of life. Weight loss in patients with cancer is associated with higher chemotherapy-related side effects, which decreases survival rate [34].

Table 3: The Prevalence of Symptoms by the Primary Site of Cancer

| | The Primary Site of Cancer, No. (%) | | | | | | | P Value | | |
|-----------------------------------|-------------------------------------|----------------|-----------------|-------------------|-------------------|----------------|----------------|-------------|-----------------|-----------------|
| | Breast (n=43) | Lung (n=13) | Cervix (n=7) | Prostate (n=7) | Lymphoma (n=5) | Liver (n=6) | Bone (n=21) | | Colon (n=39) | Other (n=45) |
| Pain, No. (%) | 30 (69.8) | 6 (46.2) | 4 (57.1) | 4 (57.1) | 1 (20) | 5 (83.3) | 11 (52.4) | 26 (66.7) | 27 (60) | 0.36 |
| Dry mouth, No. (%) | 32 (74.4) | 3 (23.1) | 4 (57.1) | 3 (42.9) | 4 (80) | 5 (83.3) | 14 (66.7) | 31 (79.5) | 21 (46.7) | 0.002 |
| Dyspnea, No. (%) | 14 (32.6) | 6 (46.2) | 1 (14.3) | 1 (14.3) | 2 (40) | 3 (50) | 8 (38.1) | 9 (23.1) | 16 (35.6) | 0.61 |
| Cough, No. (%) | 14 (32.6) | 6 (46.2) | 0 (0) | 1 (14.3) | 2 (40) | 1 (16.7) | 10 (47.6) | 9 (23.1) | 9 (20) | 0.12 |
| Nausea, No. (%) | 25 (58.1) | 5 (38.5) | 4 (57.1) | 0 (0) | 2 (40) | 3 (50) | 6 (28.6) | 19 (48.7) | 23 (51.1) | 0.13 |
| Vomiting, No. (%) | 13 (30.2) | 4 (30.8) | 2 (28.6) | 0 (0) | 0 (0) | 1 (16.7) | 2 (9.5) | 12 (30.8) | 11 (24.4) | 0.37 |
| Constipation, No. (%) | 22 (51.2) | 8 (61.5) | 4 (57.1) | 5 (71.4) | 1 (20) | 2 (33.3) | 15 (71.4) | 8 (46.2) | 24 (53.3) | 0.41 |
| Diarrhea, No. (%) | 15 (34.9) | 2 (15.4) | 2 (28.6) | 0 (0) | 0 (0) | 2 (33.3) | 2 (9.5) | 18 (46.2) | 10 (22.2) | 0.02 |
| Fatigue, No. (%) | 39 (90.7) | 10 (76.9) | 6 (85.7) | 5 (71.4) | 4 (80) | 6 (100) | 18 (85.7) | 36 (92.3) | 37 (82.2) | 0.64 |
| Anorexia, No. (%) | 29 (67.4) | 6 (46.2) | 5 (71.4) | 2 (28.6) | 1 (20) | 3 (50) | 14 (66.7) | 25 (64.1) | 30 (66.7) | 0.24 |
| Depression, No. (%) | 23 (53.5) | 5 (38.5) | 5 (71.4) | 2 (28.6) | 2 (40) | 4 (66.7) | 10 (47.6) | 22 (56.4) | 25 (55.6) | 0.75 |
| Urinary Tract Problem, No. (%) | 15 (34.9) | 3 (23.1) | 1 (14.3) | 4 (57.1) | 1 (20) | 1 (16.7) | 11 (52.4) | 16 (41) | 14 (31.1) | 0.37 |
| Insomnia, No. (%) | 30 (69.8) | 9 (69.2) | 6 (85.7) | 4 (57.1) | 3 (60) | 1 (16.7) | 13 (61.9) | 24 (61.5) | 26 (57.8) | 0.36 |
| Oral Ulcer, No. (%) | 18 (41.9) | 2 (15.4) | 2 (28.6) | 2 (28.6) | 2 (40) | 1 (16.7) | 8 (38.1) | 12 (30.8) | 14 (31.1) | 0.78 |
| Number of symptoms, mean \pm SD | 7.4 \pm 2.5 | 5.7 \pm 3.7 | 6.6 \pm 2.2 | 4.7 \pm 1.8 | 5 \pm 3.1 | 6.3 \pm 2.8 | 6.7 \pm 2.6 | 7.1 \pm 2 | 7.4 \pm 2.7 | 0.02 |

Constipation was reported by 99 (53.32%) patients. Diarrhea was less common than constipation and it was significantly more prevalent in females in current study. According to Stein, diarrhea was one of the most common problems associated with chemotherapy, especially in patients with advanced cancer, and its prevalence was 50%-80%, which was not consistent with the results of the current study [35]. The frequency of depression in the current study was 98 (52.7%). In a meta-analysis, the prevalence of major depressive disorders in patients with cancer was 3% to 40%, and the average was approximately 11% that was twice compared with that of the general population [36].

The current study findings showed that the primary site of the cancer might be related to the type and number of symptoms, which was consistent with the study by McPherson et al., [37]. A survey of Canadian oncology nurses revealed that the most common identified symptoms were fatigue, pain, nausea, constipation, and anxiety. Frequent symptoms, which nurses felt low confidence in managing them were anxiety, neuropathy, depression, anorexia, and skin alterations [38]. It shows that nurses have a clear view of cancer symptoms and limitations to management them. Many times, there are concurrent symptoms such as pain, fatigue, and sleep disturbances, or nausea, vomiting, and poor appetite that are related to each other and are called a symptom cluster. Some believe that it is quite better to look at the cluster of symptoms or nested structure of symptoms since their management is interrelated [9]. Some believe it is better to focus on core symptoms that means the symptoms that remain consistent over time [11].

The current study had some strength. First, it was about the symptoms of cancer considering gender and primary site of cancer in Iran. This view to the cancer symptom is new. Many believe that cancer is not a single disease, but a group of hundreds of illnesses sharing common biological features [13]; therefore, it might be beneficial to evaluate the symptoms in different types of the disease separately. This might provide more helpful data to clinicians in palliative care settings. Second, the data was completed with interviewing the patients that may reduce the reporting errors. The current study had some limitations. First, the sample size was small; this reduces the reliability of the results. Second, the patients were recruited from limited health care centers in one city; this non-representative sample of

the study affects the generalizability of the results. For example, the frequency of bone cancer was much higher in the current study compared with that of the literature; therefore the findings must be considered with caution. Patients diagnosed with cancer receive many curative or palliative treatments. Surgery, radiotherapy, and chemotherapy are common treatments for cancer [39]. These treatments have their own complications and symptoms. In the current study, along with many other studies mentioned here, it was not possible to attribute the symptoms to the treatments or the cancer itself. It was a limitation of the current study that should be considered when referring to the data. Further studies with the same objectives in a larger population of patients with cancer are recommended.

It seems that the primary site of cancer and gender of patients can influence the perception of symptoms. Patients and their families have uncontrolled symptoms, and this imposes additional costs on the health system. The assessment of cancer symptoms is the first step in symptom management; therefore, it is a crucial issue and needs more studies.

ACKNOWLEDGMENTS

The authors would like to thank all of the patients and nurses of chemotherapy ward of Shahid Beheshti Hospital for their support. Authors also express their gratitude to Research Council of Kashan University of Medical Sciences Research Council for funding the project (grant number: 96045).

CONFLICT OF INTEREST

The authors declared no conflict of interest.

ETHICS APPROVAL

The current study was approved by the Ethics Committee of Kashan University of Medical Sciences (IR.KAUMS.NUHEPM.REC.1396.02).

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