Original Article

Travel Burden and Clinical Profile of Cancer Patients Admitted to the Cancer Institute of Iran in 2012

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Abstract

Background: Burden of cancer is increasing in developing countries, where healthcare infrastructures and resources are limited. Evaluating the pattern of care would provide evidence for planning and improvement of the situation.

Materials and Methods: We studied the pattern of residential place and clinical information of cancer patients who were admitted to the Cancer Institute of Iran from January 1, to May 31, 2012.

Results: We studied 1,705 consecutive cancer patients admitted to the Cancer Institute in the study period. The most common cancers were breast (29.2%), colorectal (9.0%), stomach (8.3%), head & neck (8.0%) and esophageal (3.8%) cancers. Radiotherapy was the main treatment (52.1%) followed by chemotherapy (43.8%) and surgery (29.1%). We found that 60% of the patients presented in the locoregional or advanced stages. About 35% of patients travelled from other provinces mainly from Mazandaran (13.4%), Lorestan (10.6%), Zanjan (7.8%) and Ghazvin (6.6%). On average, the cancer patients travelled about 455 kilometers to receive care in the cancer institute. We found more than 38% patients who were referred from other provinces had an early stage tumor.

Conclusion: Establishment of comprehensive cancer centers in different geographical regions and implementation of a proper referral system for advanced cancer patients is needed to improve the patient outcomes and mitigate the burden of travel of patients for cancer care.

Keyword: Cancer patients, developing countries, diagnosis, Iran, travel burden, treatment

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Introduction

ancer is among the leading causes of death in both developed and developing countries. However, the burden of this lethal disease is more prominent in developing countries, where more than 80% of the world population live and the increasing trend is more striking. In spite of higher incidence rates of cancers in developed counties, the mortality to incidence ratio is considerably higher in developing countries. The disparity in cancer mortality is mainly due to detection of patients at later stages and inadequate resources for diagnosis and treatment of patients in developing countries. ^{2,3}

Access to effective and multidisciplinary approaches for diagnosis, treatment and palliative care would mitigate premature death and improve the quality of life among cancer patients. While developing countries have to deal with more than half of the world's patients, they only possess 5%-10% of the global resources for cancer treatment. 27

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The I. R. of Iran is located in the Middle East and experiences a cancer epidemic similar to other developing countries. According to Globocan 2012, about 85,000 cancers occurred in Iran, which claimed more than 50,000 died of it in 2012.8 The most common cancers in Iran include stomach, bladder, prostate and colorectal cancers among men and breast, colorectal, stomach and esophageal cancers among women. Iran is a large country and within-country disparities exist in access to diagnostic and treatment facilities. Although detailed information is not available on access to treatment, it has been reported that only 50% - 70%of patients needing radiotherapy have access to this treatment in Iran. Iran is divided into 30 provinces and high quality treatment service, including cancer surgery, chemotherapy and radiotherapy is lacking in some of these regions. Therefore, cancer patients should travel to other provinces to receive appropriate diagnosis and treatment.

The objective of this study was to study the clinical profile and referral pattern of cancer patients who were admitted to the cancer institute of Iran, the most comprehensive cancer center in the country.

Materials and Methods

Cancer Institute of Iran

The Cancer institute of Iran was established in 1949 based on collaboration between School of Medicine of Tehran University and the Red Crescent Organization of Iran. The details about history and activities of cancer institute of Iran have been

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published elsewhere.¹⁰ In summary, the cancer institute of Iran is the center of excellence for cancer in Iran, which is located in the capital city of Tehran and provides cancer care to patients referred from the entire country. Annually, about 10,000 cancer patients receive inpatient and outpatient care in different departments of the cancer institute of Iran. Several multidisciplinary tumor boards, including general, head and neck, gastroenterology, breast and sarcoma tumor boards discuss complicated patients and make decisions about the best treatment options for the presenting patients. In terms of education, the cancer institute of Iran provides residency and fellowship programs, including pathology, radiation oncology, medical oncology, cancer surgery and palliative care. In addition, nurses, radiotherapy technicians and biophysicists spend some of their training periods in the cancer institute of Iran. Three research centers including Cancer Research Center (2003), Cancer Model (Biology) Research Center (2014), and Radiation Oncology Research Center (2015) have been established in the cancer institute of Iran and coordinate cancer research in different fields including basic, epidemiology, and clinical research.

Patient information

In a case series, we reviewed archived files for all cancer patients who were successively admitted to the cancer institute of Iran between January 2012 and May 2012 and included those who fulfilled the selection criteria. Inclusion criteria were adult cancer patients who were older than 18 years old and we excluded patients with a benign tumor or non-melanoma skin cancers and those for whom the diagnosis was missing. We collected personal and clinical information of the patients including, age, sex, place of residence, cancer site, stage, and type of treatment from the information archived in the hospital.

Statistical analysis

We performed descriptive statistics to study travel burden among Iranian cancer patients. We used SSER staging method and classified the patients into four groups including 1) local (tumors are localized and there is no indication for extension outside the tumor), 2) regional (direct extension to the adjacent organ or/ and regional lymph nodes) and distant metastasis to other organs including lung, brain, bone, liver, etc., and 4) unknown stage. We used the International Classification of Diseases for Oncology (ICDO) codes to define cancer sites. Overall, we collected five months data and estimated the average number of cancer patients

admitted to the cancer institute of Iran in 2012 and applied descriptive statistics to report the frequency of different cancer types, stage of patients at diagnosis, and different treatment types including surgery, radiotherapy, chemotherapy, hormone therapy and immunotherapy. We estimated the travel distance for each patient based on the distance between their residential places to Tehran. Alborz province is located in the western part of Tehran province and recently became an independent. There is very short distance from capital city of Alborz (Karaj) to Tehran and people may commute for work or life between Tehran and Alborz and use their home or work address interchangeably in the medical records. In addition, all cancer patients travel to Tehran for diagnosis and treatment were from the capital city of Karaj. Therefore, we combined patients from Alborz province with Tehran patients in this study. Statistical analyses were performed using STATA software (Special Edition, Ver. 11.2, Stata corporation, Texas, 2009).

Results

We reviewed data from 1,705 consecutive patients who were admitted to the hospital in the first five months. We excluded 170 non-cancerous patients, 91 non-melanoma skin cancer patients and 16 patients who had no diagnostic information. From the 1428 eligible cases, 537 (38%) patients initiated their treatment in the cancer institute of Iran and the rest of them received initial treatment in other centers and were referred to our center to complete their treatment or for management of complications. Women were more frequent (N = 852, 59.7%) and the median age for all patients was 52.9 years. The most common cancer sites were breast (29.2%), colorectal (9.0%), stomach (8.3%), head & neck (8.0%) and esophageal (3.8%) cancers (Table 1).

In total, 35% of the patients were referred from other provinces, including Mazandaran (13.4%), Lorestan (10.6%), Zanjan (7.8%) and Ghazvin (6.6%) provinces for treatment in the cancer institute of Iran (Figure 1). On the average, the cancer patients had to travel about 455 kilometers to reach the cancer institute of Iran for treatment (Table 2), ranging from 132 KM travel from Ghom province to 1567 KM from Sistan-Baluchestan. Analyses by cancer sites showed that the patients were referred from other provinces for all types of cancer (Table 3). However, the main cancers referred from other provinces included melanoma, uterus, head and neck cancer, and soft-tissue sarcoma.

| Female | | Mal | e |
|---------------------|------------|------------------|-----------|
| Cancer Site | N (%) | Cancer Site | N (%) |
| Breast | 407 (47.8) | Stomach | 87 (15.1) |
| Colorectal | 53 (6.2) | Colorectal | 76 (13.2) |
| Head & Neck | 47 (5.5) | Head & Neck | 67 (11.6) |
| Cervix | 35 (4.1) | Prostate | 43 (7.5) |
| Ovary | 33 (3.9) | Esophagus | 36 (6.3) |
| Uterus | 33 (3.9) | Lymphoma | 31 (5.4) |
| Melanoma | 32 (3.8) | Soft tissue | 25 (4.3) |
| Stomach | 31 (3.6) | Lung | 22 (3.8) |
| Soft Tissue Sarcoma | 22 (2.6) | Brain | 20 (3.5) |
| Esophagus | 18 (2.1) | Hodgkin | 18 (3.1) |
| All cancer sites | 852 (100) | All cancer sites | 576 (100) |

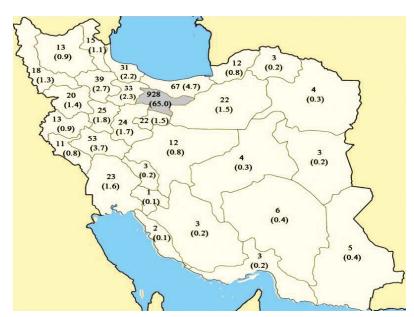


Figure 1. Number and proportion of the cancer patients who received care in the Cancer Institute of Iran by residential place from January to May 2012 (*Patients who were resident of Alborz and Tehran provinces were combined)

Table 2. The distribution and travel distance of patients who traveled from different provinces to Tehran to receive treatment in the Cancer Institute of Iran from January to May 2012

| Province* | Percentage (%) | Distance (Km) | Province | Percentage (%) | Distance (Km) | |
|-----------------|----------------|---------------|------------------------|----------------|---------------|--|
| Mazandaran | 13.40 | 267 | Esfahan | 2.40 | 414 | |
| Lorestan | 10.60 | 499 | Golestan | 2.40 | 381 | |
| Zanjan | 7.80 | 430 | Ilam | 2.20 | 723 | |
| Ghazvin | 6.60 | 150 | Kerman | 1.20 | 1064 | |
| Gilan | 6.20 | 325 | Sistan-Baloochestan | 1.00 | 1567 | |
| Hamadan | 5.00 | 336 | Khorasan Razavi | 0.80 | 894 | |
| Markazi | 4.80 | 288 | Yazd | 0.80 | 677 | |
| Khoozestan | 4.60 | 874 | Chaharmahal-Bakhtiari | 0.60 | 521 | |
| Semnan | 4.40 | 228 | South Khorasan | 0.60 | 1313 | |
| Ghom | 4.40 | 132 | North Khorasan | 0.60 | 713 | |
| Kordestan | 4.00 | 486 | Fars | 0.60 | 895 | |
| West Azarbayjan | 3.60 | 946 | Hormozgan | 0.60 | 1501 | |
| Ardabil | 3.00 | 588 | Booshehr | 0.40 | 1215 | |
| East Azarbayjan | 2.60 | 624 | Kohgilooyeh-Boyerahmad | 0.20 | 738 | |
| Kermanshah | 2.60 | 526 | Weighted Average | | 455 | |

Table 3. Frequency of different cancer types that were admitted for treatment in January–May, by residential place and annual estimation in the Cancer Institute of Iran in 2012

| Cancer type | | 1st January-31st May | | | | |
|----------------------------|---------|--|-------------|------|--|--|
| | Overall | Overall Tehran*, N (%) Other provinces N (%) | | | | |
| Breast | 417 | 313 (75.1%) | 104 (24.9%) | 1000 | | |
| Colorectal | 129 | 92 (71.3%) | 37 (28.7%) | 310 | | |
| Stomach | 118 | 70 (59.3%) | 48 (40.7%) | 282 | | |
| Head & Neck | 114 | 58 (50.9%) | 56 (49.1%) | 274 | | |
| Esophagus | 54 | 34 (63.0%) | 20 (37.0%) | 130 | | |
| Melanoma | 48 | 23 (47.9%) | 25 (52.1%) | 115 | | |
| Soft Tissue Sarcoma | 47 | 26 (55.3%) | 21 (44.7%) | 113 | | |
| Prostate | 43 | 34 (79.1%) | 9 (20.9%) | 103 | | |
| Cervix | 35 | 22 (62.9%) | 13 (37.1%) | 84 | | |
| Uterus | 33 | 16 (48.5%) | 17 (51.5%) | 79 | | |
| Ovary | 33 | 27 (81.8%) | 6 (18.2%) | 79 | | |
| Other** | 357 | 213 (60%) | 144 (40%) | 857 | | |
| Excluded patients*** | 277 | 181 (65.0%) | 97 (35.0%) | 665 | | |
| All, but excluded patients | 1428 | 928 (65.0%) | 500 (35.0%) | 3427 | | |
| Overall | 1705 | 1109 (65.0%) | 597 (35.0%) | 4092 | | |

^{*}Patients who were resident of Alborz and Tehran provinces were combined; **Patients with unknown primary site were combined in the "other" category;
***Excluded patients consist of non-cancer patients (n=170%), non-melanoma skin cancer (n=91%), and patients with missing diagnosis information (n=16%).

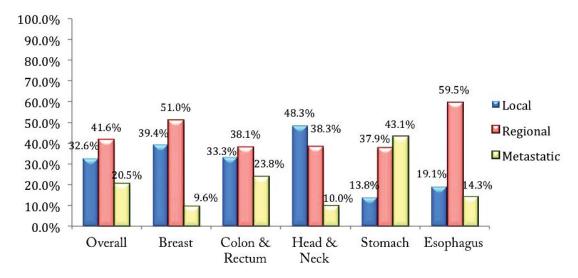


Figure 2. Stage distribution of different cancer types of cancer patients who were admitted to the Cancer Institute of Iran from January to May 2012 (Only 537 patients who received initial treatment in the cancer institute were analyzed for this figure)

Table 4. Stage distribution of different cancer types of cancer patients who initiated their cancer treatment in the Cancer Institute of Iran stratified by residential place from January to May 2012, (Only 537 patients who received initial treatment in the cancer institute were analyzed for this Table).

| Cancer type — | Tehran | | | | Other provinces | | | | | |
|------------------|------------|-------------|------------|-----------|-----------------|------------|------------|------------|-----------|------------|
| | Local | Regional | Metastatic | Unknown | Total | Local | Regional | Metastatic | Unknown | Total |
| Breast | 28 (37.4%) | 40 (53.3%) | 7 (9.3%) | 0 (0.0%) | 75 (100%) | 13 (44.8%) | 13 (44.8%) | 3 (10.3%) | 0 (0.0%) | 29 (100%) |
| Colorectal | 15 (34.1%) | 15 (34.1%) | 12 (27.3%) | 2 (4.5%) | 44 (100%) | 6 (31.6%) | 9 (47.4%) | 3 (15.8%) | 1 (5.2%) | 19 (100%) |
| Stomach | 3 (9.7%) | 12 (38.7%) | 16 (51.6%) | 0 (0.0%) | 31 (100%) | 5 (18.5%) | 10 (37.1%) | 9 (33.3%) | 3 (11.1%) | 27 (100%) |
| Head & Neck | 15 (50.0%) | 10 (33.3%) | 3 (10.0%) | 2 (6.7%) | 30 (100%) | 14 (46.7%) | 13 (43.3%) | 3 (10.0%) | 0 (0.0%) | 30 (100%) |
| Esophagus | 6 (20.7%) | 16 (55.2%) | 5 (17.2%) | 2 (6.9%) | 29 (100%) | 2 (15.4%) | 9 (69.2%) | 1 (7.7%) | 1 (7.7%) | 13 (100%) |
| All Cancer Types | 98 (29.2%) | 144 (43.0%) | 78 (23.3%) | 15 (4.5%) | 335 (100%) | 77 (38.1%) | 79 (39.1%) | 32 (15.9%) | 14 (6.9%) | 202 (100%) |

More than 60% of the patients presented at loco-regional or advanced stages. More than 30% of the breast, colorectal, and head and neck cancers were diagnosed with a local tumor. However, about 40% of the stomach cancers were diagnosed at the advanced stage (Figure 2). We found that more than 38% of referral patients from other provinces who initiated their cancer treatment in the cancer institute of Iran had an early stage tumor (Table 4). In fact, almost there was no differences in the stage distributions of the patients who came from Tehran and those who were referred from other provinces.

Radiotherapy was the main treatment (52.1%) followed by chemotherapy (43.8%) and surgery (29.1%) and hormone therapy or immunotherapy were prescribed for about 10% of the patients (Figure 3).

Discussion

In this paper, we reported the demographic and clinical characteristics as well as the type of treatment provided for cancer patients who were admitted to the cancer institute of Iran in 2012. Breast and stomach cancers were the main cancer types treated in 2012. About 65% of patients were residents of Tehran, and 35% of the patients travelled about 455 kilometers from different

provinces, mainly Mazandaran (13.5%), Lorestan (10.7%), Zanjan (7.9%), Ghazvin (6.7%) and Gilan (6.2%) to receive care in the cancer institute of Iran.

The cancer patients were referred from different cities for treatment because of several reasons. The most important reason for referring to the cancer institute of Iran is lack of an organized referral system in the country and patients may prefer to initiate their treatment in the cancer institute of Iran or come to this hospital to complete their treatment or for management of complications from previous treatments. In addition, the cancer institute of Iran is a high volume hospital for surgery, radiotherapy, and chemotherapy. Some cancers, like soft tissue sarcoma and melanoma that need critical decisions and complicated care, may generally refer to the cancer institute of Iran for initial treatment, while other cancers, like breast and prostate cancers, may be referred if the initial treatment fails. Finally, some patients may be advised to travel to Tehran for their treatment because treatment facilities such as cancer surgery, chemotherapy or radiotherapy are not available in their residential place. Although we reported that patients had travelled a long journey for treatment in the cancer institute of Iran, they may not complete treatment because of the long duration of chemotherapy and radiotherapy for most cancer patients and the high economic and social burden for

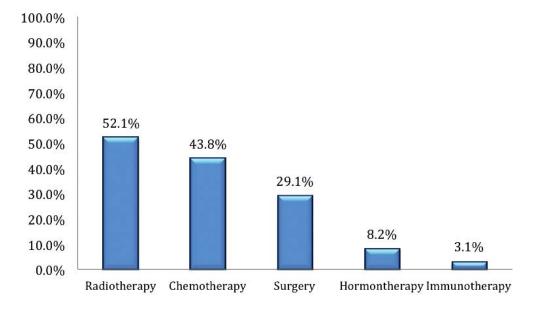


Figure 3. Treatment modality of cancer patients who were admitted for treatment to the Cancer Institute of Iran from January to May 2012

patients and their relatives who may temporary stay in the capital city of Tehran or frequently travel between the cancer institute of Iran and their hometown. Travel burden may lead to delayed or incomplete treatment for cancer patients and compromise the efficiency and survival of these patients.

We found that more than 38% referral patients from other provinces that initiated their cancer treatment in the cancer institute of Iran had an early stage tumor. These results indicate that many patients can be treated in their hometown and it is not necessary to travel for treatment and a referral system needs to be defined for cancer patients. Many studies have reported that distance could be an important barrier to cancer care. 12-15 Increasing travel requirements may cause delay in the diagnosis of cancer, and progression of tumors to an advanced stage, thereby reducing patient survival.¹⁶ In addition, it will increase economic burden to patients and the society.^{17,18} Previous studies have reported professional delays in diagnosis and treatment of oral cancer²⁰ and leukemia. ¹⁹ COSA (2006) found large disparities in cancer outcomes between people living in metropolitan versus non-metropolitan regions in Australia.²⁰ Likewise, cancer patients in remote areas were sub-optimal for continuing their cancer treatment such as adjuvant radiotherapy, hormonal therapy and conserving surgery²¹ and did not seek formal supportive care.²²

Although breast cancer and stomach cancer were respectively the most common cancers among male and female patients who were admitted for diagnosis and treatment in the cancer institute of Iran, the relative frequencies of different cancer types were not completely matched with the pattern of cancer incidence in the country. The cancer institute of Iran admits a large number of head and neck cancers, malignant melanomas, and sarcomas that are not common cancers in Iran.⁸ In addition, radiotherapy was the main treatment provided to the cancer patients in the cancer institute of Iran due to the large radiotherapy department, which provides radiotherapy service for patients referred from the cancer institute and other hospitals. After excluding non-melanoma skin cancers, we estimated that this center annually admits about 400 head and neck cancer patients for treatment. Although the incidence rate of

head and neck cancer is not high in Iran, the high volume of head and neck cancers in the cancer institute of Iran could be due to the fact that these patients need a multidisciplinary approach, which is not available in most hospitals, in particular in other provinces. ²³ As recommended by the National Comprehensive Cancer Network (NCCN),²⁴ a multidisciplinary team including oncosurgeons, otorhinolaryngologists with subspecialty in oral and maxillofacial surgery, pathologists, and oncologists provide high quality care to these patients in the cancer institute of Iran. Previous studies have reported a positive association between volume of hospitals and patient survival for head & neck cancer.25-27 In addition, the cancer institute of Iran admits a considerable number of patients with soft-tissue sarcoma and malignant melanoma that are relatively rare and should be managed in high volume centers by a multidisciplinary approach.^{28,29} In the UK, the Calman-Hine report recommended that treatment for less common cancers and where treatment is technically demanding should be offered in regional cancer centers, where there are site-specialist and multidisciplinary teams.³⁰

The incidence rate of cancer is rising in the world, in particular in low-middle income counties,³¹ indicating that the number of patients and waiting time will increase in referral hospitals like the cancer institute of Iran.³² Yun, et al. showed that the waiting time is higher at high-volume hospitals compared to low- to medium-volume hospitals.³³ Iran and other middle-income countries should establish cancer centers in different parts of the country and be prepared for the increasing number of cancers in the coming years. Providing high quality cancer care in different regions would reduce the burden of patients' trip from remote areas to large cities like Tehran.

The limitation of this study was that we used retrospective data from hospital files. It is important to conduct a prospective survey among cancer patients and figure out the reason for traveling from different parts of the country. In addition, we may need to analyze the outcome of patients treated in the cancer institute of Iran compared to those who stayed in their hometown and evaluate the benefits and harms of these travels for cancer patients. In addition,

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it will also be interesting to compare the pattern of treatment in different hospitals and regions in the country. Unfortunately, there is no hospital-based cancer registry in Iran. Establishment of hospital-based cancer registries in different hospitals may help in regular monitoring of the pattern of care and patients' outcome in the country.⁸

In conclusion, we presented data on cancer patients who were referred to the cancer institute of Iran. The cancer institute of Iran admits patients from the entire country, in particular from the neighboring regions and the provinces that lack appropriate infrastructure for diagnosis and treatment of the patients. The cancer institute of Iran remains the main referral center in the country for diagnosis and treatment of rare and advanced cancers, which require high specialty and sophisticated care. While treatment of complicated patients and specific cancer types should be still undertaken centrally in large and high volume hospitals like the cancer institute of Iran, establishment of cancer hospitals and providing essential care to patients in different parts of the country is necessary. Further evidence on pattern of care and consequences of the travel burden for cancer patients in Iran and other low- and middle-countries is warranted.

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