Original Article

Therapeutic itinerary of colorectal cancer patients treated in the state of Rio Grande do Sul

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\textbf{A B S T R A C T}

Objectives: The objective of the study was to evaluate the therapeutic itinerary of patients treated in a specialized center, including its trajectory in seeking treatment and their clinical and epidemiological characteristics.

Methods: This is a cross-sectional prospective descriptive study; patients with colorectal cancer aged over 18 years and who signed the consent form were included in the analysis. Tumor characteristics, such as staging and tumor features; epidemiological characteristics such as age, gender, profession and itinerary in the form of the number of clinical visits needed to obtain the diagnosis; the main symptoms; and mean time between diagnosis and onset of treatment were evaluated.

Results: 34\% of patients initially sought primary care (first level of care in the public health system) and 50\% were diagnosed in secondary care (second level of care in the public health system); the mean number of visits until obtaining a diagnosis was 2.5 times; and 52\% of patients received palliative therapy and 40\% had a stage IV diagnosis.

Conclusions: The mean time between the diagnosis and the onset of treatment is in line with the recommendations of the Ministry of Health of Brazil. However, it was found that the patients are unaware of the symptoms of the disease, since the mean time between the onset of symptoms until the decision to visit the doctor was 177 days – which may have been a determining factor for a diagnosis in an advanced stage of disease.

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Itinerário terapêutico de pacientes com câncer colorretal tratados no interior do Rio Grande do Sul

RESUMO

Objetivos: O objetivo do estudo foi avaliar o itinerário terapêutico de pacientes tratados em um centro especializado, compreendendo sua trajetória na busca do tratamento e as características clínicas e epidemiológicas.

Métodos: estudo descritivo prospectivo transversal; foram incluídos na análise pacientes portadores de câncer colorretal (CCR) maiores de 18 anos que assinaram o termo de consentimento. Foram avaliadas características tumorais, como estadiamento e características do tumor; características epidemiológicas como idade, sexo e profissão e a trajetória como número de consultas até o diagnóstico, principais sintomas e tempo médio entre o diagnóstico e início do tratamento.

Resultados: 34% dos pacientes procuraram inicialmente a atenção primária (primeiro nível de atenção na saúde pública), 50% receberam o diagnóstico na atenção secundária (segundo nível de atenção na saúde pública), a média de consultas até o diagnóstico foi de 2,5 vezes, 52% dos pacientes fizeram terapia paliativa e 40% fizeram diagnóstico em estádio IV.

Conclusões: o tempo médio entre o diagnóstico e o início do tratamento está de acordo com o recomendado pelo Ministério da Saúde do Brasil; entretanto, identificou-se que os pacientes desconhecem os sintomas da doença, uma vez que a média de tempo entre o início dos sintomas até a tomada de decisão de procurar um médico foi de 177 dias, o que pode ter sido determinante para um diagnóstico na fase avançada da doença.

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Introduction

Colorectal cancer (CRC) is the third most common malignancy in humans. This disease is considered a serious public health problem because of its huge socioeconomic impact on society and by requiring an individualized, long, exhausting and expensive treatment. The low income and lack of health policies to alert the population are crucial for an early diagnosis.1

As to 2014, the National Cancer Institute (INCA) estimated the occurrence of 32,600 new cases, 15,070 in men and 17,530 in women, accounting for 15.44 new cases per 100,000 men and 17.24 new cases per 100,000 women. In the South region, CRC is the third most common cancer in men (20.43/100,000) and the second most common in women (21.85/100,000). In 2011, 14,016 deaths (6818 men and 7198 women) were recorded in Brazil.2

These alarming data can be directly related to a late diagnosis of the disease and mainly related to the difficulty to offer an adequate care. The therapeutic itinerary is a way to know the trajectory of the disease process in seeking medical attention in such a way that the patient and his/her family can understand. In health care services difficulties can arise in the field of organization, which is a barrier to the provision of a more effective care for patients.3

Understanding of the timing and how individuals seek health care is a way to organize and evaluate health systems with a view to conducting a planning of the critical points.4 Studies involving knowledge of the therapeutic itinerary of patients with CRC are extremely important, but such papers are still scarce.5

Based on data in the literature and in the face of the few studies published on this topic in oncology, the aim of this study was to evaluate the therapeutic itinerary of patients with CRC treated in a specialized center, as a way to understand the trajectory of each patient and his/her clinical and epidemiological characteristics.

Methodology

This is a prospective, descriptive cross-sectional study which was conducted at the Integrated Oncology Center, Hospital Ana Nery, located in Santa Cruz do Sul/RS. This hospital is a reference in cancer treatment by the Unified Health System (Sistema Unificado de Saúde – SUS), and its database is connected to the National Cancer Institute (INCA). The hospital covers about 820,000 people in the regions of Vale do Rio Pardo, Central Serra, and Carbonifera. On average, the Integrated Oncology Center serves 950 chemotherapy patients (94% by SUS). In radiotherapy, the Center serves about 100 patients per month (84.7% SUS). The data collection began in November 2012 and was completed in April 2014. This study was approved by the Research Ethics Committee (CEP) of the Universidade de Santa Cruz do Sul.

Patients with CRC with confirmation of the disease by biopsy, aged over 18 years who and signed a free and informed consent were included in the survey. An interview face-to-face took place and, by this instrument, the participants were asked about epidemiological information such as age, gender (male/female), skin color (white/non-white), family income (number of minimum wages), profession, marital...
status (accompanied/unaccompanied), diabetes (yes/no), family history of CRC (yes/no), family history of other types of cancer (yes/no), time between onset of symptoms and the first health service sought (days), type of health care service sought for the first consultation (primary, secondary or tertiary care), number of medical visits to establish the diagnosis, and service where the diagnosis was carried out (primary, secondary or tertiary care). During the interview, the patients were informed that the data would be used for scientific research, without any personal return.

In addition to the interview, a review of medical records was performed. The data from medical records were evaluated and transcribed to a data collection form previously prepared. In the medical records, we evaluated the primary site of disease (colon/rectum), date of onset of treatment and stage of development according to the TNM classification, issued by the International Union for Cancer Control (UICC), where T refers to the extent of the tumor, N to the absence or presence of metastases in regional lymph nodes, and M to the absence or presence of distant metastases. Adding numbers to each of those letters can also indicate the extent of malignancy.6

**Statistical analysis**

All information was encoded and stored anonymously in a database created for this purpose. Clinical and epidemiological data were stored in a database created in the software Statistical Package for Social Sciences (SPSS), version 20.0 (SPSS Inc., Chicago, IL) for statistical analysis. Our results were described using descriptive statistics (absolute numbers, frequencies and means).

**Results**

In total, 50 subjects participated in this study, with a mean age of 56 (±12.7) years. Table 1 lists the epidemiological characteristics of the patients. It was found that most patients were Caucasian and had a family income of up to 3 minimum wages. Among the professions cited by patients, the most frequent were agriculture (Table 1).

Regarding the tests for the diagnosis of CRC, 38 (76%) participants had never been submitted to, or did not know, colonoscopy. 48% had never been screened for occult blood in the stool.

**Table 1 – Epidemiological characteristics of patients with colorectal cancer.**

<table>
<thead>
<tr>
<th>Characteristic</th>
<th>n</th>
<th>%</th>
</tr>
</thead>
<tbody>
<tr>
<td>Male gender</td>
<td>28</td>
<td>56</td>
</tr>
<tr>
<td>White skin color</td>
<td>39</td>
<td>78</td>
</tr>
<tr>
<td>Income (up to 3 minimum wages)</td>
<td>48</td>
<td>96</td>
</tr>
<tr>
<td>Profession (farmers)</td>
<td>19</td>
<td>38</td>
</tr>
<tr>
<td>Marital status (accompanied)</td>
<td>36</td>
<td>72</td>
</tr>
<tr>
<td>Family history of CRC</td>
<td>09</td>
<td>18</td>
</tr>
<tr>
<td>Family history of other types of cancer</td>
<td>22</td>
<td>44</td>
</tr>
</tbody>
</table>

Among the symptoms causing the visit of patients to the health service, abdominal pain, and weight loss were the most frequently cited (Table 2).

The mean time between the onset of symptoms and the decision to seek a health service was 177 ± 336.88 days, ranging from 2 to 2160 days and with a median of 60 days. On average, each patient visited the doctor at least 2.5 (±0.6) times between the initial consultation and the establishment of the diagnosis. The mean time between the first consultation with the chemotherapy service and the onset of treatment was 32 (±33.0) days. Among the patients studied, 13 (26%) were being medicated simultaneously with radiotherapy and chemotherapy.

In Fig. 1, one can see the first type of service sought by patients after making the decision to seek a doctor. Regarding the service responsible for the diagnosis of cancer, it was found that in 50% of cases this diagnosis was confirmed in a secondary care service. Fig. 2 shows the trajectory of each patient.

In Table 3, the characteristics related to the tumor and the type of treatment. 52% of patients received palliative therapy and 40% were diagnosed with a stage IV cancer.
Table 3 - Clinical characteristics of patients.

<table>
<thead>
<tr>
<th>Characteristic</th>
<th>n</th>
<th>%</th>
</tr>
</thead>
<tbody>
<tr>
<td>Site of the primary tumor – colon</td>
<td>32</td>
<td>64</td>
</tr>
<tr>
<td>Staging</td>
<td></td>
<td></td>
</tr>
<tr>
<td>II</td>
<td>16</td>
<td>32</td>
</tr>
<tr>
<td>III</td>
<td>13</td>
<td>26</td>
</tr>
<tr>
<td>IV</td>
<td>20</td>
<td>40</td>
</tr>
<tr>
<td>Purpose of treatment</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Previous</td>
<td>02</td>
<td>04</td>
</tr>
<tr>
<td>Adjuvant</td>
<td>21</td>
<td>42</td>
</tr>
<tr>
<td>Palliative</td>
<td>26</td>
<td>52</td>
</tr>
<tr>
<td>Without information</td>
<td>01</td>
<td>02</td>
</tr>
<tr>
<td>Pathology</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Well differentiated</td>
<td>05</td>
<td>10</td>
</tr>
<tr>
<td>Moderately differentiated</td>
<td>31</td>
<td>62</td>
</tr>
<tr>
<td>Poorly differentiated</td>
<td>02</td>
<td>04</td>
</tr>
<tr>
<td>Invasive</td>
<td>06</td>
<td>12</td>
</tr>
<tr>
<td>No information</td>
<td>06</td>
<td>12</td>
</tr>
</tbody>
</table>

Discussion

CRC is a prevalent malignancy worldwide, affecting mostly rich countries. In Brazil, there is a high incidence in the South and Southeast regions, especially in older individuals and in males.

In this study, the mean age was 56 years, which is in line with the literature findings. The exact cause of CRC is unknown, but there is strong involvement of genetic and environmental characteristics. Currently, clinical and biological factors have been studied in order to better understand the progression of the disease.

Malignancies have been associated with the use of pesticides. The chronic effect of these diseases occurs through the respiratory, dermal and digestive tract. Pesticides which present organochlorinated compounds in their formulation are retained in tissues, and some studies have shown that such agents are associated with malignancies. In this study, a high prevalence of farmers (38%) diagnosed with CRC was observed. The regions where this study was conducted have an economy based on tobacco and rice farming, which explains the high frequency of individuals who report farming as a professional activity. As described in a study conducted in São Paulo, the second largest related professional activity was also agriculture.

In this study, 18% of patients reported a family history of CRC. These figures are close to those found in the literature, suggesting that in 20–25% of cases one can observe a family component to CRC.

We observed in this study that patients are not aware of preventive tests such as colonoscopy and fecal occult blood, since 76% of patients had never been submitted to colonoscopy and 48% never had a fecal occult blood test. These figures are worrying because that mortality from CRC decrease and that the survival of individuals affected by this disease increase, conducting screening tests are the most effective methods.

In our study the most frequent symptoms were abdominal pain (60%) weight loss (66%) and blood in the stool (46%), which is in line with the findings described by El-Halabi et al. (2014). These authors report that weight loss, obstruction, occult blood in the stool, abdominal pain and palpable abdominal mass are common in all patients.

These data are worrying, given that the most effective method for reducing mortality from CRC and increasing survival of individuals affected by this disease is to perform screening tests. In addition to these tests, genetic tests have been developed and used as predictive and prognostic biomarkers for CRC. Knowing the therapeutic itinerary is to understand the perception of each patient and how each one behaves in the face of a certain situation. We noted that, on average, the time...
elapsed since the onset of symptoms to the decision to seek a health service was 177 (±336.88) days. This extended time is not only due to a lack of knowledge of patients, but during their questioning it was possible to identify the private fears and anxieties, such as lack of time, overwork, the fear of a diagnosis without healing chances and even the simple fact that they feel that certain symptoms would be considered normal. Therefore, the cultural and social factors, life experiences and beliefs make these individuals opt for different ways to try to solve their health problems; thus, they may resort to a specialized support or to self-treatment, resulting in a delay in their demand for medical help and, as a consequence, with a late diagnosis. Some studies have shown that some people consult the doctor as soon as the first signs and symptoms of the disease appear; on the other hand, other people with similar symptoms do not proceed in this manner, because of financial difficulties to get professional help. The time elapsed since the onset of symptoms until the first consultation is critical, because the tumors already are in an advanced T stage. The result of a late diagnosis is the palliative chemotherapy, with the tumor already in stage IV.

In this study, 34% of patients had their first consultation in primary care and 50% were diagnosed in secondary care, after two to three visits on average. These data show that the health system is not a fragmented body, and can communicate to pass along the information of its users, which facilitates the diagnosis after the patient made the decision to seek medical advice, as opposed to the findings described in other studies, i.e. Mendel et al. (2010) in which the authors report the precariousness of public health services and the lack of communication of health networks.

The mean time since the first consultation at the health service until the beginning of treatment was 32 days, which is in accordance with a Brazilian law approved in 2012 for patients with proven malignancy. This law establishes that every cancer patient should receive the first treatment within 60 days after the diagnosis confirmed by a medical report.

In their study, Arribas et al. (2014) found that in 78% of the patients studied the primary tumor site was the colon, and that 32.92% of patients had a pathology report of a well-differentiated tumor. The tumor stage at diagnosis is crucial to determine the purpose of the treatment. Very important that the doctor is aware of the pathology report (tumor characteristics), because the tumor acts as a prognostic marker, or indicates a worst (or better) result, regardless of treatment. Thus, one can evaluate each individual and choose an individualized therapy, according to the pathological features. The biological behavior of the tumor, as well as factors that induce angiogenesis, may be considered decisive in the choice of treatment and in predicting the individual’s response.

### Conclusion

We conclude that health systems are not fragmented because, on average each patient had two visits, a number that is considered appropriate. This shows that the services where the study was carried out were able to communicate so that the patient has a rapid diagnosis. Patients are not knowledgeable about screening tests; therefore, it is clear the lack of campaigns warning on the need for an early diagnosis of CRC, since the mean time between the onset of symptoms and the decision to visit a doctor was of 177 days – and this may have been decisive for a low perspective of cure, considering that 40% of subjects were in stage IV and 52% were referred to palliative chemotherapy.

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### Conflicts of interest

The authors declare no conflicts of interest.

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