Factors Associated with Delayed Diagnosis of Cervical Cancer in Iran - a Survey in Isfahan City

Fariba Behnamfar, Mahboobeh Azadehrah*

Abstract

Background: In the absence of routine screening program for cervical cancer in Iran and high rate of diagnosed cancer in its advanced stage, recognition of sociodemographic factors related to delayed diagnosis of cancer in Iran could be helpful in reducing the burden of disease in our community. The aim of this study was to determine the stage of cervical cancer at diagnosis and factors related to delayed diagnosis of cervical cancer in Isfahan, Iran.

Materials and Methods: In this cross sectional study women diagnosed with cervical cancer for the first time by histo-pathological examination were enrolled. According to the clinical and paraclinical findings and staging of the cancer, they were classified into early and delayed diagnosis of cervical cancer. Sociodemographic factors were compared in the two groups.

Results: In this study of 55 women mean age was 48.3±12.0. According to our classification 6/55 (10.9%) and 49/55 (89.1%) of them had early and delayed diagnosis of cervical cancer. Delayed diagnosis of the cancer was significantly higher in patients with lower degree of education, lower socioeconomic status, having smoker and addict husband and those who did not have a history of Pap smear test (p<0.05).

Conclusions: The results of this study indicated risk factors related to delayed diagnosis of cervical cancer. The affected women should be targeted for implementation of specialized educational programmes for improving knowledge and screening test.

Keywords: Cervical cancer - pap smear - diagnosis - delay - Isfahan, Iran

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Introduction

Cervical cancer with an estimated 529,800 new cases worldwide is considered as one of the most common cancer among women worldwide. It constitutes approximately 8% of the global burden of cancer among women (Ferlay et al., 2010; Arbyn et al., 2011). More than 85% of the cases have reported in developing countries (Jemal et al., 2011).

The estimated age-standardized incidence rate (ASR) of cervical cancer in Iran have reported to be 2.5 per 100,000 in pathology-based cancer registries and the mortality to incidence ratio was 42% according to a recent study in Iran (Khorasanizadeh et al., 2013).

Cervical cancer is a curable and preventable disease if it diagnosed in the early stage of malignancy i.e at the precancerous stage (Devi et al., 2007). Whereas advanced stage of the disease has poor prognosis and is correlated with lower survival rates. Advanced cervical cancer is one of the major causes of cancer related mortality in women specially in low- and medium-income countries mostly due to poor access to appropriate management (Vinh-Hung et al., 2007; Thomson and Forman, 2009).

Recently the concept of delayed diagnosis has become as an important issue in the cancer prevention and treatment. The concept is categorized in four components including patient delay, health care provider delay, referral delay and system delay. Though all of the mentioned delays have important role in the prevention, diagnosis and management of the disease but it seems that in our country and other developing countries patients and health care providers delay have more crucial role (Hansen et al., 2008; Berraho et al., 2012; Gwynwali et al., 2014).

Several studies worldwide have investigated the factors associated with delayed diagnosis of the cancer and disparities in its mortality rate in different racial, geographic and socio-economic groups (Allgar et al., 2005; Brewer et al., 2005; Yu et al., 2005; Ibfelt et al., 2012). However knowledge of delays for this cancer could be useful in establishing comprehensive preventative strategies.

Though the incidence rate of cervical cancer is low in Iran (Arbyn et al., 2011) but the mortality to incidence ratio is high which indicates that most of the cancer cases are diagnosed in advanced stages. Factors such as unavailability to routine screening, inadequate follow-up of abnormal Pap smears and possibly low awareness of women’s population regarding the course of the disease could explain the higher mortality of the cancer in our community (Khorasanizadeh et al., 2013). So, in the
absence of routine screening program for cervical cancer in Iran and high rate of diagnosed cancer in its advanced stage, recognition of socio-demographic factors related to delayed diagnosis of cancer in Iran could be helpful in eliminating the burden of disease in our community. The aim of this study is to determine factors related to delayed diagnosis of cervical cancer in Isfahan, Iran.

Materials and Methods

This study designed as cross sectional study with qualitative approach. In this study women aged 30-60 years diagnosed with cervical cancer for the first time by histo-pathological examination of cervix referred to three obstetrics and gynecology specialized cancer clinics in three hospitals of Isfahan during March 2013-March 2014 were enrolled.

The protocol of study was approved by Regional Bioethics Committee of Isfahan University of Medical Sciences. Written informed consent was obtained from all selected patients.

Inclusion criteria were all women who diagnosed with cervical cancer based on histopathological examination of cervix (conization or biopsy). Women with diagnosed cancer other than cervical cancer were excluded.

Table 1. Demographic Factors and Cancer Related Factors in Women with Early and Delayed Diagnosis of Cervical Cancer in Isfahan, Iran

<table>
<thead>
<tr>
<th>Variables</th>
<th>Women with early diagnosis of cervical cancer (n=6)</th>
<th>Women with delayed diagnosis of cervical cancer (n=49)</th>
<th>p value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age of marriage(years)</td>
<td>17.16±3.37</td>
<td>16.08±4.36</td>
<td>&gt;0.05</td>
</tr>
<tr>
<td>Frequency of marriage in patients</td>
<td>1.16   0.40</td>
<td>1.12   0.33</td>
<td>&gt;0.05</td>
</tr>
<tr>
<td>Frequency of marriage in husband</td>
<td>1.0    0.0</td>
<td>1.12   0.33</td>
<td>&gt;0.05</td>
</tr>
<tr>
<td>Parity</td>
<td>4.5   2.50</td>
<td>4.76   2.46</td>
<td>&gt;0.05</td>
</tr>
<tr>
<td>Gravity</td>
<td>3.0   1.26</td>
<td>4.32   2.40</td>
<td>&gt;0.05</td>
</tr>
<tr>
<td>Education of patient</td>
<td>Illiterate</td>
<td>0</td>
<td>15 (30.63)</td>
</tr>
<tr>
<td></td>
<td>Elementary-diploma</td>
<td>4 (66.7)</td>
<td>24 (48.97)</td>
</tr>
<tr>
<td></td>
<td>Higher than diploma</td>
<td>2 (33.3)</td>
<td>10 (20.40)</td>
</tr>
<tr>
<td>Education of husband</td>
<td>Illiterate</td>
<td>1 (16.7)</td>
<td>16 (32.65)</td>
</tr>
<tr>
<td></td>
<td>Elementary-diploma</td>
<td>3 (50)</td>
<td>25 (51.02)</td>
</tr>
<tr>
<td></td>
<td>Higher than diploma</td>
<td>2 (33.3)</td>
<td>8 (16.32)</td>
</tr>
<tr>
<td>Smoking</td>
<td>Patients</td>
<td>0 (0%)</td>
<td>3 (6.12%)</td>
</tr>
<tr>
<td></td>
<td>Husbands</td>
<td>1 (16.7%)</td>
<td>32 (65.3%)</td>
</tr>
<tr>
<td>Addiction</td>
<td>Patients</td>
<td>0 (0%)</td>
<td>3 (6.12%)</td>
</tr>
<tr>
<td></td>
<td>Husbands</td>
<td>1 (16.7%)</td>
<td>21 (42.85%)</td>
</tr>
<tr>
<td>Place of residence</td>
<td>Urban</td>
<td>6 (100)</td>
<td>37 (75.52)</td>
</tr>
<tr>
<td></td>
<td>Rural</td>
<td>0 (0)</td>
<td>12 (24.48)</td>
</tr>
<tr>
<td>Socioeconomic condition</td>
<td>Low</td>
<td>3 (50)</td>
<td>3 (6.12)</td>
</tr>
<tr>
<td></td>
<td>Moderate</td>
<td>3 (50)</td>
<td>8 (16.32)</td>
</tr>
<tr>
<td></td>
<td>High</td>
<td>0 (0)</td>
<td>38 (77.56)</td>
</tr>
<tr>
<td>Health insurance</td>
<td>Non</td>
<td>1 (16.7)</td>
<td>4 (8.16)</td>
</tr>
<tr>
<td></td>
<td>Rural</td>
<td>0 (0)</td>
<td>9 (18.36)</td>
</tr>
<tr>
<td></td>
<td>others</td>
<td>5 (83.3)</td>
<td>36 (73.46)</td>
</tr>
<tr>
<td>History of pap smear</td>
<td>Never</td>
<td>2 (33.3)</td>
<td>28 (57.14)</td>
</tr>
<tr>
<td></td>
<td>During last 3 years</td>
<td>2 (33.3)</td>
<td>14 (28.57)</td>
</tr>
<tr>
<td></td>
<td>More than 3 years age</td>
<td>2 (33.3)</td>
<td>7 (14.29)</td>
</tr>
<tr>
<td>Signs</td>
<td>Abnormal pap smear</td>
<td>4 (66.6)</td>
<td>0 (0)</td>
</tr>
<tr>
<td></td>
<td>Abnormal vaginal bleeding</td>
<td>0 (0)</td>
<td>28 (57.14)</td>
</tr>
<tr>
<td></td>
<td>Post-menopausal bleeding</td>
<td>1 (16.7)</td>
<td>12 (24.48)</td>
</tr>
<tr>
<td></td>
<td>Discharge</td>
<td>1 (16.7)</td>
<td>10 (20.40)</td>
</tr>
<tr>
<td>Histopathology</td>
<td>Squamous cell carcinoma</td>
<td>6 (33.3)</td>
<td>41 (83.67)</td>
</tr>
<tr>
<td></td>
<td>Adenocarcinoma</td>
<td>0 (0)</td>
<td>7 (14.28)</td>
</tr>
<tr>
<td></td>
<td>Adenosquamous carcinoma</td>
<td>0 (0)</td>
<td>1 (2.04)</td>
</tr>
</tbody>
</table>

All selected patients recalled and their medical files from mentioned three clinics were referred to the oncology clinics of Shahid Beheshti hospital for more investigation. Written informed consent was obtained from all participants after describing the aim and details of the method of the study.

Recalled patients examined clinically by gynecologist (MA). A complete pelvic and lymph nodes (axillary, supraclavicular, inguinal, femoral) examination was performed by the gynecologist in order to determine the metastatic cases. The stage of cervical cancer at diagnosis was defined according to International Federation of Gynecology and Obstetrics (FIGO) system of staging of gynecological cancer (Benedet, et al., 2000). Chest X-ray was done in all studied patients. Other complementary radiological tests were done according to the condition of the patients. If patients had a history of participating in cervical cancer screening program, if their histopathologic smears were available, they reviewed once more. According to the clinical and paraclinical findings the stage of cancer was determined in each patient. Basis of the stage of the cervical cancer at diagnosis, stage IA-1 were defined as early diagnosis and stage IA-2 and above were considered as late or delayed diagnosis. During the patients visit and after clinical examination, they interviewed by trained
nurses for following information; age, marital statuses (age of marriage and frequency), degree of education (no education, elementary/diploma, high education), smoking (patient and her husband), addiction (and her husband), socioeconomic status, assurance status, place of residence (urban or rural), hepatitis B infection, first symptom of the disease (abnormal pop smear, vaginal bleeding, vaginal discharge, pain and etc.), obstetrics history (parity and etc.). Mentioned variables were compared in patients with early and late diagnosis.

Statistical analysis
Data analyzed using SPSS ver. 18 (SPSS Inc., Chicago, Ill., USA). Quantitative and qualitative variables in two studied groups were compared using t-test and chi-square tests.

Results
In this study 55 women with cervical cancer enrolled. Mean age of studied population was 48.26±12.01. According to our classification 6/55 (10.9%) and 49/55 (89.1%) of them had early and delayed diagnosis of cervical cancer. Distributions of different pathologies of cervical cancers were as follows; Squamous cell carcinoma: 48 (85.71%), adenocarcinoma: 7 (12.5%) and adenosquamous carcinoma: 1 (1.79%). Demographic factors and cancer related factors in two studied groups are presented in Table 1. There were not any cases of hepatitis B infection.

Discussion
In this study we evaluated different socio demographic factors associated with delay diagnosis of cervical cancer among patients diagnosed with the disease in Isfahan. Patients with lower degree of education, lower socioeconomic status, having smoker and addict husband as well as those who did not history of pop smear test had higher probability for delayed diagnosis of cervical cancer.

The rate of delayed diagnosis in this study was high (89.09%). The rate was higher than that reported in Sudan (72%) (Ibrahim et al., 2011) and Nepal (80.9%) (Gyenwali et al., 2013). The rate indicated that though the overall prevalence of the cancer was not high in our population but higher occurrence of advanced stage cervical cancers in our community emphasized on the necessity of general screening program or improvement of population awareness in this regard.

As mentioned, absence of nationwide screening program in communities such as Iran led us to plan other multi factorial interventional studies to provide more proper preventative strategies for reduce both patients and health care professionals delay. The goal could be achieved if its associated risk factors identified. Some previous studies have investigated risk factors related to delay diagnosis of cervical cancer. Reported results regarding the role of different factors were not similar in different studies. It may be due to differences in the methods of studies, ethnicity, environmental and socioeconomic factors of the populations.

Gyenwali et al. (2013) in Nepal determined the factors associated with late diagnosis of cervical cancer among 110 patients with cervical cancer. They indicated that higher education and having symptoms of the disease specially abnormal vaginal bleeding as early symptom are related to lower delayed diagnosis of the cancer.

Tanturovski et al. (2013) have investigated the association between certain socio demographic factors and delayed diagnosis of invasive cervical cancer. Their studied population consists of 115 patients. Their results indicated that delayed presentation of the cancer is associated with lower degree of education, low monthly income, had no family history of invasive cervical cancer in first degree female relatives and poor genital hygiene.

Ibrahim et al. (2011) in Sudan have studied predictors of cervical cancer in its advanced stage. They showed that, delayed diagnosis of the disease is associated with older age, ethnicity (African), place of residence (rural) and not having insurance. They did not find any association between educational level and marital status.

Ma et al. (2012) in China indicated that older age (≥55 years), lower education (primary school education or illiterate), low annual income and being widow/divorced were the high risk factors for delayed reporting of cervical cancer.

In our study, socioeconomic condition of the family, degree of education, history of pop smear and smoking and addiction of husbands (i.e. passive smoking) had significant association with delay presentation of the disease.

Almost all of the studies in our investigated field have reported that literacy of women is considered as an independent risk factor for delayed diagnosis of the cancer. However, degree of education is indirectly linked with the average income, knowledge and understanding of nature of the disease, its related risk factors and health education (Galobardes et al., 2006; Franceschi et al., 2009; Lourenco et al., 2012). It is recommended to assess more educational programs regarding the importance of the cancer, its prevention and early diagnosis specially in population with low socioeconomic conditions.

Brazil: It is well established that the participation of patients in health related programs including screening or prevention activities is lower in those with lower socioeconomic status (SES). The association between SES and delayed cervical cancer diagnosis has been demonstrated in prior studies (Mitchell and McCormack, 1997; Lindau et al., 2002; McCarthy et al., 2010). The implication of this finding in our study is that the prevention or diagnostic programs should be target general population with more attention to those with lower socioeconomic conditions. Though in this study there was not significant relation between place of residence and delayed diagnosis of cervical cancer but it seems that this variables need to be investigated with larger sample size because both low educated and low SES are more prevalent among rural populations.

The association between smoking and passive smoking with cervical cancer has been reported in many studies. Accordingly passive smoking as a carcinogen potentially could progress transition of persistent
infection/preinvasive lesions to invasive cervical cancer. Though the mechanisms by which passive as well as active smoking induces cervical cancer is not understood clearly but possible mechanism is that tobacco smoke contains carcinogen which could cause immunosuppression and consequently progression of HPV infection to cancer (IARC, 2004; Louie et al., 2011). It seems that addiction could cause cervical cancer by similar mechanism. The lack of association between smoking of the patients and the cancer may be due to that smoking was not prevalent among Iranian women at that age. Considering the role of passive smoking and increased rate of smoking in developing countries and globally increasing trend of smoking in young females aged less than 20 (WHO, 2010), it seems that in order to perform appropriate cervical cancer prevention strategies effective tobacco control programs should be implemented.

In current study, there was significant relation between not doing Pap smear test and delayed diagnosis. Pap smear is considered the screening test of cervical cancer and has an important role in early detection, or prevention or delay in progression of cervical abnormalities to invasive cancer. It could result in significant decrease in cancer related morbidity and mortality. Many studies worldwide have studies the barriers related to poor utilization of Pap smear tests among women even in those countries with organized cervical cancer screening programs. Accordingly lack of knowledge and awareness about the cancer, its risk factors, its preventability and different cultural and geographical barriers are responsible in this regard (Chigbu and Aniebue, 2011; Bebis et al., 2012; Augusto et al., 2013; Maar et al., 2013). Though studies relieved that the barriers are presented even in women’s with appropriate knowledge, but it seems that the most important barrier specially in our population is lack of knowledge about the cancer and its risk factors as well as its prevention. So considering the results of this study it is recommended to improve the women’s knowledge in this regard using more comprehensive educational programs.

The limitations of current study were its cross sectional design and low sample size. It seems that in order to achieve more conclusive results further studies with larger sample size and prospective design is needed.

The results of this study indicated the risk factors related to delayed diagnosis of cervical cancer. Women with mentioned risk factors should be targeted for implementation of specialized educational programmes for improving knowledge and screening test. In addition the findings of this study could serve as baseline information for planning further large studies and performing large scale educational programs for general population. The consequences will be early detection, proper management and reducing disease related mortality.

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