Cervical Cancer Screening: Recommendations for Muslim Societies

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Abstract

**Background:** The overall cervical cancer incidence rate is low in Iran; however, because of a higher risk of death for cervical cancer, a disease that kills women in middle age, a cervical cancer control program is needed. The aim of this study was to provide consensus recommendation for cervical cancer prevention in Iran and other Muslim societies with low incidences of cervical cancer. **Materials and Methods:** Through a practical guideline development cycle, we developed six questions that were relevant to produce the recommendation. We reviewed 190 full text records of cervical cancer in Iran (1971 to 2013) of which 13 articles were related to the data needed to answer the recommendation questions. We also reviewed World Health Organization, IARC, GLOBOCAN report, Iran Ministry of Health cancer registry report and 8 available foreign countries guidelines. Lastly, we also evaluated the Pap smear results of 825 women who participated in the Iranian HPV survey, of whom 328 were followed-up after a 5-year interval. **Results:** The obtained data highlighted the burden of HPV and cervical cancer situation in Iran. Experts emphasized the necessity of a cervical cancer screening program for Iranian women, and recommended an organized screening program with a cytological evaluation (Pap smear) that would start at the age of 30 years, repeated every 5 years, and end at the age of 69 years. Meanwhile, there is no need for screening among women with a hysterectomy, and screening should be postponed to post-partum among pregnant women. **Conclusions:** An organized cervical cancer screening is a necessity for Iran as more than 500-900 women in middle age diagnosed with an invasive cervical cancer every year cannot be ignored. This recommendation should be taken into account by the National Health System of Iran and Muslim countries with shared culture and behavior patterns. CUBA HPV test could be consideration in countries Muslim country with appropriate budget, resources and facility.

**Keywords:** Guideline - cervical cancer - screening - pap smear - Iran - muslim societies
diagnosis of 10 squamous cell carcinoma (2/1000 CIN and 1.2/100000 cervical cancer rate). The evaluation included the overall cervical neoplasm detection, its efficiency and the costs of this screening program (split into the fixed costs and the changing costs, costs of smear taking, costs of cytological evaluation, and costs of registration). Finally, it resulted in a new policy rule in which since year 2000, the cervical cancer screening was no longer covered by the public health services. The opportunity of screening was still provided for the women based on their own request or the health workers’ (midwives, gynecologists) recommendation. It is still partly covered by the same insurance companies (M. o. Health, 2001; Allahverdipour and Emami, 2008).

The national studies showed that more than 80% of the women who were diagnosed with an invasive lesion never experienced an early detection procedure in their lifetime or had no appropriate follow up, while some other Iranian women in central cities were over-screened due to personal demand (Karimi et al., 2010; Farajzadegan et al., 2012). The national cancer registry program, which has run since 2003, showed that cervical cancer death rate (ASR) was less than 2.6% in Iran. Meanwhile, the death rate of this cancer among the women, diagnosed with an invasive cervical cancer, is about 40% (I. M. o. Health, 2012). Some studies reported an increase in abnormal pap smear results in Iran during the past few years with 2 to 4 folds growth, compared to the previous reports (Maryam and F.M.D., 2007; Farajzadegan et al., 2012). Also among the women diagnosed with cervical cancer, HPV infection prevalence, which is the primary causal agent for cervical cancer, is 70-84%. Its prevalence is 6-8% in general population (Khodakarami et al., 2012), which is lower than in many other settings, but is still higher than in some other Asian populations e.g. Pakistan, rural Vietnam (Crobbie et al., 2013).

Iran has a population of 27 million women aged 15-69 years who are at a risk for cervical cancer. Considering the potential of cervical cancer as a screen able disease we justified for developing a systematic, efficient and affordable guideline to cover majority of Iranian women population for cervical cancer screening. It should be an effective way to control cervical cancer and contribute to future considerations of the cost-effectiveness and feasibility of HPV-base vaccination.

The main object of this study was to draft a recommendation guideline for cervical cancer screening among sexually active women, based on the territorial demographic, cultural and socioeconomic data in Iran that could be work in other Muslim societies.

Materials and Methods

This study was carried out as a joint program between cancer research center, ministry of health, and health deputy of Shahid Beheshti University of Medical Sciences through a practical guideline development protocol, searching systematically for pertinent evidence to answer eight central questions addressed in the guideline (Turner et al., 2008).

The study was approved by the Review Board of SBMU, supported by policy council of Iranian Ministry of Health and supervised by a steering committee. We conducted a systematic review of previous studies, related to cervical cancer in Iran and reviewed the international guidelines, the result of a follow up study and the results of an expert panel discussion. Key words were picked up both for domestic and foreign search, based on the study goals.

The evidence was reviewed by two groups of experienced researchers of the cancer research center and supervised by its head who was also the head of Guideline development in Iranian Ministry of Health Policy Council. The idea behind having two groups was to minimize the failure chance of selecting and reviewing the studies.

Most of the studies were chosen from the medical literature databases like Pub MED, Science Direct, OVID, ISI, Islamic World Science Citation Center (ISC), Iranian Data base (SID) and Iranian medical journals (English and Persian). National studies were sorted into four categories; cervical cancer incidence-mortality, screening, cost -effectiveness studies, cervical cancer risk factors, and HPV prevalence both in women diagnosed with cervical cancer and the healthy women. Collected data were summarized and indicated with four quality grades: good, moderate, poor and inadequate. The following step started through cervical cancer screening and prevention guidelines in eight countries, which had organized cervical cancer screening program. The results of this systematic review were formulated by the Grading of Recommendations, Assessment, Development and Evaluation (GRADE) system to determine the quality of the evidences for consideration (Schunemann et al., 2008). Seven key subjects, questioned by a research group, to find appropriate evidences for a clean and rational answer to every query were as follows: i) Is an organized cervical cancer screening program needed for Iranian women? ii) What is the appropriate tool for the screening [Conventional Pap smear, HPV DNA testing, Liquid based cytology or visual Inspection with acetic acid (VIA)]? iii) What is the reasonable age to begin the screening program? iv) What is the reasonable age to finish the screening program? v) How often should the screening test be repeated? vi) Is it necessary to call post hysterectomy and pregnant women for screening? vii) Should Visual inspection with Acetic Acid (VIA) be used as an alternative test to pap smear in low resource location?

In order to achieve an appropriate answer, based on the evidences of literature review, a structured questionnaire was prepared to make the draft of the guideline, which contained multiple choice questions for the main question to be rated through a Likert’s scale (1-5) (Malhotra and Peterson, 2006). Then, the guideline developers reviewed all the data to make a decision about which outcomes would be crucial and which would be important (the balance between the ideal and the feasibility), and came to a final conclusion regarding the rating of the overall quality of the evidences, drafted as a strong recommendation.

For the translation the evidences into recommendations, we designed a questionnaire by the systematic review results of the national and international data which supported by sufficient evidence both grade A and B recommendations. The questionnaire was attached to a
Results

For the national evidences systematic review, we found 190 full text records about cervical cancer in Iran (1971 to 2013 of which 23 articles were eligible for inclusion in the review. Among them, 13 articles were directly or indirectly related to the data needed to develop the recommendation questions (Adli, 1976; Mortazavi et al., 2002; Eftekhar et al., 2005; Tbrizi et al., 2006; Maryam and F.M.D, 2007; Noorbala and Kafaie, 2007; Allahverdipour and Emami, 2008; Esmaeili et al., 2008; Farnaz et al., 2008; Nokiani et al., 2008; Zarchi et al., 2010; Khodakarami et al., 2011; Farajzadegan et al., 2012; Khodakarami et al., 2012; Khorasanizadeh et al., 2013). We also reviewed World Health Organization (Data, 2013), IARC (Bray et al., 2013). GLOBOCAN report (Ferlay and Forman, 2013), Iran Ministry of Health cancer registry report (I. M. o. Health, 2012) and nine available international guidelines (Debbie et al., 2012; Lonnberg, 2010; Levin et al., 2010; Saslow et al., 2012; van Rosmalen et al., 2012; Kulasingam et al., 2013; Peirson et al., 2013) and has been sent via email to 40 expert (Oncologist, gynecologists, epidemiologists, midwives, physicians, Pathologists and reproductive health specialists) for their comments and rating. After analyzing the result of the survey a recommendation for the guideline was developed. Also we considered the result of a 5 years follow up of 361 women who participated in a population base survey for incidence of HPV prevalence among Iranian women on 2008. We invited them again on 2012 for a repeated cytological test (conventional pap smear) after 6 years .The aim of this follow up study was to gather more appropriate national evidences for the cervical cancer screening program. To improve and confirm the recommendation, two technical advisory panel were conducted. The first one, that held in the Cancer Research Center. The draft was reviewed by twelve invited experts. The second was an external panel that discussed about the draft and results of the first expert panel recommendations. we consider the number of voted for every recommendation of the expert consultants group, then appropriates recommendation took in to account. When the guideline was refined, we conducted a symposium with combination of steering committee and all leading experts who were consultants for the guideline developer team. In addition we invited representatives from Ministry of Health, Center of Disease and Cancer Control, UNFPA, stakeholders and advocacy organizations for a one day symposium. Tow week before the symposium we sent a copy of the drafted guideline for all invited experts .Symposium was held at Academic Medical Sciences of Iran. We presented the methods of the guideline development and explore the summery of the recommendation for the cervical cancer screening in Iran. The result of the symposium discussion was analyzed and took in to account for the final version of the guideline that we written and sent to the Iran Ministry of Health for approving.

Steering committee controlled conflict of interest (COI) for all working groups concerning not having a direct or indirect relation with companies that market or sells the screening tests or devices. For the final decision panel in addition to steering committee we invited representatives from Ministry of Health, CDC, UNFPA, and advocacy organizations for a one day symposium.

The target for this guideline was the majority of women in the general population who were married, widowed or divorced. Usually unmarried females are not sexually active in Muslim Societies.

Table 1. Cervical Cancer Screening Program in Different Countries

<table>
<thead>
<tr>
<th>Country Name</th>
<th>age to Start (Year)</th>
<th>Screening Interval</th>
<th>Screening Cessation</th>
<th>Screening Tool</th>
<th>HPV DNA Test</th>
<th>Mortality (ASR)/100000</th>
</tr>
</thead>
<tbody>
<tr>
<td>Australia</td>
<td>18-20</td>
<td>every 2 yr</td>
<td>69 Yr</td>
<td>Pap Smear</td>
<td>yes</td>
<td>0.6</td>
</tr>
<tr>
<td>Canada</td>
<td>25-30</td>
<td>every 5 yr</td>
<td>69 yr</td>
<td>Pap Smear</td>
<td>No recommendation</td>
<td>1.6</td>
</tr>
<tr>
<td>Europe</td>
<td>22-30</td>
<td>every 3-5 yr</td>
<td>60-65</td>
<td>Pap Smear</td>
<td>recommendation for 30-35 Yr every 5 yr</td>
<td>10.5</td>
</tr>
<tr>
<td>Finland</td>
<td>30</td>
<td>every 5 yr</td>
<td>60</td>
<td>Pap Smear</td>
<td>No recommendation</td>
<td>1</td>
</tr>
<tr>
<td>Netherland</td>
<td>30</td>
<td>Every 5 yr for age 30-49 yr every 10 yr for age 50-60</td>
<td>60</td>
<td>Pap Smear</td>
<td>for women ≥ 30 yr</td>
<td>2.5</td>
</tr>
<tr>
<td>New Zealand</td>
<td>20</td>
<td>every 3 Yr</td>
<td>70</td>
<td>Pap smear</td>
<td>No recommendation</td>
<td>1.4-2</td>
</tr>
<tr>
<td>Singapore</td>
<td>25</td>
<td>3 yr</td>
<td>69</td>
<td>Pap Smear</td>
<td>No recommendation</td>
<td>1.8</td>
</tr>
<tr>
<td>United Kingdom</td>
<td>25</td>
<td>Every 3 yr for age 25-49 and every 5 yr after 50 yr</td>
<td>64</td>
<td>Liquid based cytology (LBC)</td>
<td>combination with Pap smear from 30-60</td>
<td>2.1</td>
</tr>
<tr>
<td>United State</td>
<td>21</td>
<td>every 3 Yr after 30 co test every 5 Yr</td>
<td>65</td>
<td>Pap Smear</td>
<td>combination with Pap smear from 30-60</td>
<td>Table1</td>
</tr>
</tbody>
</table>
to the Iranian studies and international guidelines, some international articles were identified that addressed various aspects, related to the technology assessments and cost-effectiveness of cervical cancer screening (Hsairi et al., 2000; Castle et al., 2008; Castellsague et al., 2009; Arbyn et al., 2011; de Bekker-Grob et al., 2012; Ronco et al., 2012; Sankaranarayanan et al., 2012; van Rosmalen et al., 2012; Berkhof et al., 2013; Khozaim et al., 2013; Rashid et al., 2013; Rustagi et al., 2013; Xu et al., 2013).

The first question was the necessity of cervical cancer screening program in Iran. Guideline developer and experts were completely agree that an organized cervical cancer screening is a necessity, as over 500-900 new cases with invasive cervical cancer who are diagnosed every year in Iran cannot be ignored (the annual number of new cases of cervix uteri cancer was 2.4 per 100,000 women) (I. M. o. Health, 2012; Ferlay and Forman, 2013). The guideline would be applied to all women who are or have ever been married. General information about the cervical cancer situation, according to a five-year National Cancer Registry report that presented in (Figure 1). Research shows that the main risk factor such as HPV infection, and then, smoking, passive smoking, multiparity, divorce, use of oral contraceptives for more than 9 years, low socioeconomic status and husband’s polygamy play an important role in incidence of cervical cancer in Iran (Esmaeili et al., 2008; Momtahen et al., 2009; Karimi et al., 2010; Allameh et al., 2011; Ghaemmaghami et al., 2012; Khodakarami et al., 2012; Haghshenas et al., 2013). The expert’s high listed that these risk factors should be considered in prevention programs.

**Table 2. Evidence for Tool of Cervical Cancer Screening in Iran**

<table>
<thead>
<tr>
<th>Author(s)</th>
<th>Country</th>
<th>suggestion for age to start</th>
<th>Interval</th>
<th>screening technique</th>
</tr>
</thead>
<tbody>
<tr>
<td>Burger (2012)</td>
<td>The Netherlands</td>
<td>31</td>
<td>5</td>
<td>HPV DNA</td>
</tr>
<tr>
<td>Chuck (2010)</td>
<td>Canada</td>
<td>after 30</td>
<td>3</td>
<td>Conventional Pap test, HPV, LBC</td>
</tr>
<tr>
<td>De Bekker-Grob(2012)</td>
<td>Europe</td>
<td>after 30</td>
<td>5</td>
<td>Cytology /HPV DNA</td>
</tr>
<tr>
<td>Kulasingam (2013)</td>
<td>USA</td>
<td>21</td>
<td>3 for age 21-29 and 5 for after 30 years</td>
<td>Cytology /HPV DNA</td>
</tr>
<tr>
<td>Saslow (2012)</td>
<td>USA</td>
<td>21</td>
<td>3(3 time)</td>
<td>Conventional Pap Test, LBC Pap, VILI/VIA</td>
</tr>
<tr>
<td>Levin (2010)</td>
<td>China</td>
<td>35</td>
<td>5</td>
<td>cytology test</td>
</tr>
<tr>
<td>Peirson (2013)</td>
<td>Canada</td>
<td>30</td>
<td>5</td>
<td>Co-Screening</td>
</tr>
<tr>
<td>Scrozynski (2011)</td>
<td>Germany</td>
<td>30</td>
<td>2</td>
<td></td>
</tr>
<tr>
<td>Vijayaraghavan (2010)</td>
<td>USA</td>
<td>30</td>
<td>30</td>
<td>HPV DNA</td>
</tr>
</tbody>
</table>

**Table 3. The Studies on HPV Prevalence among Healthy Iranian Women**

<table>
<thead>
<tr>
<th>Author</th>
<th>year</th>
<th>Number of tested women</th>
<th>HPV Positive N(%)</th>
<th>HPV 16 N(%)</th>
<th>HPV 18 N(%)</th>
<th>HPV 31 N(%)</th>
<th>any</th>
<th>Study Methodology</th>
</tr>
</thead>
<tbody>
<tr>
<td>Khodakarami</td>
<td>2012</td>
<td>825</td>
<td>7.8</td>
<td>2.1</td>
<td>0.2</td>
<td>0.6</td>
<td>4.9</td>
<td>Population Base</td>
</tr>
<tr>
<td>Yousefzadeh</td>
<td>2014</td>
<td>851</td>
<td>262(31.1)</td>
<td>7.3</td>
<td>2.8</td>
<td>not reported</td>
<td>not reported</td>
<td>Retrospective among Gyn Clinic</td>
</tr>
<tr>
<td>Safaei</td>
<td>2010</td>
<td>402</td>
<td>5.5</td>
<td>2</td>
<td>0</td>
<td>not reported</td>
<td>not reported</td>
<td>Population base survey</td>
</tr>
<tr>
<td>Ghafari</td>
<td>2006</td>
<td>127</td>
<td>10(13)</td>
<td>42</td>
<td>12.7</td>
<td>not reported</td>
<td>not reported</td>
<td>Retrospective among Gyn Clinic</td>
</tr>
<tr>
<td>Afshar</td>
<td>2013</td>
<td>410</td>
<td>108(26.34)</td>
<td>10.1</td>
<td>0</td>
<td>not reported</td>
<td>not reported</td>
<td>population base survey</td>
</tr>
<tr>
<td>Zandi</td>
<td>2010</td>
<td>200</td>
<td>5.5</td>
<td>3.5</td>
<td>1.5</td>
<td>0</td>
<td>not reported</td>
<td>women who attending for opportunity screening</td>
</tr>
</tbody>
</table>
consideration of national studies, which determined cost-effectiveness, sensitivity, specificity and accuracy of different technologies for cervical cancer screening (Ghaemmaghami et al., 2004; Eftekhar et al., 2005; Nokiani et al., 2008; Karimi et al., 2011; Khodakarami et al., 2011), World Health organization and international guidelines (Table 1) (David, 2010; Debbie et al., 2012; Canadian Task Force on Preventive Health et al., 2013; Data, 2013; Lowe, 2013; Simonella and Canfell, 2013). Although many evidences show that HPV test has a higher sensitivity and adding an HPV test to cytology (Co testing), or doing HPV as a primary screen followed by triage by cytology, increases detection of cervical precancerous lesions (Aoki et al., 2012; Trope et al., 2012; Untiet et al., 2013; Abraham and Stenger, 2014). Nevertheless, the HPV test has lower specificity and more expensive than cytology (van Rosmalen et al., 2012). Therefore we focused on a program that makes screening available all over the country and its feasibility, manageability and cost beneficence for the National Health System (Iran spends $346 per capita (USD) on health care) (Bank, 2013). The cytological (PAP smear) method got a good grade as a simple and low cost test for the cervical cancer screening in Iran.

We came to the conclusion that the conventional Pap smear test provides a suitable balance between the benefits and harms in a country with low HPV and cervical cancer incidence (Table 3) and a small budget for financing the health screening programs. Expert panel indicated that if liquid-Based Cytology (LBC) method is available, LBC can be a cost-effective alternative for the Pap smear in preparation of cervical cells sampling, which may make the slides easier to read, reduce the rate of inadequate smear tests, improve efficiency of handling laboratory samples and result in an increased laboratory productivity. It allows for easy combination with HPV testing in the future.

Age to start and lay off the screening
Cervical cancer screening for Iranian women should start at the age of 30 years and stop at the age of 69 years. The choice of such age points is rational and evidence-based with regard to the mean age at which a woman gets married for the first time (23.5 years), international evidences (Table 3) (Chuck, 2010; Levin et al., 2010; Vijayaraghavan et al., 2010; Sroczynski et al., 2011; de Bekker-Grob et al., 2012; Burger et al., 2012; Kaufman et al., 2013; Kulasingam et al., 2013; Peirson et al., 2013), prevalence and the mean age of diagnosis of cervical cancer among Iranian women that is 53.5 years (Karimi et al., 2010; Ghaemmaghami et al., 2012; I. M. o. Health, 2012) and the age recommended by WHO to begin cervical cancer screening (Data, 2013).

Screening interval
The committee recommended screening interval of every 5 years, and a back up by an adequate recall. Women with negative tests will be invited for rescreening by the five-year interval. Screening should continue with a 15-year interval after three five- year period with negative Pap tests. Rational and evidence for this screening interval was determined by the review of the international guidelines and recent meta analysis studies (Hamashima et al., 2010; Lonnberg et al., 2012; Canadian Task Force on Preventive Health et al., 2013; Kulasingam et al., 2013; Peirson et al., 2013), and WHO recommendation (Data, 2013). Also ,we looked at the cytological results of 328 healthy women aged 25-65 years who already interned in an HPV prevalence survey with 825 samples in 2007-2008.We followed up the women who accepted our recall for a Pap smear test 5 years after the HPV survey (Khodakarami et

<table>
<thead>
<tr>
<th>Question</th>
<th>Choices by GRADE rating</th>
<th>Grade rating (Strong, moderate, weak)</th>
<th>final experts Recommendation</th>
</tr>
</thead>
<tbody>
<tr>
<td>Is organized cervical cancer screening a requisite program for Iranian women?</td>
<td>Yes/No</td>
<td>Moderate</td>
<td>yes</td>
</tr>
<tr>
<td>What is the adequate tool for the screening by considering: cost-effectiveness, feasibility, fair access?</td>
<td>Conventional Pap smear, HPV DNA testing, liquid based cytology or visual inspection with acetic acid (VIA)</td>
<td>Strong</td>
<td>Cytology test(Conventional pap smear or LBC)</td>
</tr>
<tr>
<td>What is the reasonable time for beginning the screening program?</td>
<td>3 years after marriage at 21, 25 or 30 years</td>
<td>Moderate</td>
<td>30 years</td>
</tr>
<tr>
<td>What is the reasonable time to stop the screening program?</td>
<td>Ages: 59, 65, 69 or 75 years</td>
<td>Strong</td>
<td>69 years</td>
</tr>
<tr>
<td>At what time interval should the screening be repeated?</td>
<td>Annually, every 3,5 years</td>
<td>Moderate</td>
<td>every 5 years</td>
</tr>
<tr>
<td>Is it necessary to invite post hysterectomy and pregnant women for screening?</td>
<td>Yes/No</td>
<td>strong</td>
<td>No</td>
</tr>
<tr>
<td>Could Visual inspection with Acetic Acid(VIA) be used as an alternative test to pap smear in low resource location?</td>
<td>Yes/No</td>
<td>strong</td>
<td>Yes</td>
</tr>
<tr>
<td>Is it necessary HPV vaccination come to the national guideline</td>
<td>Yes/No</td>
<td>Moderated</td>
<td>No</td>
</tr>
</tbody>
</table>

Table 4. Questions Details of the Studies Regarding A Cervical Cancer Screening Program in Iran and Final Recommendation
Screening during pregnancy and post hysterectomy. Evidences for the screening during pregnancy and post hysterectomy were confirmed expert panel idea and highlighted that cervical cancer screening is not a necessity for women who had a total hysterectomy due to a noncancerous condition. Expert panel members believed that pregnant women do not need screening during pregnancy, except the cases with compulsory screening for whom collecting cervical cell smear by Cyto Brush should be avoided (soaking a cotton swab is the preferred sampling technique). (Debbie et al., 2012). Related to question 7 of the study: Should Visual inspection with Acetic Acid (VIA) be used as an alternative test to pap smear in low resource location? Three national evidences help to expert panel to approve VIA for rural area that may have not access to pap smear facility.

Discussion

As cervical cancer is preventable, early detectable and is potential for treatment of precancerous lesions, its screening, is an excellent opportunity, even in a region with relatively low incidence. Previous cervical cancer screening program in Iran was an edited copy of the US guideline, modified by a national expert panel. This is the first recommendation for a guideline, that developed by national evidences and an international guideline systematic review. During the review process, we found that national data did not have adequate information to get high quality evidences, therefore, guideline developers suggested giving an appropriate reply to the proposed questions through reviewing international evidences and guidelines too. We reviewed international guidelines of the countries that have successful outcome for control cervical cancer incidence by an organized screening program, such as Australia (Simonella and Canfell, 2013), Canada (Canadian Task Force on Preventive Health et al., 2013), Europe (Arbyn et al., 2010) Finland (Lonnberg, Nieminen, Luostarinen and Anttila, 2013), Netherland (van Rosmalen et al., 2012), New Zealand (Simonella and Canfell, 2013), Singapore (Jin, Louange, Chow, and Fock, 2013), United Kingdom (Duggan, 2012), USA (Karjane and Chelmon, 2013; Kulasingam et al., 2013), and involved experts and the stakeholders for a better solution. The role of opportunity cervical cancer screening that has already been proven in Iran is not clear for reducing the mortality of the cervical cancer. To reduce the mortality from cervical cancer, development an appropriate screening program and advisable management are required. In addition, to achieve its aims, a quality assurance system should be considered. The effects of new technologies, including liquid-based cytology and HPV testing, must be evaluated at the community level in Iran.

Some study indicated that LBC is not expected to increase sensitivity (Confortini et al., 2010) and others believe that liquid-based cytology is expected to decrease unsatisfactory samples compared with conventional cytology. The US Food and Drug Administration (FDA) for the first time approved a Human Papilloma Virus (HPV) test for use as a first-line primary screening test for cervical cancer screening in women 25 and older (Abraham and Stenger, 2014). Also the result of a recent study in European Union shown that HPV-based screening would provided a better protection against invasive cervical cancer compared with cytology and support initiation of HPV-based screening from age 30 years with at least 5 years intervals (Ronco et al., 2014). According the result of final round of experts panel discussion, they came to the conclusion that in countries such as Iran with limited resources, mass screening with HPV test is not best choices for now. However for an effective evaluation of the recommendation, we were advised to design a pilot study within the family physician program (that is now running in Fars and Mazandaran provinces of Iran as a pilot study). The family physician pilot program would be an opportunity to evaluate this recommendation as a guideline for cervical cancer screening within public health services and can be updated further by new national and international evidences. We should proceed to act collaboratively with national health system to create the models, protocols, and implement issues to develop an efficient, cost-effective recall and a follow-up system for cervical cancer screening that Government would be convinced to invest appropriately for it.

We recommended an organized population-based program for average risk women, aiming at controlling the cervical cancer in Iran (Table 4). According this recommendation the screening with cytology test would be start at ages 30 among women who are married. The interval is 5 years and the end of screening ages is 69 years we expect that application of this recommendation would result in a lower incidence of cervical cancer in Iran and other Muslim countries (EMRO, Middle East, and west Asia). In parallel developing awareness and empowerment programs on HPV infection as a sexual transmitted disease by the aim of the disease prevention and control for youth, could come to into account and make the behavior, culture, faith and religion issues as an asset for set up the screening guideline. By provision of an online registration software and creation of a permanent facility for regular monitoring, rating and timely updating of this guideline, we can revise the present version to a better one that will be developed completely by national and regional data in future. It is expected that this recommendation would permit less wasteful cervical screening, while take the opportunity to National Health System towards ensuring adequate coverage of high-risk women. We hope to achieve further improvements to better technology for cervical lesions early detection and prevention, such as HPV DNA test and HPV vaccination for all eligible women.

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Microbiol, 50, 1927-35.