



Trends in Incidence of Prostate Cancer in Iran and Its 31 Provinces During 2003 - 2016

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Abstract

Background: Prostate cancer is the most common malignancy in men, which is the second leading cause of cancer-related death. The prevalence of prostate cancer in Iran is lower than in Western and European countries.

Objectives: Our study presents an updated incidence rate of prostate cancer in Iran and its provinces, displays its trends, and adds its spatial distribution to the literature.

Methods: We investigate the age-standardized incidence rates (ASIRs) per annum of prostate cancer from 2003 to 2016 in Iran, using the data of the cancer registration available by the Ministry of Health and Medical Education. Crude incidence rates were calculated by dividing incident cases by the population of the whole country and each province which was available on the website of Iran Statistic Center. Age standardization was done by the World Health Organization (WHO) standard population and presented according to age and province.

Results: The crude number of incident prostate cancer cases in Iran was 49 188 between 2003 and 2016. Age-standardized incidence rate of prostate cancer showed an increasing trend from 4.46 in 2003 to 18.44 per 100 000 in 2016. Mazandaran (29.57 per 100 000 in 2008) and Esfahan (27.5 per 100 000 in 2016 and 26.51 per 100 000 in 2015) were provinces with the highest reported ASIRs, while Sistan and Baluchestan province had the lowest mean of ASIR (1.59 per 100 000) in the period of the study.

Conclusions: The incidence of prostate cancer displays an increase of more than 4 times at the end of the study period. However, the increase in coverage of data recording and greater access to diagnostic tests during these years are effective in the reported ASIRs, but the steep slope in the incidence trend of prostate cancer compared to other cancers in Iran shows its capacity to become one of the health problems that underscore the necessity of implementation of control and prevention programs, especially in high-incidence provinces.

Keywords: Prostate Cancer, Incidence, Standardized Age Incidence Rate, ASIR, Trend

1. Background

Prostate cancer is introduced as the second most common cancer of men after lung cancer; 127 610 6 new cases were reported in 2018, which caused 3.8% of all cancer-related deaths in men (1, 2). It is estimated that 2 293 818 new cases will be found by 2040 and only a small increase in mortality will happen (3).

In men, the risk of prostate cancer development is 11.6% (4, 5) and it is a major public health concern, but it should be noted that its incidence and mortality are associated with its determining factors. For instance, higher age is

an important risk factor for the development of prostate cancer; also, family history, smoking history, increasing height, higher BMI, higher intake of calories, calcium, and alpha-linolenic acid are reported to be significantly associated with the evaluated risk of fatal prostate cancer, whereas a higher level of physical activity is reported as a preventive factor (6).

Notably, the incidence rates of African-American men are higher than Caucasian men and the mortality rate is almost doubled. This finding indicates that the underlying mechanisms of prostate cancer are intricate and still need to be understood. It is hypothesized that genetic,

social, and environmental reasons are also effective in the incidence rates (7).

In Iran, a detailed understanding of prostate cancer epidemiology is of great value for health administrators and helps in the early detection and prevention of the disease. Epidemiological studies may also provide essential information for assessing the disease disparities in different provinces of the country.

2. Objectives

In the present study, we aimed at reporting national and subnational incidence of prostate cancer in Iran between 2003 and 2016.

3. Methods

The incidence of prostate cancer was assessed in Iran and its 31 provinces between 2003 and 2016 according to pathology-confirmed cases of the national cancer registry system. This is the analysis of the last updated data from the cancer registry system of Iran Ministry of Health and Medical Education. Iranian cancer registry system has been developed systemically since 2000, using software-based data from all pathology centers with coverage of 100% of the country's provinces (8). In the Iranian cancer registry system, cancer types are coded according to the International Classification of Diseases for Oncology, 3rd edition (ICD-O-3) (9), and the topography of cancers is determined based on the International Classification of Diseases, 10th edition (ICD-10). In this system, prostate cancer is detectable by C61, the malignant neoplasm of the prostate (10).

In the next step, all national data were entered into an Excel software file. To simplify the assessments of age-specific epidemiological rates through populations with diverse age structures, it is beneficial to determine short statistics that take away the effects of variation in age composition. To keep the comparability of the incidence rates in different societies, the World Health Organization (WHO) has introduced a new standard population as a basis to facilitate epidemiological research worldwide (11).

A census of data from the National Cancer Registry Center was used to accomplish this study. The population of the whole country and each province were obtained from the website of the Iran Statistics Center (12). Then, age-standardized incidence rates (ASIRs) per 100 000 per annum were calculated for the country and all provinces according to the age groups, and incident year. Age-standardized incidence rates were calculated by a direct age-standardization method based on the WHO

standard population, using STATA software version 12. To make epidemiological rates easier to compare through populations with diverse age compositions, it is useful to determine short statistics that remove the effects of age variation. Because of the importance of the comparability of the incidence rates in different populations, WHO has presented a new standard population to facilitate epidemiological studies (11).

Due to a lot of data recording defects from 2010 to 2013, the available data did not have the necessary quality to use in this study. Therefore, we decided to do a time series analysis for this period.

To find an appropriate primary model, *sc* and *pac* graphs were used. The likelihood ratio test (*Lrttest*) was carried out to compare the results. The autoregressive integrated moving average (ARIMA) model was used as the most common kind of model for predicting time series events (13). Autoregressive integrated moving average (*pdq*) is comprised of autoregressive (*p*), difference (*d*), and moving average (*q*). Seasonal trend in the disease of this model is presented as SARIMA (*p d q*) (*P D Q*)_s, where *P*, *D*, and *Q* are the same components as before, with the difference that they are considered for the seasonal delay and the component *s* is the time corresponding to one data in the previous year, which is equivalent to 12 in monthly data.

4. Results

The number of incident prostate cancer cases in Iran was 49 188 between 2003 and 2016. The ASIR of prostate cancer showed an increasing trend from 4.46 per 100 000 in 2003 to 18.44 per 100 000 in 2016 (Figure 1). The incidence rate of prostate cancer increased with age and the highest ASIR of prostate cancer was observed in the age group of more than 80 years old (Figure 2).

Mazandaran (29.57 per 100 000 in 2008), Esfahan (27.5 per 100 000 in 2016 and 26.51 per 100 000 in 2015), and Tehran (24.47 per 100 000 in 2014) are provinces with the highest reported ASIRs in recent years. Detailed ASIRs of prostate cancer in each province are summarized in Figure 3. The mean of ASIR of prostate cancer in the years of the study shows that Mazandaran province with the highest ASIR of 16.06 per 100 000 was the riskiest area in Iran followed by Esfahan (14.87 per 100 000) and Fars (14.58 per 100 000), while Sistan and Baluchestan province had the lowest mean of ASIR (1.59 per 100 000) from 2003 to 2016 in the country (Figure 4 and Figure 5).

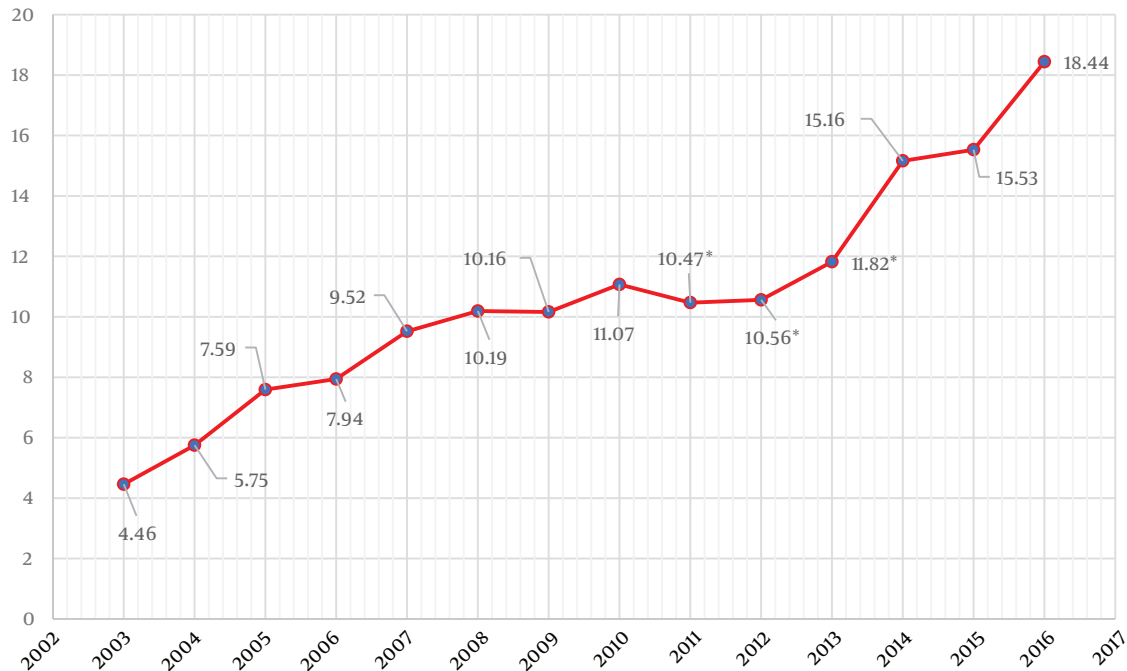


Figure 1. The age-standardized incidence rate (ASIR) per 100 000 prostate cancer in Iran between 2003 and 2016. *: ASIRs of 2011, 2012, and 2013 were provided by time series analysis due to lack of data.

5. Discussion

Cancer is the third leading cause of death in Iran (14). However, few studies have been conducted on the epidemiology of cancer in developing countries such as Iran. In Iran, national and subnational policymakers need to know basic cancer epidemiology in each area to develop comprehensive cancer prevention plans. This pathology-based study, using national data from the Ministry of Health, has examined the exact incidence of prostate cancer in all provinces of the country and provides vital information about the trend of prostate cancer incidence in the whole country and all provinces in each year from 2003 to 2016.

The annual incidence of age-adjusted prostate cancer is reported to be 152 per 100 000 men, but the incidence varies widely between different races and ethnicities. For example, prostate cancer is now the second leading cause of cancer death in the United States, affecting 1 in 7 men (6). Approximately, 1 in 36 men in the United States dies each year from prostate cancer (2). The annual incidence of prostate cancer in Caucasian men is 144.9 per 100 000, while for African-American men, it is 228.5 per 100 000. Alaskan Native Americans have the lowest incidence (77.8 per 100 000) in the Americas (6).

Prostate cancer most often affects older men, with 60% of patients being men over 65 years of age or older, and the average age of patients at the time of diagnosis is estimated at 67 years (2).

According to WHO, in 2012, the incidence of this disease in East and South Asia was 10.5% and 4.5%, respectively (6). Although the incidence of prostate cancer in Asia is lower than in Western countries, the prevalence of prostate cancer in Asian countries has increased dramatically in the last few decades. Prostate cancer is currently the 6th most common cancer in Asian men and West Asia has the highest of all cancers and South Asia has the lowest (15).

So far, there are no accurate statistics on the prevalence of prostate cancer in Iran, but in limited studies, its prevalence is lower than in Western and European countries. In Iranian men, prostate cancer is the third most common visceral cancer and the 7th deadliest cancer (16, 17).

According to the results of this study, the raw number of patients and ASIR prostate cancer in Iran shows an increasing trend from 2003 to 2016. Changing the ASIR of prostate cancer from 4.46 in 2003 to 18.44 per 100 000 in 2016 indicates an increasing trend of 4-fold. This

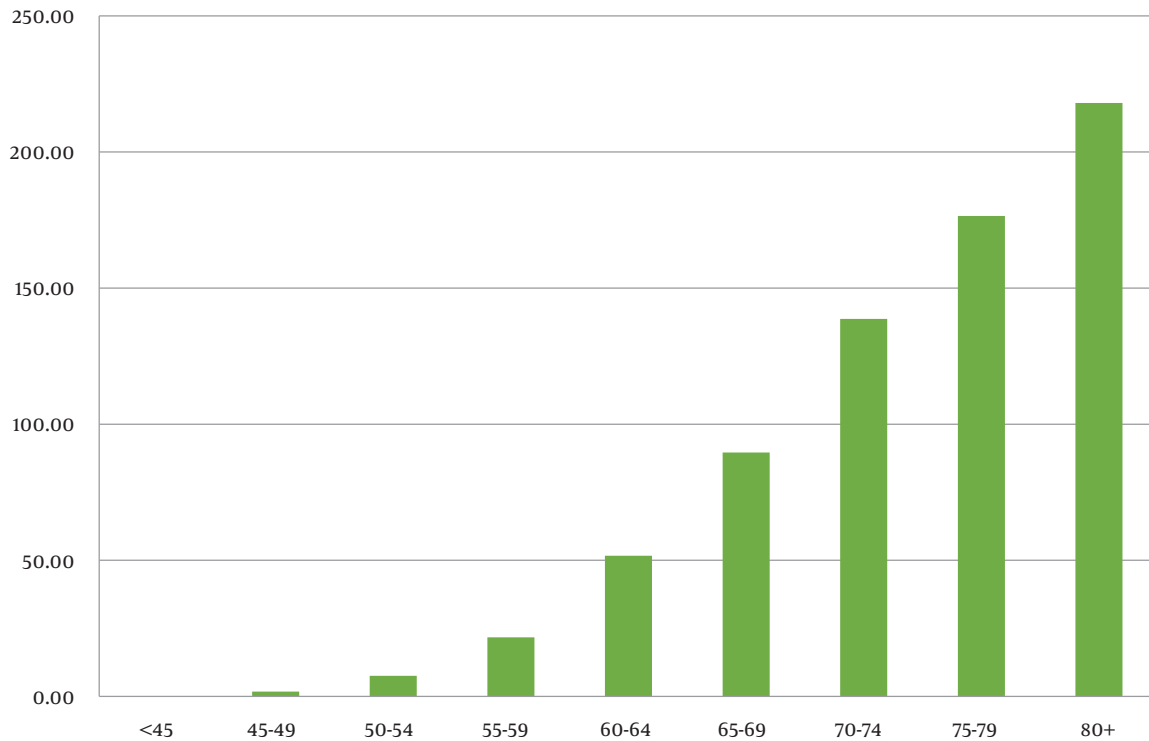


Figure 2. The age-standardized incidence rate (ASIR) per 100 000 prostate cancer in different age groups in Iran from 2003 to 2016

increasing trend is following the results of a systematic review and meta-analysis on the incidence of prostate cancer in Iran. The researchers showed a significantly higher pooled estimate of the ASIR (11 per 100 000) from 2004 to 2012 compared with those conducted from 1996 to 2003 (6.3 per 100 000) ($P < 0.001$) (18).

A noteworthy point in the study of the incidence in the provinces of the country is the presence of high ASIR levels of prostate cancer at the end of the study period (2016) in some provinces, including Esfahan 27.50, Mazandaran 24.44, East Azerbaijan 21.59, and Tehran 21.07 per 100 000, indicating an increased risk of prostate cancer in the coming years in Iran.

Although Iran has a mild incidence of prostate cancer compared to other parts of the world, the results of this study show that the incidence trend of prostate cancer in Iran has a different pattern of changes in the world. This trend can be due to the lack of a systematic screening program in Iran, which underscores the need for a regular and comprehensive screening program as well as the development of treatment guidelines.

Investigating the risk factors of prostate cancer in high-risk provinces is valuable to prevent and control the

increasing trend of prostate cancer incidence.

Worldwide, factors such as population growth, population aging, and changes in the incidence and mortality rates highlight the importance of paying attention to urological cancers, including prostate cancer; as such cancers are more common in older age groups in countries. In developing countries like Iran, where life expectancy is higher, the burden of these malignancies may increase significantly by increasing the mean age of the population. Common and controllable risk factors such as smoking and obesity in prostate cancer underscore the importance of preventive programs in this regard.

5.1. Conclusions

Although the increase in coverage of data recording and greater access to diagnostic tests during these years is effective in the reported ASIRS, the steep slope in the incidence trend of prostate cancer compared to other cancers in Iran (8) shows its capacity to become one of the health problems in the future that emphasize the duty of the health officials to reduce the serious consequences of the disease in the coming years by evaluating risk factors, implement screening programs, and develop national therapeutic guidelines across the country.

Year	2003	2004	2005	2006	2007	2008	2009	2010	2014	2015	2016	
Province												
Ardebil	Number	21	2	12	21	19	21	23	28	70	61	88
	ASIR	5.81	0.55	3.32	5.81	5.26	5.81	3.64	4.43	10.87	9.43	13.52
Esfahan	Number	123	129	192	207	264	232	345	368	553	678	7093
	ASIR	6.33	6.64	9.89	10.66	13.60	11.95	13.93	14.86	21.79	26.51	27.50
Alborz*	Number	-	-	-	-	-	113	125	94	173	189	250
	ASIR	-	-	-	-	-	9.24	10.22	7.69	13.43	14.43	18.77
Ilam	Number	8	3	3	4	22	21	23	24	44	33	34
	ASIR	4.74	1.78	1.78	2.37	13.04	12.45	8.14	8.50	15.17	11.26	11.49
East Azarbaijan	Number	8	6	54	54	143	159	152	177	366	366	419
	ASIR	0.65	0.49	4.42	4.42	11.69	13.00	8.08	9.40	19.08	18.96	21.39
West Azarbaijan	Number	8	42	52	57	56	69	93	95	221	228	236
	ASIR	0.92	4.80	5.95	6.52	6.41	7.89	5.98	6.11	13.70	13.96	14.28
Bushehr	Number	2	8	25	14	16	15	24	23	48	62	53
	ASIR	0.67	2.66	8.31	4.66	5.32	4.99	4.28	4.10	8.03	10.16	8.51
Tehran	Number	123	251	364	431	670	1312	1235	1166	1549	1430	1361
	ASIR	1.96	4.00	5.80	6.86	10.67	20.89	20.12	19.00	24.47	22.36	21.07
Chaharmahal Bakhtiari	Number	6	19	17	16	23	20	23	31	36	51	49
	ASIR	2.69	8.53	7.64	7.19	10.33	8.98	5.10	6.87	7.76	10.90	10.36
South Khorasan*	Number	-	-	-	-	-	14	12	26	44	26	42
	ASIR	-	-	-	-	-	8.45	3.61	7.81	11.52	6.72	10.71
Razavi Khorasan	Number	29	125	178	218	240	238	261	340	398	405	392
	ASIR	1.51	6.51	9.27	11.35	12.49	12.39	8.70	11.34	12.70	12.74	12.16
North Khorasan*	Number	-	-	-	-	17	11	10	16	34	27	36
	ASIR	-	-	-	-	8.63	5.58	2.32	3.71	7.62	6.00	7.91
Khuzestan	Number	39	63	83	95	122	221	197	167	278	344	354
	ASIR	2.66	4.30	5.66	6.48	8.32	15.08	8.62	7.30	11.70	14.29	14.53
Zanjan	Number	16	12	33	24	18	29	39	42	78	84	44
	ASIR	5.67	4.25	11.68	8.50	6.37	10.27	7.66	8.25	14.89	15.58	8.24
Semnan	Number	10	13	11	8	19	37	45	40	55	45	47
	ASIR	4.44	5.78	4.89	3.55	8.44	16.44	14.09	12.53	16.47	13.24	13.62
Sistan and Baluchestan	Number	2	6	8	7	24	7	20	16	27	25	29
	ASIR	0.33	0.98	1.30	1.14	3.91	1.14	1.58	1.26	1.99	1.80	2.04
Fars	Number	68	107	168	146	233	223	274	367	537	547	422
	ASIR	5.05	7.94	12.46	10.83	17.29	16.55	11.83	15.85	22.52	22.71	17.35
Qom	Number	10	23	20	23	36	41	16	28	0	0	88

	ASIR	1.99	4.58	3.98	4.58	7.16	8.16	2.72	4.76	0	0	13.73
Kordestan	Number	15	22	38	28	36	54	56	59	58	45	91
	ASIR	3.45	5.06	8.75	6.45	8.29	12.43	7.46	7.85	7.61	5.87	11.82
Kerman	Number	27	17	23	44	30	50	98	109	152	132	200
	ASIR	3.40	2.14	2.89	5.53	3.77	6.29	6.61	7.35	9.83	8.41	12.57
Kermanshah	Number	34	49	67	36	66	76	65	69	91	123	115
	ASIR	5.35	7.71	10.55	5.67	10.39	11.96	6.62	7.03	9.19	12.40	11.57
Kohgiluyeh and Buyer Ahmad	Number	0	10	12	8	16	21	17	31	42	34	25
	ASIR	0	6.46	7.76	5.17	10.34	13.57	5.17	9.42	12.14	9.69	7.00
Goolestan	Number	13	23	34	35	28	41	40	48	0	93	109
	ASIR	3.26	5.78	8.54	8.79	7.03	10.29	4.51	5.41	0	9.77	11.25
Gilan	Number	22	55	71	65	92	87	111	192	194	247	243
	ASIR	3.39	8.48	10.95	10.02	14.19	13.42	9.01	15.59	15.47	19.59	19.15
Lorestan	Number	17	41	34	33	85	73	71	105	59	43	62
	ASIR	3.30	7.96	6.60	6.41	16.51	14.18	8.03	11.88	6.52	4.71	6.75
Mazandaran	Number	51	97	84	108	129	231	184	178	266	346	393
	ASIR	6.53	12.42	10.75	13.83	16.52	29.57	11.93	11.54	17.41	21.69	24.44
Markazi	Number	22	23	33	30	39	40	84	90	71	121	141
	ASIR	4.66	4.87	6.98	6.35	8.25	8.47	11.74	12.58	9.63	16.26	18.75
Hormozgan	Number	16	8	28	21	24	28	30	30	46	58	70
	ASIR	4.60	2.30	8.05	6.04	6.90	8.05	3.74	3.74	5.39	6.69	7.88
Hamedan	Number	19	29	48	58	40	65	52	90	151	125	218
	ASIR	3.85	5.88	9.73	11.75	8.11	13.17	5.89	10.19	16.89	13.94	24.22
Yazd	Number	34	37	20	17	31	66	84	91	130	107	124
	ASIR	8.30	9.03	4.88	4.15	7.57	16.12	15.17	16.44	23.64	19.07	21.68
Undetected province	Number	832	819	983	1006	820	1	0	0	0	0	0

Blue cells indicate ASIRs less than 5, green cells indicate ASIRs 5 to < 10, yellow cells indicate ASIRs 10 to < 15, orange cells indicate 15 to < 20, and red color cells indicate ASIRs ≥ 20 per 100 000.

- Empty cells are due to the separation and naming of the provinces after the respective years.

Figure 3. Crude number of incident cases and the age-standardized incidence rate (ASIR) per 100 000 prostate cancer in different provinces of Iran from 2003 to 2016

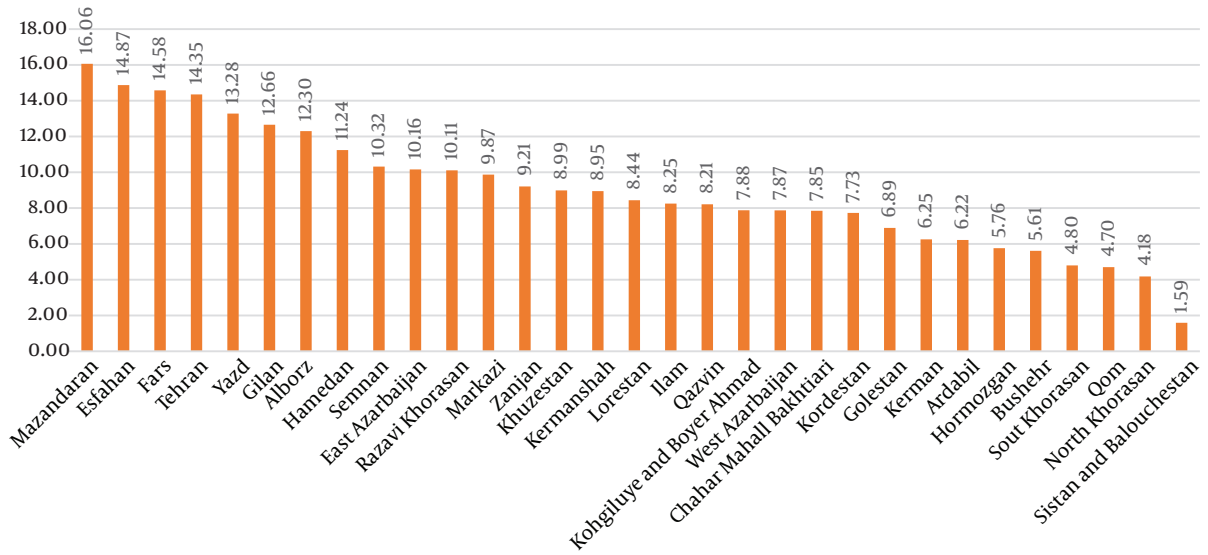


Figure 4. The mean age-standardized incidence rate (ASIR) per 100 000 prostate cancer in different provinces in Iran from 2003 to 2016

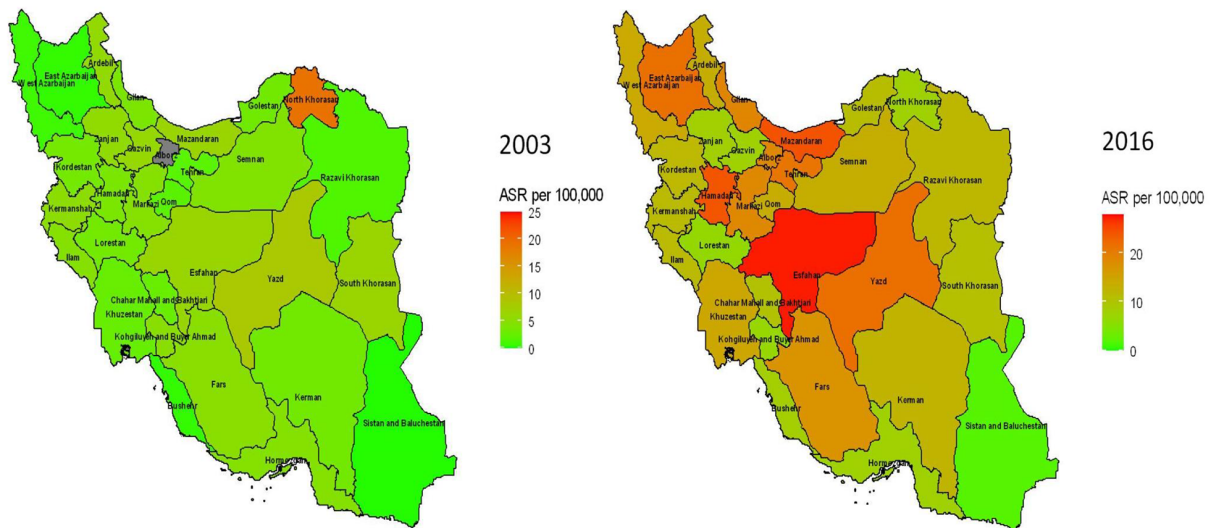


Figure 5. The age-standardized incidence rate (ASIR) per 100 000 prostate cancer in different provinces in Iran in 2003 and 2016

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Footnotes

Authors' Contribution: Study design: MRN and SB; data collection: EMF, LS, and SHI; data analysis: KH, EMF, and SB;

drafting the manuscript: LS, EA, SHI, and SAM; intellectual input and manuscript revision: MRN, EA, and SAM; all authors have read and approved the final manuscript. They also accepted responsibility for all aspects of the work.

Conflict of Interests: The authors declare that they have no competing interests.

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