Risk Factors of Ovarian Cancer in Eastern Province - Saudi Arabia

Azza Ali El-Mahalli Ali, ¹ Sahar Ahmed A. Al-Shamandy, ² And Mohamad Khaled Abdel-Hadi ³

¹Professor - Public Health Department. College of Public Health in Imam Abdulrahman Bin Faisal University-Dammam- Kingdom of Saudi Arabia

²Assistant Professor - Obstetrics and gynecological Nursing – Fundamental of Nursing Department - College of Nursing in Imam Abdulrahman Bin Faisal University- Dammam- Kingdom of Saudi Arabia

³Gynecologic Oncology Consultant- Deputy Chair, Department of Obstetrics & Gynecology- King Fahad Specialist Hospital- in Dammam, Saudi Arabia.

Corresponding Author: Azza Ali El-Mahalli Ali

Abstract: The leading cause of death from gynecologic malignancies remains ovarian cancer therefor; preventive strategies are instantly needed to decrease mortality. Project intended to determine the risk factors of ovarian cancer among Saudi females, in Eastern Province. Setting: was at King Fahd Specialist Hospital in Dammam- Eastern Province. Design: was case-control, hospital-based: Cases were counted to 60 ovarian cancer Saudi patients. Controls were counted to 240 Saudi women without ovarian cancer, who attended outpatient clinics of the study hospital, (ratio was1:4). Pre-coded data collection sheet was designed to collect data from cases and controls through interview and from medical records from the first of January, 2016 to the end of December, 2016. Data included the socio- demographic characteristics, reproductive history, and lifestyle attributes of participants. Data was analyzed using SPSS (V.20). Cases & controls were compared for risk factors. Results demonstrated that risk factors of ovarian cancer were illiteracy, late menopause, low parity and eating high fat diet. However, history of ovarian cancer in the family and lack of daily physical exercise were surprisingly protective factors. Recommendation: Further research to investigate why these possible risk factors raise the likelihood of ovarian cancer is recommended.

Key words: Ovarian cancer- Risk factors- Eastern Province -Saudi Arabia

Date of Submission: 19-03-2018

Date of acceptance: 07-04-2018

.

I. Introduction

Ovarian Cancer is considered the 8th most common malignancy among ladies worldwide. It has the greatest mortality rate of gynecological malignancies; hence, it has serious effects on the socioeconomic and society ^(1,2) The 5-year survival rate ranges from 30% to 45% without significant improvement over the last three decades, even with the new methods have been used in treatment. The American Cancer Society estimated that 22,240 women were going to be diagnosed with ovarian cancer in the US in 2013, ⁽³⁾

Women who are older in age ">50" and those with critical family history have a higher risk of developing ovarian cancer; The risk also increases by around threefold when having 1st or 2nd degree relative diagnosed with ovarian malignancy. Older age, history of ovarian cancer in the family, infertility medications, hormonal replacement therapy during menopause, and obesity were established to be risk factors related to ovarian cancer. Women with ovarian cancer are less likely to have reported a tubal ligation, and more possible to account a first-degree relative with breast cancer. In contrast, oral contraceptives use is a protective factor, (4-5) specifically this occurs owing to the decreased estrogen levels in ovaries and stoppage of ovulation.

Physical activity is another protective factor for ovarian cancer. (4-5) Moorman et al. also stated breast-feeding to be a defensive factor for the existence of ovarian cancer. Study founds a protective role for some reproductive parameters (parity, twin pregnancy, and oral contraceptive usage) against non-mucinous ovarian cancer, but not mucinous tumors. Exclusive associations comprise an opposite relation of serious cancer risk to body mass index, however a positive relation of mucinous cancer risk to cigarette smoking, and a weakly positive relation of endometrioid cancer risk to body mass index. Risk of all histologic types was unassociated with age at menopause, history of infertility, non-contraceptive estrogen use, and alcohol consumption. (9) **The rationale for the current research** is that, once better understanding of the risk factors of ovarian cancer, it is

expected that it becomes possible to introduce procedures that reduce prevalence of the disease by either improving or preventing these risk factors.

Study design was case control. Case: control ratio was 1:4. Cases counted to 60 and controls counted to 240. **Results of the current research** demonstrated that risk factors of ovarian cancer were illiteracy, late menopause, low parity and eating high fat diet. However, family history of ovarian cancer and lack of practicing sports daily were surprisingly protective factors. No obstacles faced research team. **Obstacles** for this study are: the researchers using only living cases to measure the ovarian cancer risk factors, science some risk factors could be identified in severe cases, and using the controls from the hospital (relatively sick) and not from the community (relatively healthy)

II. Literature review:

The fifth most common reason of cancer death in women worldwide is ovarian cancer. Globally, more than 200,000 women are recognized with this disease. According to National Cancer Registry in KSA, ovarian cancer is the seventh most common cancer diagnosed in females representing 3.3% of all female cancer patients diagnosed during 2013. (10) Detailed descriptive data concerning the clinical course of the disease among Saudi women is largely unknown. (11)

In KSA, numerous studies were done on broad analysis of ovarian cancer features, assessment of the disease development, and also the prognostic factors that may influence survival. One research studied integration of Human Papilloma Virus - in Saudi women with ovarian cancer. (12-15) As no research studied the association of ovarian cancer with lots of independent variables; socio- demographic, reproductive history, and lifestyle attributes among Saudi women in Eastern Province- KSA, the research intended to determine predictors (risk factors) of ovarian cancer among Saudi patients attending King Fahd Specialist Hospital (KFSH) in Dammam - Eastern Province, KSA for treatment / follow up.

Currently, there is a lack of effective early diagnostic tools for ovarian cancer. Also, there is an apparent major knowledge gap of ovarian cancer and its suggestive symptoms among both public and many health specialists. These factors have significantly contributed to the late stage diagnosis of most ovarian cancer cases (63% are diagnosed at stage III or IV), where the 5-year survival rate is less than 30%. (1) Thus preventive strategies are urgently required to decrease incidence of ovarian cancer. One of these strategies is determining its risk factors.

III. Material and Methods

Study Setting: King Fahd Specialist Hospital (KFSH) in Dammam - Eastern Province, KSA, is a large specialized hospital for treatment of cancer. It has 640 beds. The hospital has a "**Cancer registry center**" that started registration of cancer cases since 2006. The hospital provides different specialties including medical oncology, surgical oncology, nuclear medicine, radiation oncology, and rehabilitation.

Study Design: Case- control hospital- based study.

Sample Size: Case: Control ratio was 1:4 to increase study power ⁽¹⁶⁾. Cases counted to 60 patients and controls counted to 240 patients. Duration of the study: One year (from the first of Muharam 1437 to the first of Muharam 1438).

Inclusion Criteria:

- 1. Ovarian cancer diagnosis is confirmed at KFSH.
- 2. Saudi women.

Exclusion Criteria:

- 1. Control subjects with bilateral oophorectomy.
- 2. Patients refuse to participate in this study.

Tool:

A pre-coded data collection sheet was designed $^{(12-15)}$ to collect data from cases and controls, which included the following parts:

Part I: Socio-demographic characteristics, which comprised: woman's age; marital status; educational level; ...etc.

Part II: Reproductive history, which involved: woman's age at menarche & age at menopause, parity, use oral contraceptive method, tubal ligation, use fertility drugs (e.g. Clomid), infertility, use hormone replacement therapy during menopause, family history of ovarian, breast, or colorectal cancer, individual history of breast

cancer, use of talc powder on genital area, history of multiple births, endometriosis, ovarian cyst, polycystic ovary, use of intrauterine device, and breast feeding.

Part III: Lifestyle attributes e.g. Body Mass Index (BMI), low fat diet, daily eating of fresh fruits and vegetables, smoking, alcohol use, tea consumption, and daily physical exercise.

IV. Methods

For Cases: Saudi patients with ovarian cancer, who fulfilled the inclusion criteria, were interviewed during their treatment/ follow up visits after taking their written consent. Moreover, the reproductive history was obtained from both their paper and electronic health records.

For Controls: Saudi women without ovarian cancer were interviewed while attending the outpatient clinics of the study hospital. Their written consent was taken first. Their medical records/ investigation reports were reviewed to make sure that they have no ovarian cancer diagnosed.

Ethical Consideration: Before conducting the research, formal approval from the institutional review board (IRB) of the study hospital was taken and confidentiality of the data collected was maintained. Informed written consent was taken from participants.

Pilot Study: It was conducted after obtaining the IRB. Fifteen medical records were reviewed to investigate documentation of risk factors. Number of menstrual cycles when Clomid was prescribed was removed due to recall problems.

Data Analysis: SPSS program (version 20) was used for data analysis. Quantitative data were presented as mean & standard deviation and qualitative data were presented as number & percentage. Cases and controls were compared and test of significance at <0.05% P- Value was used. The measures of risk were calculated by using univariate analysis to study the relation between risk factors and ovarian cancer. Then multivariate logistic regression was used.

V. Results

(**Table 1**) Revealed that most of cases aged 30 and above (81.7%) vs control (70.4%). Cases who were ever married (66.6%) vs controls (71.2%). Percent of cases who completed their secondary school and university education or more was low (55%) in comparison to controls (77.9%), difference was significant (P <0.0001). Cases who had early menarche at or less than 13 years were (65%) vs controls (80.8%), difference was significant (P 0.008). Cases with family history of ovarian cancer represented (8.3%) vs controls (2.5%); difference was significant (P 0.032). Majority of cases had high fat diet (66.7%) vs controls (32.9%), difference was significant (P <0.0001). High percent of cases had BMI 30+ (61.7%) vs controls (57.1%), difference was significant (P <0.0001). Cases who did not have daily physical exercise represented (76.7%) in comparison to controls (87.9%), difference was significant (P 0.026).

Multivariate regression analysis (**Table 3**) demonstrated that odds of getting ovarian cancer for illiterate were 68.781 times more than the odds for non- exposed to illiteracy (OR: 68.781, 95% CI: 10.141-466.501). Women with late menopause (> 51 years) was getting the risk of ovarian cancer by 2.979 times more than those with menopause (\le 51) (OR: 2.979, 95% CI: 1.177-6.947). The risk of having ovarian cancer for women with parity \le 3 was 3.066 times higher than those having > 3 parity (OR: 3.066, 95% CI: 1.186-7.928). The odds of getting ovarian cancer was 7.536 times higher for women with high fat diet than those who had low fat diet (OR: 7.536, 95% CI: 3.300-17.210). In addition, the risk of having ovarian cancer for women with BMI 18.5-24.9 was 28.758 times more than those non-exposed. In addition, odds of getting ovarian cancer for women with family history of ovarian cancer were 92.7 % times less than those non-exposed (OR: 0.073, 95% CI: 0.017-0.326). Similarly, the odds of getting ovarian cancer for women who did not have daily exercise were 83.6% times less than women who exercised daily (OR: 0.164, 95% CI: 0.061-0.442). Difference was statistically significant for previously mentioned independent variables.

VI. Discussion

The leading cause of death from gynecological malignancies remains ovarian cancer. However, most women present with advanced stage of ovarian cancer due to lack of effective screening approaches. Therefore preventive strategies are urgently needed to reduce mortality. Researchers reported several risk factors for developing ovarian cancer, such as menstrual and reproductive factors, with the exception of parity, remains uncertain. (17)

Parity is considered a well-established protective factor for ovarian cancer. There were 6952 incident cases of ovarian cancer reported in a prospective cohort study of middle-aged women in UK, which was done on 1.1 million women. Overall, women with multiparity had a 26% lower risk of ovarian cancer than nulliparous women (OR: 0.74, 95% CI: 0.69–0.79). (18)

Between 2006 and 2008, a hospital-based case-control study was undertaken in Guangzhou, Guangdong Province in a sample of 500 ovarian cancer patients and 500 controls (mean age, 59 years). By using unconditional logistic regression models, OR and 95% CI were calculated. High parity was founded not associated with ovarian cancer, with an adjusted OR 0.43 (95% CI, 0.30- 0.62) for women who had delivered three or more births compared to women who had delivered no more than one birth. (19) The present findings are parallel to the published research as the risk of having ovarian cancer for women with parity ≤ 3 was 3.066 times higher than those having > 3 parity (Table 3).

Family history of ovarian or breast cancer is remains the most important risk factor for epithelial ovarian cancer. ⁽²⁰⁾ On the contrary, the present study showed that odds of getting ovarian cancer for women with family history of ovarian cancer were 92.7 % less than in non-exposed (Table 3). This may be attributed to the fact that researchers collected data from all ovarian cancer patients regardless the histological type of ovarian cancer

The risk of developing ovarian cancer gets higher with woman's age. Ovarian cancer is rare in women younger than 40. Nearly, half of all ovarian cancers are found in women 63 years of age or older. (21) Age at menopause was also shown to be positively associated with the risk of ovarian cancer (>52 vs. \leq 47 years: HR 1.46, 95 % CI 1.06–1.99). (22) This goes parallel to the present findings as the risk of getting ovarian cancer for women with late menopause > 51 years was 2.979 times higher than those with menopause \leq 51 (Table 3).

The present findings demonstrated that odds of getting ovarian cancer for illiterate were 68.781 times higher than non- exposed to illiteracy (Table 3). Literature reported that socioeconomic characteristics play an important role in survival after ovarian cancer, such as; low level of education; lower income and living without a partner were related to poorer survival after ovarian cancer. Among women with early cancer stage, HR (95% CI) for death was 1.75 (1.20-2.54) in shorter compared to longer educated women.

A numerous studies have found a relation between obesity and hormone dependent cancers including, endometrial cancer and postmenopausal breast cancer, but the relation with ovarian cancer is unclear. (24-25) Researchers in a systematic literature review and meta-analysis, found that 24 of 28 studies reported a positive association between obesity and ovarian cancer, and 10 of this reached statistical significance. The pooled effect estimate for adult obesity was 1.3 (95%CI 1.1 -1.5) with a smaller increased risk for overweight (OR 1.2; 95% CI 1.0 -1.3). The pooled OR was stronger among case—control studies (OR= 1.5) than cohort studies (OR= 1.1).

In 2001 A systematic review and meta-analysis conducted by Purdie and colleagues, concluded that there was a small-to-moderate positive relation between high BMI and ovarian cancer risk with statistically significant differences. (27) Results of the present study revealed that the risk of having ovarian cancer for women with BMI 18.5-24.9 (healthy range) was 28.758 times higher than non-exposed (Table 3). The contradicting findings could be attributed to the fact that BMI in the current study is that measured at time of data collection and not at time of diagnosis as done in the published research.

Dietary factors could explain some of the observed differences in ovarian cancer incidence worldwide. Research revealed that greater consumption of fat may increase risk of ovarian cancer overall. $^{(28)}$ In the Women's Health Initiative Dietary Modification randomized controlled trial, a low fat and high fruit, vegetable and grain dietary pattern (vs usual diet) was associated with a reduced risk for ovarian cancer in postmenopausal women after 4–8 years of follow-up (RR= 0.60, 95% CI 0.38–0.96). $^{(29)}$ This goes hand in hand with the current findings as the odds of getting ovarian cancer was 7.536 times higher for women with no low fat diet than those who had low fat diet (Table 3).

Between 2006 and 2008, eighty consecutive patients were enrolled in a study in two national referral centers for ovarian cancer in Serbia. Results revealed that patients with ovarian cancer had physical exercise for 6.3 ± 2.1 years and controls for 11.8 ± 9.9 years. Physical exercise was statistically significant protective factor (OR = 0.2, p = 0.011 and OR = 0.4, p = 0.019 respectively). (30) In contrast, present findings revealed that the odds of getting ovarian cancer for women who did not have daily exercise were 83.6% less than women who had daily exercise (Table 3). This may be attributed to lack of data concerning duration of physical exercise.

Conclusions and Recommendations: Risk factors of ovarian cancer were illiteracy, late menopause, low parity and eating high fat diet. This may be due to small sample size of 60 cases evaluated in this study and the short approval time that taken from the institutional review board (IRB) of the study hospital for conducting the research. However, family history of ovarian cancer and lack of daily physical exercise were surprisingly protective factors.

Finally further research to investigate why these risk factors increase the likelihood of developing ovarian cancer is recommended. In addition, researchers recommend that health care providers must formulate different prevention strategies for women attending the outpatient clinics about the risk factors of ovarian cancer.

Acknowledgement

This project was funded by King Abdulaziz City for Science and Technology (SG-36-30). Without the help and support of helpful KACST team, this study could not have been achieved. We would like to express our thanks to everyone who helped us in making this study successful. All thanks go to KACST team who has supported us all along our journey to complete this study. All the gratefulness can't describe how grateful we are to the support they gave us. This work could not have been possible without the help of Imam Abdulrahman Bin Faisal University (Before-University of Dammam) and King Fahd Specialist Hospital in Dammam city. Moreover.

this work cannot be completed without the help of the research assistants: Nora Fahad Ibrahim AlShuwayer, Sara Fayez Jaber Aldawood and Maha Saad Alhumoud, (Graduates from the University of Dammam) and they were responsible for data collection. Also we would like to acknowledge all women who enrolled in this study for their valuable participation.

Funding source: The project is funded by King Abdulaziz City for Science and Technology (KACST) {SG-36-30}. Sponsor has no role in the study plan; and in the gathering, investigation & interpretation of data, writing of the manuscript, or in the decision to submit the manuscript for publication.

Conflicts of Interest:

The authors declare that no financial or personal conflict of interest for this work. However, the limitations of the study that may affect on the study results are: choosing only living cases of ovarian cancer because difficulties of getting all information related to ovarian cancer risk factors from the medical records (as not recorded in the patient medical record). And related to the controls were choosing from the general outpatient clinics of the study hospital not from the community.

Tables:

Table 1: Distribution of Ovarian Cancer Cases Vs Controls attending King Fahd Specialist Hospital in Dammam- Fastern Province - KSA

Independent variables	Cases	110,111	Controls		Chi-	P-value	
	(n=60) (n:		(n=240)		squar		
	n	%	n	%	e		
I. Socio-demographic characteristics							
Age	Mean ± SD Mean		Mean ± SI	Mean ± SD			
	45.3±14.6		41.6±15.9				
<30	11	18.3	71	29.6	4.182	0.124	
30-	25	41.7	100	41.7			
50+	24	40.0	69	28.7			
Marital status							
Married	29	48.3	142	59.2	3.304	0.347	
Single	20	33.3	69	28.8			
Widow	7	11.7	15	6.2			
Divorced	4	6.7	14	5.8			
Education level							
Illiterate	10	16.7	3	1.2	36.10	< 0.0001	
Completed primary/intermediate school	17	28.3	50	20.8	2		
Completed secondary school	22	36.7	80	33.3			
Completed university or more	11	18.3	107	44.6			
II. Reproductive history & other h	ealth-related	l data					
Age at menarche	Mean ± SI 12.6±1.7)	Mean ± SI 12.7±1.4	Mean ± SD 12.7±1.4			
≤ 13	39	65.0	194	80.8	6.937	0.008	
> 13	21	35.0	46	19.2	1		
Age of menopause*	Mean ± SI	Ò	Mean ± SI)			
	43.5± 7.1		47± 5.2				
≤51	34	87.2	67	87.0	0.001	0.980	
>51	5	12.8	10	13.0			

	Parity	Mean ± SD		Mean ± SD			
No	< 3	3.6± 4.1	59.2	2.8± 2.7	63.2	0.511	0.475
History of multiple births						- 0.511	0.713
Yes	_	23	41.7	00	30.7	<u> </u>	
No		3	5.0	23	9.6	1.274	0.259
Yes 1		57	95.0	217	90.4		
No		1 00	10.5		1 40 :	1.65-	0.000
Ves					1	1.235	0.266
Yes		31	31.7	143	39.0	L	
No		6	10.0	19	7.9	0.273	0.602
Yes	No	54		221			
No							
No						0.079	0.779
Yes		53	88.3	215	89.6	1	
No	· ·	3	5.0	3	1.2	3.444	0.063
Yes			_				
No	Use of fertility drugs e.g. clomid						
No						4.270	0.039
Yes 1 1.7 5 2.1 23.00 <0.0001 Family history of ovarian cancer Teamily history of ovarian cancer S 8.3 6 2.5 4.624 0.032 No 55 91.7 234 9.75 1 Is relative degree* 1° degree 5 8.3 5 2.1 6.045 0.04 Family history of breast cancer Yes 8 13.3 35 14.6 0.061 0.805 No 52 86.7 205 85.4 0.081 0.805 Is relative degree* 1° degree 7 11.7 23 9.6 1.448 0.485 8 13.3 20 8.8 0.273 0.602 Is relative degree* 7 11.7 23 9.6 1.448 0.485 Personal history of colorectal cancer 8 4 6.7 21 8.8 0.273 0.602 Is relative de			88.3	229	95.4	<u> </u>	
No			1.7	5	2.1	23.06	<0.0001
Yes							(0.0001
No							
The second color of the		_				4.624	0.032
		55	91.7	234	97.5		
Para		1 5	83	1 5	2.1	6.045	0.049
Pamily history of breast cancer Yes 8 13.3 35 14.6 0.061 0.805 No						0.043	0.047
No			0.0	-		1	
The stative degree The state The sta	Yes	_				0.061	0.805
1st degree 7 11.7 23 9.6 1.448 0.485 2nd degree 1 1.7 12 5.0 1.448 0.485 Family history of colorectal cancer Yes 4 6.7 21 8.8 0.273 0.602 No 56 93.3 219 91.2 4 0.62 Its relative degree* Ustable degree 2 3.3 20 8.3 5.742 0.057 Personal history of other cancer Yes 11 18.3 22 9.2 4.120 0.042 No 49 81.7 218 90.8 4.120 0.042 Talcum powder on genital area Yes 9 15.0 79 32.9 7.433 0.006 History of endometriosis 3 5.0 6 2.5 1.031 0.310 Yes 3 5.0 6 2.5 1.031 0.310 </td <td></td> <td>52</td> <td>86.7</td> <td>205</td> <td>85.4</td> <td><u> </u></td> <td></td>		52	86.7	205	85.4	<u> </u>	
This color of the color of the cance of the color of th	Its relative degree*	1 7	11.7	1 22	100	1 440	0.405
Parily history of colorectal cancer Yes	1 uegree 2 nd degree					1.448	0.485
Yes		1	1./	12	3.0		
Its relative degree*		4	6.7	21	8.8	0.273	0.602
1st degree 2 3.3 20 8.3 5.742 0.057 Personal history of other cancer Yes 11 18.3 22 9.2 4.120 0.042 No 49 81.7 218 90.8 7.433 0.006 Talcum powder on genital area Yes 9 15.0 79 32.9 7.433 0.006 No 51 85.0 161 67.1 0.006 6 History of endometriosis 3 5.0 6 2.5 1.031 0.310 No 57 95.0 234 87.5 1.031 0.310 History of ovarian cyst 7 95.0 229 95.4 0.019 0.891 No 57 95.0 229 95.4 0.019 0.891 History of polycystic ovary 7 1 1 4.6 0.042 0.849 0.357 No 59 98.3 230 95.8		56	93.3	219	91.2	<u> </u>	
Personal history of other cancer Yes			1 2 2	1 20	100	1 5 5 10	0.055
Personal history of other cancer Yes						5.742	0.057
Talcum powder on genital area Yes 9		1 2	3.3	1	0.4		
No		11	18.3	22	9.2	4.120	0.042
Talcum powder on genital area Yes 9 15.0 79 32.9 7.433 0.006 No 51 85.0 161 67.1 0.006 History of endometriosis Yes 3 5.0 6 2.5 1.031 0.310 No 57 95.0 234 87.5 0.019 0.891 History of ovarian cyst 3 5.0 11 4.6 0.019 0.891 No 57 95.0 229 95.4 0.019 0.891 Mistory of polycystic ovary 7 95.0 229 95.4 0.019 0.891 No 59 98.3 230 95.8 0.499 0.357 No 59 98.3 230 95.8 0.494 0.357 History of diabetes mellitus 75.0 168 70.0 0.583 0.445 No 45 75.0 168 70.0 12.89 0.0001	No		21-			1	
No	Talcum powder on genital area				_		
History of endometriosis Yes 3 5.0 6 2.5 1.031 0.310 No 57 95.0 234 87.5 0.019 0.891 History of ovarian cyst 3 5.0 11 4.6 0.019 0.891 No 57 95.0 229 95.4 0.019 0.891 History of polycystic ovary 8 1 1.7 10 4.2 0.849 0.357 No 59 98.3 230 95.8 0.357 History of diabetes mellitus 8 15 25.0 72 30.0 0.583 0.445 No 45 75.0 168 70.0 0.583 0.445 History of ever breast feeding 8 27 45.0 53 22.1 12.89 <0.0001		_				7.433	0.006
Yes 3 5.0 6 2.5 1.031 0.310 No 57 95.0 234 87.5 1.031 0.310 History of ovarian cyst Very See 3 5.0 11 4.6 0.019 0.891 No 57 95.0 229 95.4 0.019 0.891 History of polycystic ovary Very 1 1.7 10 4.2 0.849 0.357 No 59 98.3 230 95.8 0.357 History of diabetes mellitus 15 25.0 72 30.0 0.583 0.445 No 45 75.0 168 70.0 0.583 0.445 History of ever breast feeding 27 45.0 53 22.1 12.89 <0.0001 No 33 55.0 187 77.9 1 1 III. Lifestyle attributes 55 91.7 237 98.8 16.31 <0.0001		51	85.0	161	67.1		
No		3	5.0	6	2.5	1.031	0.310
History of ovarian cyst Yes 3 5.0 11 4.6 0.019 0.891 No 57 95.0 229 95.4 0.891 History of polycystic ovary Yes 1 1.7 10 4.2 0.849 0.357 No 59 98.3 230 95.8 0.445 History of diabetes mellitus Yes 15 25.0 72 30.0 0.583 0.445 No 45 75.0 168 70.0 0.583 0.445 History of ever breast feeding 27 45.0 53 22.1 12.89 <0.0001						1.031	0.510
No 57 95.0 229 95.4 Image: control of the property							
History of polycystic ovary Yes 1 1.7 10 4.2 0.849 0.357 No 59 98.3 230 95.8 0.849 0.357 History of diabetes mellitus Yes 15 25.0 72 30.0 0.583 0.445 No 45 75.0 168 70.0 0.583 0.445 History of ever breast feeding 27 45.0 53 22.1 12.89 <0.0001	Yes				1	0.019	0.891
Yes 1 1.7 10 4.2 0.849 0.357 No 59 98.3 230 95.8 0.849 0.357 History of diabetes mellitus Ves 15 25.0 72 30.0 0.583 0.445 No 45 75.0 168 70.0 0.583 0.445 History of ever breast feeding 27 45.0 53 22.1 12.89 <0.0001 No 33 55.0 187 77.9 1 <0.0001 HI. Lifestyle attributes Smoking 55 91.7 237 98.8 16.31 <0.0001 Current smoker 1 1.7 3 1.2 0 Was smoker then stopped 4 6.7 0.0 0.0 0		57	95.0	229	95.4		
No 59 98.3 230 95.8 History of diabetes mellitus Yes 15 25.0 72 30.0 0.583 0.445 No 45 75.0 168 70.0 <td></td> <td>1</td> <td>17</td> <td>10</td> <td>1.2</td> <td>0.940</td> <td>0.257</td>		1	17	10	1.2	0.940	0.257
History of diabetes mellitus Yes 15 25.0 72 30.0 0.583 0.445 No 45 75.0 168 70.0 0.583 0.445 History of ever breast feeding Yes 27 45.0 53 22.1 12.89 <0.0001						0.049	0.337
Yes 15 25.0 72 30.0 0.583 0.445 No 45 75.0 168 70.0 0.583 0.445 History of ever breast feeding Yes 27 45.0 53 22.1 12.89 <0.0001		1 37	70.5		75.0		
History of ever breast feeding Yes 27 45.0 53 22.1 12.89 <0.0001	· ·			72		0.583	0.445
Yes 27 45.0 53 22.1 12.89 <0.0001 No 33 55.0 187 77.9 1 <0.0001 III. Lifestyle attributes Smoking Nonsmoker 55 91.7 237 98.8 16.31 <0.0001 Current smoker 1 1.7 3 1.2 0 Was smoker then stopped 4 6.7 0.0 0.0 0.0		45	75.0	168	70.0		
No 33 55.0 187 77.9 1 III. Lifestyle attributes Smoking Nonsmoker 55 91.7 237 98.8 16.31 <0.0001 Current smoker 1 1.7 3 1.2 0 0 Was smoker then stopped 4 6.7 0.0 0.0 0.0		1 0-	1 42 -	1 40		1.5.5.	0.000
III. Lifestyle attributes Smoking Smoking Nonsmoker 55 91.7 237 98.8 16.31 <0.0001		_				-	< 0.0001
Smoking Nonsmoker 55 91.7 237 98.8 16.31 <0.0001 Current smoker 1 1.7 3 1.2 0 Was smoker then stopped 4 6.7 0.0 0.0 0.0		33	55.0	187	17.9	1	
Nonsmoker 55 91.7 237 98.8 16.31 <0.0001 Current smoker 1 1.7 3 1.2 0 Was smoker then stopped 4 6.7 0.0 0.0 0.0							
Current smoker 1 1.7 3 1.2 0 Was smoker then stopped 4 6.7 0.0 0.0		55	91.7	237	98.8	16.31	< 0.0001
Was smoker then stopped 4 6.7 0.0 0.0	Current smoker		_			_	
	Was smoker then stopped	4			1	<u> </u>	

Yes	0	0.0	9	3.8	2.320	0.128
No	60	100.0	231	96.2		
Daily eating fresh fruit & vegetables						
Yes	40	66.7	132	55.0	2.671	0.102
No	20	33.3	108	45.0		
Low fat diet						
Yes	20	33.3	161	67.1	22.84	< 0.0001
No	40	66.7	79	32.9	6	
BMI	Mean ± SI)	Mean ± SD			
	30.9 ± 8.1		32.0± 7.3			
< 18.5	7	11.7	2	.8	22.64	
< 18.5 18.5-24.9		11.7 13.3		.8 15.4	22.64 6	< 0.0001
	7		2			<0.0001
18.5-24.9	7 8	13.3	2 37	15.4		<0.0001
18.5-24.9 25-29.9	7 8 8	13.3 13.3	2 37 64	15.4 26.7		<0.0001
18.5-24.9 25-29.9 30+	7 8 8	13.3 13.3	2 37 64	15.4 26.7		<0.0001
18.5-24.9 25-29.9 30+ Daily tea consumption	7 8 8 8 37	13.3 13.3 61.7	2 37 64 137	15.4 26.7 57.1	6	
18.5-24.9 25-29.9 30+ Daily tea consumption Yes	7 8 8 8 37	13.3 13.3 61.7 56.7	2 37 64 137	15.4 26.7 57.1 64.2	6	
18.5-24.9 25-29.9 30+ Daily tea consumption Yes No	7 8 8 8 37	13.3 13.3 61.7 56.7	2 37 64 137	15.4 26.7 57.1 64.2	6	

^{*} remaining participants are not applicable P value = ≤ 0.05

Table 2: Univariate Regression Analysis for Risk Factors of Ovarian Cancer of Saudi Patients attending King Fahd Specialist Hospital in Dammam- Eastern Province - KSA

Independent variables	variables B P-value OR			95% CI	
•				Lower	Upper
I. Socio-demographic ch	aracteristics				
Age		0.281			
<30	Reference				
30-	-2.576	0.113	0.076	0.003	1.842
50+	-1.291	0.294	0.275	0.025	3.070
Marital status					
Ever married	Reference				
Single	-1.198	0.277	0.302	0.035	2.618
Education level		0.010			
Illiterate	5.785	0.008	325.263	4.621	22896.678
Completed primary/intermediate	1.948	0.042	7.016	1.074	45.814
school	<u> </u>				
Completed secondary school	2.563	0.004	12.971	2.219	75.822
Completed university or more	Reference				
II. Reproductive history &		related data	1		
•					
Age at menarche					
≤ 13	1.020	0.337	2.772	0.346	22.181
> 13	Reference		•	•	•
Age of menopause	•				
≤51	Reference				
>51	-2.922	0.034	.054	0.004	0.798
Parity	•				<u> </u>
≤3	2.184	0.030	8.879	1.233	63.959
>3	Reference	1	1	I.	1
History of multiple births					
Yes	2.511	0.085	12.314	0.704	215.336
No	Reference			1 01.0	1 ======
Use of oral contraceptive pills					
Yes	Reference				
No	-1.138	0.250	0.320	0.046	2.228
Use of intrauterine device		,			
Yes	Reference				
No	-2.083	0.104	0.125	0.010	1.530
History of tubal ligation	2.003	3.101	0.125	0.010	1.550
Yes	Reference				
No	-1.942	0.060	0.143	0.019	1.082
History of infertility	1.7.12	3.000	0.110	0.017	1.002
Yes	0.729	0.905	2.072	0.000	330116.907
No	Reference	0.703	2.072	0.000	330110.707
Use of fertility drugs e.g. Clomid					
Yes	0.247	0.893	1.280	0.036	45.942
No	Reference	0.073	1.200	0.030	+3.744
INO	Reference				

Hormone replacement therapy a	fter menopaus	e						
Yes	0.685	0.678	1.983	0.079	50.054			
No	Reference	0.070	1.505	0.077	20.02			
Family history of ovarian cancer								
Yes	-5.569	< 0.0001	0.004	0.000	0.067			
No	Reference	<0.0001	0.004	0.000	0.007			
Family history of breast cancer	Reference							
Yes	- 0.338	0.703	.713	.125	.4.064			
No	Reference	0.703	./13	.123	.4.004			
Family history of colorectal cand								
Yes	- 0.421	0.757	0.656	0.045	9.487			
No	Reference	0.737	0.030	0.043	9.407			
Personal history of other cancer	Reference							
Yes	1.599	0.312	4.950	0.223	109.971			
No	Reference	0.312	4.930	0.223	109.971			
	Reference							
Talcum powder on genital area Yes	0.838	0.303	2.312	0.469	11.395			
No	Reference	0.505	2.312	0.409	11.373			
History of endometriosis	Kelefelice							
Yes	1 729	0.209	0.178	0.007	4 504			
No	-1.728 Reference	0.298	0.170	0.007	4.594			
History of ovarian cyst	Keierence							
Yes	-3.288	0.036	0.037	0.002	0.804			
No	Reference	0.030	0.037	0.002	0.804			
History of polycystic ovary	Reference							
Yes	5.536	0.020	690 602	1.022	247360.502			
No	Reference	0.029	689.693	1.923	24/300.302			
History of diabetes mellitus	Reference							
Yes	-2.062	0.025	0.127	0.021	0.774			
No	Reference	0.023	0.127	0.021	0.774			
History of ever breast feeding	Reference							
Yes	Reference							
No	- 0.589	0.555	0.555	0.079	3.924			
III. Lifestyle attributes	- 0.369	0.555	0.555	0.079	3.924			
m. Encstyle attributes								
Smoking								
Nonsmoker	Reference							
Ever smoker	- 0.851	0.668	0.427	0.009	20.916			
Alcohol use	0.001	0.000	01.127	0.007	20.710			
Yes	22.085	0.998	3902020594	0.000				
No	Reference	0.770	370 <u>2</u> 02027 1	0.000	II.			
Daily eating of fresh fruit & vege								
Yes	Reference							
No	1.414	0.073	4.114	0.879	19.264			
Low fat diet								
Yes	Reference							
No	1.265	0.055	3.543	0.973	12.901			
BMI			, ,,, ,,		1			
< 18.5	Reference							
18.5-24.9	5.518	0.000	249.163	11.358	5466.187			
25-29.9	- 0.690	0.467	0.501	0.078	3.229			
30+	-1.572	0.088	0.208	0.034	1.267			
Daily tea consumption			,	,				
Yes	1.342	0.099	3.828	0.776	18.891			
No	Reference	0.077	3.020	0.770	10.071			
Practicing sports daily	Reference							
Yes	Reference							
No	-3.854	< 0.0001	0.021	0.003	0.143			
1 110	-5.054	<0.0001	0.041	0.003	0.143			

^{*}P value = ≤ 0.05

Table 3: Multivariate Regression Analysis for Risk Factors of Ovarian Cancer of Saudi Patients attending King Fahd Specialist Hospital in Dammam- Eastern Province - KSA

Independent variables				95% CI of OR			
_	В	P-value	OR	Lower	Upper		
Education level							
Illiterate	4.231	< 0.0001	68.781	10.141	466.501		
Completed primary/intermediate school	1.852	0.001	6.372	2.119	19.157		
Completed secondary school	1.470	0.004	4.348	1.606	11.776		

Completed university or more	Reference						
Age of menopause							
≤ 51	Reference						
>51	1.091	0.012	2.979	1.177	6.947		
Parity							
≤3	1.120	0.021	3.066	1.186	7.928		
>3	Reference						
History of tubal ligation							
Yes	Reference						
No	-1.184	0.060	0.306	0.089	1.050		
Family history of ovarian cancer							
Yes	-2.611	0.001	0.073	0.017	0.326		
No	Reference						
History of ovarian cyst							
Yes	-1.047	0.239	0.351	0.061	2.008		
No	Reference						
History of polycystic ovary							
Yes	1.521	0.274	4.578	0.300	69.912		
No	Reference						
History of diabetes mellitus							
Yes	0.488	0.286	1.628	0.665	3.986		
No	Reference	•		•			
Low fat diet							
Yes	Reference						
No	2.020	< 0.0001	7.536	3.300	17.210		
BMI							
< 18.5	Reference						
18.5-24.9	3.359	0.001	28.758	3.920	210.973		
25-29.9	- 0.284	0.616	0.753	0.248	2.285		
30+	-1.095	0.042	0.334	0.116	0.963		
Practicing sports daily							
Yes	Reference						
No	-1.807	< 0.0001	0.164	0.061	0.442		

References

- [1] Njoud A Alhedyani, Abeer Alsaigh, Laura Alrumayyan, Nada Bohligah, Noura Aldhawi, Alanoud A Alhedyani and Reham Elsayed. (2016); Women's Awareness and Attitude toward Screening of Ovarian Cancer in Riyadh, Saudi Arabia. International Journal of Advanced Research (IJAR) 4 (12). 2189-2198] (ISSN 2320-5407). www.journalijar.com.
- [2] Alghamdi IG, Hussain II, Alghamdi MS, Alghamdi MM, Dohal AA, El-Sheemy MA: (2014); Incidence rate of ovarian cancer cases in Saudi Arabia: an observational descriptive epidemiological analysis of data from Saudi Cancer Registry 2001-2008. Jun 25; 6:639-45. doi: 0.2147/IJWH.S63636. www.Int J Womens Health.
- [3] Cancer Incidence Report Saudi Arabia: (2006) Kingdom of Saudi Arabia, Ministry of Health, Saudi Cancer Registry.
- [4] DeVita VT, Hellman S, Rosenberg SA. Cancer: Principles and practice of Oncology. 8th ed. Baltimore: Lippincott Williams and Willkins; 2008.
- [5] Gates AM, Rosner BA, Hecht JL, Tworoger SS. Risk factors for Epithelial Ovarian Cancer by Histologic Sybtype. Am J Epidemol 2009; 171(1): 45-53.
- [6] Moorman PG, Calingaert B, Palmieri RT, Iversen ES, Bentley RC, Halabi S, et al. Hormonal risk factors for ovarian cancer in premenopausal and postmenopausal women. Am J Epidemiol 2008, 167(9): 1059-69.
- [7] Salehi F, Dunfield L, Phillips KP, Krewski D, Vanderhyden BC. Risk factors for ovarian cancer: an overview with emphasis on hormonal factors J Toxicol Environ Health B Crit Rev 2008;11(3-4): 301-21.
- [8] Modugno F, Ness RB, Wheeler JE. Reproductive risk factors for epithelial ovarian cancer according to histologic type and invasiveness. Ann Epidemiol 2001;11:568–74.
- [9] Kelemen LE, Bandera EV, Terry KL, Rossing MA, Brinton LA, Doherty JA et al. Recent alcohol consumption and risk of incident ovarian carcinoma: a pooled analysis of 5,342 cases and 10,358 controls from the Ovarian Cancer Association Consortium. BMC Cancer. 2013 Jan 22; 13:28. doi: 10.1186/1471-2407-13-28.
- [10] Kingdom of Saudi Arabia, Saudi Health Council, Saudi Cancer Registry, Cancer Incidence Report 2010; p.18 (2013; p.19).
- [11] Nazer A, Al-Badawi IA. Incidence of gynecological malignancy among the Saudi population. Hematol Oncol Stem Cell Ther. 2012; 5:69-70.
- [12] Al-Badawi IA, Munkarah AR, Tulbah A, Babic II, Al Husaini H, Ahmad S. A detailed study of patients and tumor characteristics of epithelial ovarian cancer in Saudi Women. Int J Gynecol Cancer 2013; 23: 456- 460.
- [13] Abu-Zaid A, Nazer A, Alomar O, Azzam A, Al-Eid HS, Elhassan TA, Al-Badawi IA. Incidence of malignant ovarian germ cell tumors (MOGCTs) in Saudi Arabia. Hematol Oncol Stem Cell Ther 2014 Jan 8. pii: S1658-3876(13)00097-6. doi: 10.1016/j.hemonc.2013.12.001. [Epub ahead of print]
- [14] Abdullah LS, Bondagji NS. Histopathological pattern of ovarian neoplasms and their age distribution in the western region of Saudi Arabia. Saudi Med J 2012 Jan;33(1):61-5.
- [15] Al-Shabanah OA, Hafez MM, Hassan ZK, Sayed-Ahmed MM, Abozeed WN, Al-Rejaie SS, Alsheikh AA. Human papillomavirus genotyping and integration in ovarian cancer Saudi patients. Virol J 2013 Nov 20; 10(1):343. doi: 10.1186/1743-422X-10-343.
- [16] Hennessy S, Bilker W B, Berlin J A, Strom BL. Factors influencing the optimal control-to-case ratio in matched case-control studies. American Journal of Epidemiology 1999; 149:195-7.
- [17] Titus-Ernstoff L, Perez K, Cramer DW, Harlow BL, Baron JA, Greenberg ER. Menstrual and reproductive factors in relation to ovarian cancer risk. British Journal of Cancer 2001; 84(5): 714–21.

- [18] Gaitskell K, Green J, Pirie K, Reeves G, Beral V. OP26 Parity and ovarian cancer in the Million Women Study: variation by histological subtype. J Epidemiol Community Health 2014;68:A15-A16
- [19] Pasalich M, Su D, Binns CW, Lee AH. Reproductive factors for ovarian cancer in southern Chinese women. J Gynecol Oncol 2013 Apr; 24(2):135-40.
- [20] Meindl A, Hellebrand H, Wiek C, Erven V, Wappenschmidt B, Niederacher D, et al. Germline mutations in breast and ovarian cancer pedigrees establish RAD51C as a human cancer susceptibility gene. Nat Genet 2010; 42: 410–14.
- [21] Huerta JM, Ardanaz E, Larranaga N, Jirstrom K, Manjer J, Idahl A, et al. Oral contraceptive use and reproductive factors and risk of ovarian cancer in the European Prospective Investigation into Cancer and Nutrition. Br J Cancer 2011; 105(9):1436–42.
- [22] American Cancer Society. Cancer Facts and Figures 2016. Atlanta, GA: American Cancer Society; 2016.
- [23] EH, Dalton SO, Høgdall C, Fagö-Olsen CL, Steding-Jessen M, Osler M, et al. Do stages of disease, comorbidity or access to treatment explain socioeconomic differences in survival after ovarian cancer? A cohort study among Danish women diagnosed 2005-2010. Cancer Epidemiol 2015 Jun; 39(3):353-9.
- [24] Rapp K, Schroeder J, Klenk J, Stoehr S, Ulmer H, Concin H, et al. Obesity and incidence of cancer: a large cohort study of over 145,000 adults in Austria. Br J Cancer 2005 Oct 31; 93(9):1062-7.
- [25] World Health Organization. Obesity: preventing and managing the global epidemic. Report of a WHO consultation. World Health Organ Tech Rep Ser 2000; 894: (i–xii), 1–253.
- [26] Olsen CM, Green AC, Whiteman DC, Sadeghi S, Kolahdooz F, Webb PM. Obesity and the risk of epithelial ovarian cancer: a systematic review and meta-analysis. Eur J Cancer 2007 Mar; 43(4):690-709.
- [27] Purdie DM, Bain CJ, Webb PM, Whiteman DC, Pirozzo S, Green AC. Body size and ovarian cancer: case-control study and systematic review (Australia). Cancer Causes Control 2001 Nov; 12(9):855-63.
- [28] Merritt MA, Cramer DW, Missmer SA, Vitonis AF, Titus LJ, Terry KL. Dietary fat intake and risk of epithelial ovarian cancer by tumour histology. Br J Cancer 2014 Mar 4; 110(5):1392-401.
- [29] Prentice RL, Thomson CA, Caan B, Hubbell FA, Anderson GL, Beresford SA, et al. low-fat dietary pattern and cancer incidence in the women's health initiative dietary modification randomized controlled trial. J Natl Cancer Inst 2007 Oct 17; 99(20):1534-43.
- [30] Gazibara T, Filipović A, Kesić V, Kisiĉ-Tepavcević D, Pekmezović T. Risk factors for epithelial ovarian cancer in the female population of Belgrade, Serbia: a case-control study. Vojnosanit Pregl 2013 Dec; 70(12):1097-102.

Azza Ali El-Mahalli Ali "Risk Factors of Ovarian Cancer in Eastern Province - Saudi Arabia"." IOSR Journal of Nursing and Health Science (IOSR-JNHS), vol. 7, no.2, 2018, pp. 54-63.

DOI: 10.9790/1959-0702065463 www.iosrjournals.org 63 | Page