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## **FULL PAPER**

# **Risk factors of different types of HR-HPV in** patients suffering from high-grade cervical intraepithelial neoplasia (CIN)

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Cancer is a chronic disease with a high prevalence both worldwide and in Iran. The present study was done to investigate the relationship between the frequency of different types of HR-HPV in patients suffering from high-grade cervical intraepithelial neoplasia (CIN) and the demographic characteristics of patients. In this descriptive analytical study, 302 patients with female gender were included. Out of 302 patients examined, the mean (standard deviation) of the patients' age was 34.12 (7.72) years. Regarding education, 25 (8.3%), 71 (23.5%), 97 (32.1%), 101 (33.4%), and 6(2%) had below diploma, diploma, between diploma and bachelors, bachelors and Masters, and PhD, respectively. Regarding Gravid, 63 (20.9%) had grade 1, 50 (16.6%) had grade 2, and 25 (8.3%) had grade 3. The biopsy from lesions was performed by a gynecologist under sterilized conditions according to the guidelines available in gynecology and obstetrics reference books. Next, the taken specimens were sent to the laboratory and their results were investigated. After collection, the data were introduced into SPSS 16, and analyzed through descriptive as well as analytical statistical methods. The results of this study can help clinical physicians to implement evidence-based practice.

* <b>Corresponding Author:</b> Seyedeh Atefeh Kashani Email: atykashani@gmail.com	KEYWORDS
Tel.: +98 21 23871	Cancer; HPV; cervical intraepithelial neoplasia; risk factors.

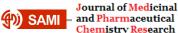
## Introduction

Women's health as a group of individuals claiming a large population of society is notable as one of the most significant issues in the public health. Considering the role of women at home, place of residence, society, as well as the vulnerability of these individuals, in case of presence of any disease in them, numerous problems can occur in the patient and their family. Different chronic diseases afflict this group of society, including

cardiovascular disease. lung diseases. infectious diseases, psychological disorders and cancer [1-8].

Cancer is a chronic disease with a high prevalence both worldwide and in Iran [9-12]. One type of this disease is cervical cancer, which is the fourth most common and the fourth main cause of mortality by cancer in women in 2020 [13]. Cervical cancer is one of the causes of mortality resulting from cancer in developing countries. World Health Organization has estimated that by 2030 this

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cancer would be the cause of death of around 474000 women, with 95% of these mortalities occurring in low- and middle-income countries. Meanwhile, according to the report issued by Iran's cancer registration national Center, in 20909, its prevalence was around 2.19 cases per 100,000, standing 11<sup>th</sup> among all cancers of Iranian women [14-16].

Most cases of cervical cancer results from infection with HPV as the most common sexually transmitted infection worldwide. Regular screening if accompanied by treatment in the early stages alongside vaccination against human papilloma virus (HPV) can be prevented [17-19]. HPV is one of the most common sexually transmitted infections, which is the main cause of incidence of infection attributed cancer cases worldwide [20]. HPV is a small double-stranded DNA virus which infects the patient. Different types of HPV are classified according to genomic structure as well as the propensity to the human epithelial tissue [21].

The coverage of HPV vaccination is also affected by different factors, which is lower in developing and less-developed countries [19]. The aggressive cervical cancer because of having a long period of no invasion, availability of suitable screening programs, and effective treatment of the initial lesions has been known as a preventable cancer [22].

# Methodology

The present study was done to investigate the relationship between the frequency of different types of HR-HPV in patients suffering from high-grade cervical intraepithelial neoplasia (CIN) and the demographic characteristics of patients.

In this descriptive analytical study, with the ethics code of IR.SBMU.RETEC.REC.1400.1196, 302 patients with female gender were included.

In this study, the sampling method was census whereby the women referring to the governmental gynecology clinics because of existence of genital lesions were included. The inclusion criteria were informed consent for participation in the study, presence of the risk factors confirmed for cervical neoplasia, some evidence suggesting abnormal examination, as well as abnormal Pap smear test result. The exclusion criteria included having other diseases that would influence the conditions of the study as well as lack of patient consent to continue their participation (at any time of conducting the research).

The instrument utilized in the research included two sections of the demographic characteristics form and the patient clinical examination (Pap smear, HPV type and colposcopy, test, etc.). The mentioned form included questions about age, education, marital status, employment status, type of method of contraception, number of sexual partners, history of taking corticosteroid drugs, number of pregnancies and methods of delivery, history of STDs in the spouse, immunodeficiency, etc.

The biopsy from lesions was performed by a gynecologist under sterilized conditions according to the guidelines available in gynecology and obstetrics reference books. Next, the taken specimens were sent to the laboratory and their results were investigated. After collection, the data were introduced into SPSS 16, and analyzed through descriptive as well as analytical statistical methods.

# Results

Out of 302 patients examined, the mean (standard deviation) of the patients' age was 34.12 (7.72) years. Regarding education, 25 (8.3%), 71 (23.5%), 97 (32.1%), 101 (33.4%), and 6 (2%) had below diploma, diploma, between diploma and bachelors, bachelors and Masters, and PhD, respectively. Regarding Gravid, 63 (20.9%) had grade 1, 50 (16.6%) had grade 2, and 25 (8.3%) had grade 3 (Tables 1 and 2).

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TABLE 1 Status	of variables bas	sed on HPV1 result 66
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	All CIN (n=302)	colpo1 result CIN1	colpo1 result CIN2	colpo1 result CIN3	colpo1 result	colpo1 result CIN23
HPV types						
HPV1result16		25	6	8	2	4
HPV1result18		2	0	0	1	0
HPV1result31		2	1	1	0	0
HPV1result33		1	1	0	0	0
HPV1result35		1	0	0	0	0
HPV1result39		2	0	1	0	0
HPV1result45		1	0	0	0	0
HPV1result51		3	0	1	0	0
HPV1result52		4	0	0	0	0
HPV1result56		0	1	0	0	0
HPV1result58		1	0	0	0	0
HPV1result59		2	0	0	0	0
HPV1result66		1	0	0	0	0
HPV1result1618HI		1	1	0	0	0
HPV1result1618LOW		1	0	0	0	0
HPV1result1618HILOW		1	1	0	0	0
HPV1result68		1	0	0	0	0
HPV1resultOTHER		23	7	7	0	0
HPV1resultHIOTHER		22	2	3	0	0
HPV1resultNEG		3	0	0	0	1
HPV1resultHIRISK		15	4	3	0	1
HPV1resultLowRisk		11	0	0	2	0
HPV1result16LowHiRisk		4	2	0	0	0
HPV1result16HiRisk		13	4	1	2	1
HPV1result18LowHiRisk		0	3	0	0	0
HPV1result18HiRisk		3	0	0	0	0
HPV1result16LOW		3	3	1	0	1
HPV1result18Low		2	1	0	0	0
HPV1result1618		0	1	1	0	1

**TABLE 2** Status of variables used in cancer

-	Yes	No	None	Total
HPV types				
HPV1 result 16	51(16.9)	227(75.2)	23(7.6)	301(99.7)
HPV1 result 18	4(1.3)	274(90.7)	23(7.6)	301(99.7)
HPV1 result 31	5(1.7)	273(90.4)	23(7.6)	301(99.7)
HPV1 result 33	2(0.7)	276(91.4)	23(7.6)	301(99.7)
HPV1 result 35	1(0.3)	277(91.7)	23(7.6)	301(99.7)
HPV1 result 39	3(0.1)	275(91.1)	23(7.6)	301(99.7)
HPV1 result 45	2(0.7)	276(91.4)	23(7.6)	301(99.7)
HPV1 result 51	5(1.7)	273(90.4)	23(7.6)	301(99.7)
HPV1 result 52	4(1.3)	274(90.7)	23(7.6)	301(99.7)
HPV1 result 56	1(0.3)	277(91.7)	23(7.6)	301(99.7)
HPV1 result 58	1(0.3)	277(91.7)	23(7.6)	301(99.7)
HPV1 result 59	2(0.7)	276(91.4)	23(7.6)	301(99.7)
HPV1 result 66	1(0.3)	277(91.7)	23(7.6)	301(99.7)
HPV1 result 1618 HI	2(0.7)	1(0.3)	275(91.1)	301(99.7)
HPV1 result 1618 LOW	1(0.3)	277(91.7)	23(7.6)	301(99.7)
HPV1 result 1618 HILOW	2(0.7)	276(91.4)	23(7.6)	301(99.7)
HPV1 result 68	2(0.7)	276(91.4)	23(7.6)	301(99.7)

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HPV1 result OTHER	41(13.6)	237(78.5)	23(7.6)	301(99.7)
HPV1 result HI OTHER	31(10.3)	247(81.8)	23(7.6)	301(99.7)
HPV1 result NEG	8(2.6)	270(89.4)	23(7.6)	301(99.7)
HPV1 result HI RISK	24(7.9)	254(84.1)	23(7.6)	301(99.7)
HPV1 result Low Risk	14(4.6)	264(87.4)	23(7.6)	301(99.7)
HPV1 result 16 Low Hi Risk	6(2)	272(90.1)	23(7.6)	301(99.7)
HPV1 result 16 Hi Risk	23(7.6)	255(84.4)	23(7.6)	301(99.7)
HPV1 result 18 Low Hi Risk	4(1.3)	274(90.7)	23(7.6)	301(99.7)
HPV1 result 18 Hi Risk	3(1)	275(91.1)	23(7.6)	301(99.7)
HPV1 result 16 LOW	9(3)	269(89.1)	23(7.6)	301(99.7)
HPV1 result 18 Low	3(1)	275(91.1)	23(7.6)	301(99.7)
HPV1 result 1618	4(1.3)	274(90.7)	23(7.6)	301(99.7)

## Conclusion

When a patient develops chronic diseases, all of their life dimensions are affected [6,23-25]. In different studies, the prevalence of cancer as well cervical cancer has been examined in Iran. In the study by Hajbagheri *et al.*, to examine the frequency of human papilloma virus as well as its genotypes in women suffering from genital lesions, 50 women living in Sanandaj, Iran were included. According to the findings, in 56% of the studied samples, HPV genome was identified; the frequency of the genotypes presents for genotype 6 was 32%, 18 was 12%, 16 was 6%, and 11 was 2% (2%) [26,27].

Cervical cancer is one of the most important groups of cancer in women, whose prevalence has been examined in studies conducted in Iran as well as in other countries. In the study by Kalliala *et al.*, investigating 27 eligible studies, the incidence of cervical cancer following CIN treatment in the year was 39 per 100,000. The relative risk was higher for women older than 50 years, and the relative risk of vaginal and rectal cancer was higher [28].

In the study by Loopik *et al.*, examining the cervical neoplasia and risk of cervical cancer, it was found that in case of developing cervical intraepithelial neoplasia (CIN) at grade 3, the risk of developing cervical cancer would double. Furthermore, at older ages, advanced age and women with recurrent CIN3 had higher chance of developing cervical cancer [29]. In the study by Damgaard *et al.*, examining 23-to-40-year-old women under screening, the results of histopathological

examinations of patients were inspected. According to the findings, out of 3623 patients with CIN2, the prevalence of any type of highrisk HPV was 87%, and HPV 16 was found as the most common genotype; HPV 16 in women of 30 years or younger was equal to 39.3%, while being 25.2% in women at age of 30 and above. Ultimately, the high prevalence of HPV 16 and the high-grade cytology in women was regarded a high-risk population [30].

## Suggestions for further research

The results of this study can help clinical physicians to implement evidence-based practice.

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#### **Authors' Contributions**

All authors contributed to all stages of the article.

## **Conflict of Interest**

The authors declare that they have no conflict of interest in this study.

## **Data Availability**

All data produced and examined are incorporated in this article.

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# References

[1] M. Shokri, A. Tarjoman, M. Borji, L. Solaimanizadeh, Investigating psychological problems in caregiver of pediatrics with cancer: A systematic review, *Journal of Child and Adolescent Psychiatric Nursing*, **2020**, *33*, 229-238. [Crossref], [Google Scholar], [Publisher]

[2] E. Khalighi, A. Tarjoman, A. Abdi, M. Borji, The prevalence of delirium in patients in Iran: a systematic review and meta-analysis, *Future Neurology*, **2019**, *14*, 34. [Crossref], [Google Scholar], [Publisher]

[3] B. Kazeminezhad, A. Tarjoman, M. Borji, Relationship between praying and self-care in elderly with heart failure: A cross-sectional study in west of Iran, *Journal of Religion and Health*, **2020**, 59, 19-28. [Crossref], [Google Scholar], [Publisher]

[4] M. Hatefi, R. Parvizi, M. Borji, A. Tarjoman, Effect of self-management program on pain and disability index in elderly men with osteoarthritis, *Anesthesiology and Pain Medicine*, **2019**, *9*. [Crossref], [Google Scholar], [Publisher]

[5] F. Tahmasbi, A. Madani Neishaboori, M. Mardani, A. Toloui, K. Komlakh, Y. Azizi, M. Yousefifard, Efficacy of polyarginine peptides in the treatment of stroke: A systematic review and meta-analysis, *Brain and Behavior*, **2023**, *13*, 2858. [Crossref], [Google Scholar], [Publisher]

[6] K. Komlakh, A. Karbasfrushan, The effect of Pregabalin on the pain status of patients with disc and spinal surgeries: A systematic review of drug therapy, *Eurasian Chemical Communications*, **2022**, *4*, 1147-1155. [Crossref], [Google Scholar], [Publisher]

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Journal of Medicinal

and Pharmaceutical Chemistry Research

[7] A. Rahmatian, E. Bastani, F. Shokri, A. Karbasfrushan, Prevalence of hemiplegic shoulder pain in Iran: a systematic review and meta-analysis, *Anesthesiology and Pain Medicine*, **2023**, *13*. [Crossref], [Google Scholar], [Publisher]

[8] S. Fazelipour, F.A. Moghadam, P. Davudi, Z. Tootian, F. Assadi, Histometrical study of ovarian follicles of immature mice treated with methylphenidate, **2015**. [Google Scholar], [Publisher]

[9] E. Bastani, F. Shokri, Incidence trend of lung cancer in Iran: A systematic review and meta-analysis, *International Journal of Cancer Management*, **2023**, *16*. [Crossref], [Google Scholar], [Publisher]

[10] H. Nourmohammadi, M. Motaghi, M. Borji, A.Tarjoman, B. Soltany, The effects of reflexology on fatigue severity of patients with cancer, *Asian Pacific Journal of Cancer Prevention: APJCP*, **2019**, *20*, 391. [Crossref], [Google Scholar], [Publisher]

[11] M. Borji, A. Tarjoman, A. Abdi, M. Otaghi, Efficacy of implementing home care using eye movement desensitization and reprocessing in reducing stress of patients with gastrointestinal cancer, *Asian Pacific Journal of Cancer Prevention: APJCP*, **2019**, *20*, 1967. [Crossref], [Google Scholar], [Publisher]

[12] E. Bastani, M. Rizehbandi, F. Shokri, Prevalence of pain and factors affecting it in patients with lung cancer in Ilam, *International Journal of Cancer Management*, **2024**, *17*. [Crossref], [Google Scholar], [Publisher]

[13] H. Sung, J. Ferlay, R.L. Siegel, M. Laversanne, I. Soerjomataram, A. Jemal, F. Bray, Global cancer statistics 2020: GLOBOCAN estimates of incidence and mortality worldwide for 36 cancers in 185 countries, *CA: a Cancer Journal for Clinicians*, **2021**, *71*, 209-249. [Crossref], [Google Scholar], [Publisher]

Page | 1127





[14] S. Ozcelik, N. Ozpinar, S. Karakus, F. Akyildiz, O. Karakaya, Metronidazole resistance in Trichomonas vaginalis determined by molecular and conventional methods, **2018**. [Google Scholar], [Publisher]

[15] M. Sharifi, Z. Mohammadi, Z. Makvandi, P. Rostami, A. Moradi, Assessment of cervical cancer screening and its barriers in 18-50 year old women referring to Asad Abad comprehensive health centers, *Pajouhan Scientific Journal*, **2018**, *16*, 35-45. [Google Scholar], [Publisher]

[16] F. Farzaneh, S. Mohammadi, E. Ghaffari, A. Hosseini, S. Younesi, M.M. Taheri Amin, P. Balvayeh, F. Navid Pour, S. Noori Ardebili, Frequency of HR-HPV types in patients with high grade cervical intraepithelial neoplasia (CIN), *Journal of Obstetrics, Gynecology and Cancer Research*, **2021**, *6*, 122-127. [Crossref], [Google Scholar], [Publisher]

[17] L. Peirson, D. Fitzpatrick-Lewis, D. Ciliska, R. Warren, Screening for cervical cancer: a systematic review and meta-analysis, *Systematic Reviews*, **2013**, *2*, 1-14. [Crossref], [Google Scholar], [Publisher]

[18] N.G. Campos, M. Sharma, A. Clark, K. Lee, F. Geng, C. Regan, J. Kim, S. Resch, The health and economic impact of scaling cervical cancer prevention in 50 low-and lower-middle-income countries, *International Journal of Gynecology & Obstetrics*, **2017**, *138*, 47-56. [Crossref], [Google Scholar], [Publisher]

[19] J.M. Lemp, J.W. De Neve, H. Bussmann, S. Chen, J. Manne-Goehler, M. Theilmann, M.E. Marcus, C. Ebert, C. Probst, L. Tsabedze-Sibanyoni, L. Sturua, Lifetime prevalence of cervical cancer screening in 55 low-and middle-income countries, *Jama*, **2020**, *324*, 1532-1542. [Crossref], [Google Scholar], [Publisher]

[20] D. Gökengin, T. Noori, A. Alemany, C. Bienkowski, G. Liegon, A.Ç. İnkaya, J. Carrillo, G. Stary, K. Knapp, O. Mitja, J.M. Molina, Prevention strategies for sexually transmitted infections, HIV, and viral hepatitis in Europe, *The Lancet Regional Health–* 

*Europe*, **2023**, *34*. [Crossref], [Google Scholar], [Publisher]

[21] M. Shahi, S.A.A. Shamsian, M. Ghodsi, A. Shafaei, Prevalence and genotype distribution of human papillomavirus in a population of Iranian men, *Journal of Mazandaran University of Medical Sciences*, **2022**, *32*, 147-154. [Google Scholar], [Publisher]

[22] Z. Vafaeinezhad, Z. Kazemi, M. Mirmoeini, H. Piroti, E. Sadeghian, M. Mohammad Ali-Vajari, N. Fattah, M. Jafari, Trends in cervical cancer incidence in Iran according to national cancer registry, *Journal of Mazandaran University of Medical Sciences*, **2018**, *28*, 108-114. [Google Scholar], [Publisher]

[23] M. Karimian, R. Ranjbar, M. Salamati, A. Adibi, F. Kazemi, M. Azami, Prevalence of dyspepsia in Iran: a systematic review and meta-analysis, *Archives of Iranian Medicine*, **2021**, *24*, 568-578. [Crossref], [Google Scholar], [Publisher]

[24] S. Tardeh, A. Adibi, A.A. Mozafari, Prevalence of suicide ideation and attempt during COVID-19 pandemic: A systematic review and meta-analysis, *International Journal of Preventive Medicine*, **2023**, *14*, PMC10023846. [Crossref], [Google Scholar], [Publisher]

[25] M. Khosravi, D. De Berardis, S. Mazloom, A. Adibi, N. Javan, Z. Ghiasi, M. Nafeli, N. Rahmanian, Oropharyngeal microbiome composition as a possible diagnostic marker for true psychosis in a forensic psychiatric setting: A narrative literature review and an opinion, *Electronic Journal of General Medicine*, **2023**, *20*, em486. [Crossref], [Google Scholar], [Publisher]

[26] M. Hatefi, K. Komlakh, The effect of Atorvastatin on chronic subdural hematoma status: A systematic review of drug therapy, *Eurasian Chemical Communications*, **2022**, 4, 1130-1137. [Crossref], [Google Scholar], [Publisher]

[27] K. Hajibagheri, A. Abaszade, S. Afrasiabian,F. Verdi, D. Roshani, F. Abdi, M. Mohsenpour,Frequency of human papilloma virus genotypes Among woman with genitalia lesion,

Sanandaj, Iran, *Scientific Journal of Kurdistan University of Medical Sciences*, **2018**, *23*, 46-52. [Google Scholar], [Publisher]

[28] I. Kalliala, A. Athanasiou, A.A. Veroniki, G. Salanti, O. Efthimiou, N. Raftis, S. Bowden, M. Paraskevaidi, K. Aro, M. Arbyn, P. Bennett, Incidence and mortality from cervical cancer and other malignancies after treatment of cervical intraepithelial neoplasia: a systematic review and meta-analysis of the literature, *Annals of Oncology*, **2020**, *31*, 213-227. [Crossref], [Google Scholar], [Publisher]

[29] D.L. Loopik, J. IntHout, R.M. Ebisch, W.J. Melchers, L.F. Massuger, A.G. Siebers, R.L. Bekkers, The risk of cervical cancer after cervical intraepithelial neoplasia grade 3: A population-based cohort study with 80,442 women, *Gynecologic Oncology*, **2020**, *157*, 195-201. [Crossref], [Google Scholar], [Publisher]

[30] R.K. Damgaard, D. Jenkins, M.H. Stoler, M.M. van de Sandt, K.D. Lycke, M.N. de Koning,



Page | 1129

W.G. Quint, T. Steiniche, L.K. Petersen, A. Hammer, High prevalence of HPV16 and highgrade cytology in women undergoing active surveillance for cervical intraepithelial neoplasia grade 2, *Acta Obstetricia et Gynecologica Scandinavica*, **2023**, *102*, 1227-1235. [Crossref], [Google Scholar], [Publisher]

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