

## **Research Article**

# HPV Vaccination Prevalence among Lebanese Female University Students: A Cross-Sectional Study

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Background. Cervical cancer is the fourth most common cancer in women, worldwide. Most often, cervical cancer is caused by a human papillomavirus (HPV) infection. A lack of studies concerning HPV knowledge and vaccination among the Lebanese population is noticed. We aim to assess the prevalence of HPV vaccine administration among female university students in Lebanon alongside determining the predictors influencing vaccine uptake. Lastly, knowledge scores concerning HPV and HPV vaccination are also computed. Methods. This is a cross-sectional analytical study. It is an anonymous, online-based, close-ended questions survey conducted between the 24th of February 2021 and the 30th of March 2021. Our questionnaire was addressed to females aged between 17 and 30 years and enrolled in a Lebanese university. The collected data were analyzed using Statistical Package for Social Sciences (SPSS) v.26. We used bivariate analysis to compare the rate of vaccination with different variables. We used the chi-square test for categorical variables and Student's t-test for continuous variables. Logistic linear regression was conducted between the state of vaccination and other statistically significant variables from the bivariate analysis. Results. We have received a total of 454 questionnaires. Among the respondents, only 18.9% had received a minimum of one dose of the HPV vaccine. The mean age of up taking the first dose of the vaccine was 17.5 years. In addition, 48% of the respondents were not willing to take the HPV vaccine during the upcoming year. Barriers to not taking the HPV vaccine were mainly the low levels of knowledge concerning HPV and its vaccine. In the multivariate analysis, three predictors were found to affect the rate of HPV vaccination: university type, paternal educational level, and HPV vaccine knowledge score. In detail, a student enrolled in a public university had 77% likelihood of not being vaccinated. Furthermore, female students with a paternal educational level higher than a university degree had 88% probability of being vaccinated. Finally, every increase in the HPV vaccination knowledge of one point increased the likelihood of being vaccinated by 37%. Conclusion. A low vaccination rate among female university students in Lebanon was noticed in our study. In addition, a lack of HPV and HPV vaccine knowledge was found in our population. An awareness campaign alongside public vaccination programs is recommended in order to accomplish higher rates of HPV immunization.

## 1. Background

Human papillomaviruses (HPVs) are the predominant etiologies of sexually transmitted diseases. It is estimated that close to 80% of people worldwide would have contracted at least one strain of HPV by the age of 45 [1] and that the estimated global HPV prevalence is 11.7% across all countries. Studies on different Arab nations revealed an HPV infection prevalence of 10.2% and an overall prevalence of 3.7% for high-risk oncogenic HPV types [2]. More than 20 HPV types fall into that category, especially HPV 16 and 18, which are incriminated in the occurrence of cervical

cancer. HPV types 6, 11, 42, and 43 fall into the low-risk category [3]. Sexual behavior was identified as the main risk factor for HPV acquisition [4].

Alongside sexual hygiene, preventive measures can be taken to reduce HPV infectivity and transmission. The World Health Organization (WHO) clearly endorses HPV vaccines and encourages nationwide vaccination campaigns, as these vaccines contribute to low mortality rates attributable to cervical cancer. By 2017, according to the WHO worldwide HPV vaccination report, 71 countries (37% of all nations) introduced HPV vaccines in their national immunization schedule for girls, among them only 11 countries (6% of all nations only) also included vaccines for boys [5]. To date, Lebanon lacks a national strategy for mass HPV vaccination, which is unfortunately not mentioned in vaccination calendars.

Since 2006, three vaccines were licensed and put on the market for public use. The first to be licensed is the quadrivalent vaccine or Gardasil, targeting HPV types 6, 11, 16, and 18. The licensure of Gardasil was subsequently followed by the bivalent Cervarix in 2007 directed solely against high-risk types 16 and 18. A more inclusive Gardasil 9 was licensed in 2014, targeting the same strains as Gardasil, in addition to less prevalent types, HPV-31, 33, 45, 52, and finally 58.

In 2016, researchers investigated the evolution of sexual life in Lebanese youth across 17 universities. They found that 15% of surveyed subjects had already engaged in sexual activity and that 20% were regularly sexually active, while only 36% used condoms. This particular risky behavior shows that turning a blind eye toward HPV and the importance of prophylaxis is no longer an option [6].

Local studies set the prevalence of HPV vaccination in female Lebanese school children at 2.5% and 16.5% in a small sample of 215 students of a prestigious private school in Beirut [7, 8]. Unfortunately, a severe lack of literature on the subject exists, since the available studies only concerned a specific population, making any generalization on the subject biased.

Unfortunately, both official national recommendations for cervical cancer screening practices and a national HPV immunization program do not exist in Lebanon [9]. With sexually transmitted disease rates on the rise due to risky behavior among university students [6], specifically HPV, and given the lack of a clear national vision for combatting cervical cancer through vaccination, this study aimed to estimate the prevalence of HPV vaccine uptake in female university students in Lebanon. In addition, we aimed to highlight the determinants of HPV vaccine uptake decision in an effort to ultimately launch the appropriate campaigns.

### 2. Methods

2.1. Study Design. This study was an observational crosssectional study conducted between the 24th of February 2021 and the 30th of March 2021 using the online questionnaire platform, Microsoft Forms.

2.2. Ethical Consideration. The study was reviewed and approved by the ethical review board of INSPECT (IRB Approval No. 2022REC-005-INSPECT-01-21). Data were

collected anonymously with no identifying or sensitive information. Students were introduced to the subject and the aim of our study. Moving forward in the online questionnaire was considered to be informed consent.

2.3. Study Population and Sampling. We included in this study female university students aged between 17 and 30 years and residing in Lebanon. Due to the pandemic and the unreachable target population, we were obliged to resort to a nonprobability sampling method, which is "snowball" sampling, following a "cascading progression." A network of informants (trustworthy university students) in all the governorates was recruited to aid our sampling efforts and spread our survey to their peers. Our questionnaire was distributed via a URL in addition to an informative text illuminating the aim of our study. This URL was sent via messages to potential respondents and class representatives.

2.4. Minimal Sample Size Calculation. Considering that 119638 female students were enrolled in Lebanese universities [10] and that the expected prevalence of HPV vaccination in Lebanon is around 16.5% [7], a minimal sample size of 211 was calculated using the Epi Info<sup>TM</sup> software v7.2 (confidence level of 95% and an acceptable margin of error of 5%).

2.5. Questionnaire. A well-structured questionnaire was developed after a rigorous literature review related to the following research objective: expert opinions on the importance, intelligibility, and clarity of the questionnaire's content were considered before the final version was distributed. An initial stage of preliminary testing took place before surveying to make sure that the phrasing of the questions is clear and to check the length of time till completion of the questionnaire. On average, the questionnaire took 7 minutes to complete.

The questionnaire was divided into 8 sections and a total of 79 questions. The initial four sections collected data related to the sociodemographic and behavioral characteristics as well as the medical history and sexual behavior of the participants. The remaining sections of the survey included questions on attitudes towards HPV vaccination, in addition to questions evaluating participants' knowledge concerning HPV and its vaccine.

2.6. Statistical Analysis. The collected data were assembled, coded, and analyzed using SPSS v.26 (IBM SPSS Inc., Chicago, IL, USA). Continuous variables were reported as means and standard deviations (SD), while categorical variables were summarized as frequencies and percentages. Two indexes on the knowledge of HPV and the HPV vaccine were calculated. Every correct answer to single response questions, every correct answer was given 1 point. However, for multiple response questions, every correct answer was given 1 point while not choosing incorrect responses was also rewarded a point. HPV knowledge index was over 28 and the HPV vaccine knowledge index was over 12. In the bivariate analysis, the chi-square test was used for categorical independent variables and an independent sample *T*-test and Fisher's exact

test for continuous variables. Finally, a binary logistic regression was used with having been vaccinated with at least one HPV vaccine dose as the dependent variable of interest. We included in the model all the independent variables with a *p* value <0.2 in the bivariate analysis. A *p* value of  $\leq 0.05$  as a significant statistical difference and a confidence interval of 95% were considered.

#### 3. Results

3.1. Sociodemographic, Medical, Behavioral, and Sexual Characteristics. A total of 454 questionnaires were filled by participants who met the inclusion criteria. The age range varied between 17 years and 30 years with a mean of 22.7 years of age (±3.33). The majority (95.2%) of participants were Lebanese and almost half of them were majoring in nonmedical fields. Less than half of the students (39.4%) were enrolled in the Lebanese university (public). Almost half of the participants (44.7%) had a maternal educational level below a university degree and 58.6% had a paternal educational level below a university degree. Concerning previous medical history, only 1.3% had a personal history of cancer and only 1.5% had a familial history of cancer(s) related to HPV. The majority of the participants (94.3%) have never had a sexually transmitted disease. Regarding behavioral characteristics, around half of the students consume alcohol occasionally and 78.9% do not smoke. Concerning sexual behavior characteristics, almost twothirds of the responders (65.9%) were not sexually active. In addition, only 10% of the currently sexually active participants never used a condom during sexual intercourse. Regarding the usual number of partners, only 3.3% had two or more sexual partners. Additional and detailed results about sociodemographic characteristics, medical history, and behavioral characteristics are found in Table 1.

3.2. HPV Vaccination. The prevalence of taking at least one dose of the HPV vaccine was 18.9%. The mean age of taking the first dose was 17.5 years ( $\pm$ 4.89 years). The age of first dose administration ranged between 9 years and 30 years. The main reported reasons for not taking the HPV vaccine were not knowing about HPV (31.2%) and its vaccine (26.6%) and not being sexually active (26.3) (Figure 1). Additional descriptive data concerning the HPV vaccination are available in Table 2. In addition, barriers to not taking the vaccine are represented in Figure 1.

3.3. HPV Knowledge. The HPV knowledge index varied from 3 to 25 with a mean score of 13.9 over 28. Detailed participants' responses concerning HPV knowledge are presented in Table 3. Regarding the modes of transmission of HPV, 45.3% of the participants reported sexual intercourse, only 12.8% reported skin-to-skin contact, and 18.6% and 19.5% reported nonpenetrative sex and childbirth, respectively. More than half of the participants (58.4%) reported that HPV can be transmitted even in the asymptomatic phase, 26.4% reported that an HPV infection can lead to cervical cancer, and 35.5% reported that HPV can lead to infertility. The majority of the respondents (78.9%) reported that condom use will reduce the possibility of HPV transmission and know that multiple sexual partners can increase the risk of infection (80.6%). Only 11.7% reported that circumcision prevents HPV infection and transmission. More than half of the respondents (58.6%) reported that HPV infections can be present for many years asymptomatically. Finally, only 5.9% of the participants are aware that an HPV infection cannot be tested in men.

3.4. HPV Vaccine Knowledge. HPV vaccine knowledge index had a mean of 4.4 over 12 with a standard deviation of 3.4 and ranged between 0 and 12. Detailed respondents' answers to HPV vaccine knowledge questions are presented in Table 4. Around a third of the respondents (33.9%) reported that the youngest age for HPV vaccine administration is 9 years. Only 27.1% reported that HPV vaccination can prevent oropharyngeal cancer, 33.9% reported that it can prevent penile cancer, and 33.3% reported that it can prevent anal cancer. In addition, only 16.5% reported that the HPV vaccine cannot prevent prostate cancer. Less than half (45.4%) reported that HPV vaccination is more effective if given before the first sexual intercourse. Only 28% reported that HPV-vaccinated persons are not fully immunized against HPV infection. Finally, around two-thirds (67.4%) reported that despite a full course of HPV vaccination, it is not safe to have unprotected sex.

3.5. Bivariate Analysis. Tables 5–7 showcase the bivariate analysis and different associations between HPV vaccination status and sociodemographic characteristics, medical history and habits (Table 5), sexual behavior (Table 6), and knowledge scores (Table 7).

3.6. Multivariate Analysis. Logistic binary regression was performed using the vaccination status variable as the dependent variable. Given that the smallest group of this variable was the HPV-vaccinated group with N = 86, we chose the maximal number of predictors (10%) as 8. From the previously performed bivariate analysis, we had a total of eleven variables with a p value of less than 0.2. Therefore, the predictors of our multivariate analysis were chosen depending on their relevance and correlation with our dependent variable. Table 8 shows the final variables that were entered into the model using the forward likelihood ratio (LR) method.

Table 8 also shows the factors associated with having been vaccinated with at least one HPV vaccine dose. Receiving at least one dose of the HPV vaccine was found associated with being enrolled in a private university compared to enrollment in a public university (ORa = 0.228, p < 0.001). Higher paternal educational level was significantly associated with having been vaccinated (ORa = 1.881, p = 0.027). Finally, a higher HPV vaccine knowledge index was significantly associated with being vaccinated where an increase of 1 point in the index was related to a 37% increase in the odds of having been vaccinated with at least one HPV vaccine dose.

Variable		n (%)	Mean ± SD	Min	Max
Sociodemographic characteristics					
Age (years)			$22.7 \pm 3.33$	17	30
Nationality	Lebanese	432 (95.2)			
Wationality	Others <sup>1</sup>	22 (4.8)			
Major	Medical <sup>2</sup>	219 (48.2)			
1111/01	Nonmedical	235 (51.8)			
University	Public	179 (39.4)			
Oniversity	Private	275 (60.6)			
	Married	39 (8.6)			
Family status	Single	278 (61.3)			
	In a relationship/engaged	137 (30.2)			
	Beirut	95 (20.9)			
Residence	Mount Lebanon	277 (61)			
	Others <sup>3</sup>	82 (18.1)			
	Christian	287 (63.2)			
Religion	Muslim/Druze	179 (27.7)			
-	Nonreligious	18 (4)			
	Less than university degree	203 (44.7)			
Maternal educational level	University degree or higher	251 (55.3)			
	Less than university degree	266 (58.6)			
Paternal educational level	University degree or higher	188 (41.4)			
Medical history					
Demograph biotomy of someon	Yes <sup>4</sup>	6 (1.3)			
Personal history of cancer	No	448 (98.7)			
Family history of cancer related to HPV <sup>5</sup>	Yes	7 (1.5)			
Family mistory of cancer related to HPV	No history	447 (98.5)			
Histomy of convelley transmitted disease	At least once	26 (5.7)			
History of sexually transmitted disease	Never	428 (94.3)			
Behavioral characteristics					
	No	167 (3)			
Alcohol consumption	Frequently	88 (19.4)			
	Occasionally	199 (43.8)			
Smoking	Yes	96 (21.1)			
	No	358 (78.9)			
Sexual characteristics					
Age on first sexual intercourse (years)			$21.08 \pm 4.02$	14	30
	Always	38 (8.4)			
Condom use	Frequently	35 (7.7)			
Condonii use	Never	46 (10.1)			
	Not currently sexually active	335 (73.8)			
	One	140 (30.8)			
Number of partners in general	Two or more	15 (3.3)			
	Not sexually active	299 (65.9)			
	Man	145 (31.9)			
Derther(a) gender	Woman	6 (1.3)			
Partner(s) gender	Both	4 (0.9)			
	No sexual partner	299 (65.9)			
	Anal giving or receptor	7 (1.5)			
Correct maniforming of (mariletical and the start and the	Oral giving or receptor	73 (16.1)			
Sexual preference (multiple choice question)	Vaginal sex	140 (30.8)			
	Not sexually active	299 (65.9)			

TABLE 1: Sociodemographic, behavioral, and sexual characteristics and medical history of study participants (n = 454).

SD: standard deviation. LBP: Lebanese pound. <sup>1</sup>Other nationalities include Palestinian (n = 9), Syrian (n = 8), Iraqi (n = 3), Filipino (n = 1), and Brazilian (n = 1). <sup>26</sup>Medical students" include students of medicine, nursing, and pharmacy. <sup>3</sup>Aakar, North, Baalbeck-Hermel, Beqaa, South, and Nabatiyeh. <sup>4</sup>Breast cancer; CML, non-Hodgkin lymphoma; hodgkin lymphoma (x2); and kidney cancer. <sup>5</sup>Cervical cancer; penile cancer; and coropharyngeal cancer.

#### 4. Discussion

Our study explored the prevalence of HPV vaccine administration and associated predictors among 454 Lebanese female university students aged between 17 and 30 years. On the margin, we aimed to calculate knowledge scores of HPV and HPV vaccines to draw the appropriate assumptions.

According to our results, 18.9% of our target population have received at least one vaccination dose by the time of the survey whilst around 13.9% have received the full regimen of

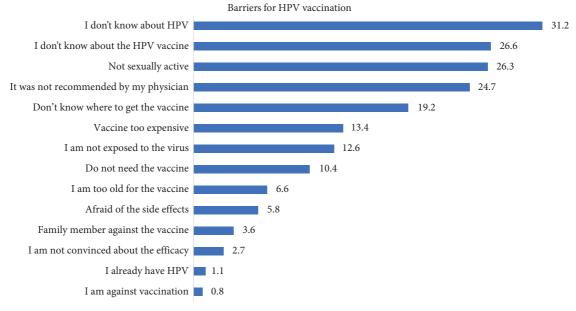


FIGURE 1: Barriers for HPV vaccination.

TABLE 2:	Description	of HPV	vaccination	variables.
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Variable		N (%)	Mean ± SD	Min	Max
Age of HPV vaccination uptake (1 <sup>st</sup> dose) (years)			$17.5 \pm 4.89$	9	30
Age of HPV vaccination uptake (1 <sup>st</sup> dose) in a participan	t with history of sexual activity	(years)	$19.6\pm5.15$	10	30
UDV marine untaka (at least 1 <sup>st</sup> deca)	Yes	86 (18.9)			
HPV vaccine uptake (at least 1 <sup>st</sup> dose)	No	368 (81.1)			
	Yes	82 (18.1)			
HPV vaccine uptake (2 <sup>nd</sup> dose)	No	4 (0.9)			
	Did not take the first dose	368 (81.1)			
	Yes	63 (13.9)			
HPV vaccine uptake (3 <sup>rd</sup> dose)	No	19 (4.2)			
<b>*</b> · · · ·	Did not take any doses	372 (81.9)			
	Yes	25 (16.1)			
HPV vaccination before first sexual intercourse	No	130 (83.9)			
	Yes	150 (33)			
Do you intend to get the HPV vaccine in the next year	No	218 (48)			
, , , , , , , , , , , , , , , , , , , ,	Already took a dose	86 (18.9)			
	Yes	318 (70)			
Have you ever heard about HPV	No	136 (30)			
	Yes	303 (66.7)			
Have you ever heard about HPV vaccine	No	151 (33.3)			

SD: standard deviation.

3 consecutive doses. We opted to consider that 18.9% is the prevalence of HPV vaccination among the surveyed, even though the vaccinated might not have reached optimal immunization status. That was decided given that no clear vaccination program was put in motion by the Lebanese state and that the number of doses needed to be considered fully vaccinated is undefined, varying between 2 and 3. A thorough literature search revealed concerning lack of papers on that matter. The only other significant paper of a study among Lebanese female students at the American University of Beirut found that 16.5% of their sample had received the HPV vaccine [7]. However, compared to a younger target population, such as Lebanese female school girls of private schools in Beirut, with an HPV vaccination prevalence of 2.5%, the age gap becomes more apparent. We could assume that, in general, Lebanese females tend to get vaccinated later in life [8]. In fact, we found that the mean vaccination age was 17.5 ( $\pm$ 4.89) with an age range between 9 and 30 years. Merely 15.3% of subjects received their first vaccination dose at age 12 or earlier, conforming to the guidelines stated by the Advisory Committee on Immunization Practices in the United States. Another published paper on HPV vaccination rates in neighboring countries of the Middle East (Jordan, Iraq, UAE, and Qatar) found that

Second introduct       Second introduct       66 (43)         If the hadrone       16 (53)       Nonperentifience       16 (53)         Can you transmit HV if you are symptomated       Vest of the above       24 (33)       16 (33)         Can you transmit HV if you are symptomated       None of the above       13 (53)       13 (53)         Can you transmit HV if you are symptomated       None of the above       13 (53)       13 (53)         Can smoking increase the risk of HPV?       Do not know       23 (53)       13 (53)         Can smoking increase the risk of HPV?       Do not know       23 (53)       13 (53)         HV can cause (multiple response accepted)       No       No       26 (54)       14 (01)         HV can cause (multiple response accepted)       Cervical causer       13 (51)       16 (53)       16 (53)         HV can cause (multiple response accepted)       Public causer       15 (12)       16 (13)       16 (13)         HV can cause (multiple response accepted)       Public causer       26 (54)       16 (13)       16 (13)         HV can cause (multiple response accepted)       Public causer       26 (54)       16 (13)       16 (13)         Condorn use will reduce the possibility of HV infection <t< th=""><th>Variable       Nariable       Nariable</th><th></th><th>(%) <i>u</i></th><th>Mean±SD</th><th>Min</th><th>Max</th></t<>	Variable       Nariable       Nariable		(%) <i>u</i>	Mean±SD	Min	Max
ed) Skin-to-skin contact Nonpenetrative sex During birth None of the above No No No No No No No No No No No No No		Sevual intercourse	400 (45 3)			
ed) None of the above None restan contact Nonperentitive sex Dome of the above No entending the above No enterding						
ed) houring birth None of the above None None None None None None None Non		MIII-10-SKIII COIILIACI	(0.21) 011			
During During     None of the above     None of the above     Yes     No     Do not know     No     Do not know     Anal cancer     Genital warts     Cervical cancer     Penile cancer     Prostate cancer     Prostate cancer     None of the above     None of the above     Yes     No     Do not know     No     None of the above     Yes     No     Do not know     No     No     No     No     No     Set the likelihood of HPV?     Do not know     HPV?     Do not know     Do not know	rir v transmission (mumpic responses accepted)	Nonpenetrative sex	100 (10.0)			
titć? No the more and the more not know No		Uuring birth None of the showe	(C.61) 0/1 34 (3.8)			
tic <sup>2</sup> Do not know No No Do not know Anal cancer Cervical cancer Cervical cancer Propharyngeal cancer Prostat cancer Postat cancer Do not know No No Do not know No No No No Do not know No No No No No No No No No No No No No						
tic? Do not know Yes Yes No Do not know Do not know Anal cancer Genital warts Cervical cancer Penile cancer Prostate cancer Prostate cancer Prostate cancer Do not know None of the above Yes No Do not know No Do not know No Do not know No Ection Do not know No Do not know Do not know No Do not know No Do not know Do not know No Do not know Do NO		Yes	265 (58.4)			
Do not know     Yes     No     Do not know     Do not know     Anal cancer     Genial warts     Cervical cancer     Penile cancer     Pointe know     No     Pointe know     Pointe know     No     Pointe know	Can you transmit HPV if you are asymptomatic?	No	13 (2.9)			
Yes     No       Do not know     Anal cancer       Anal cancer     Cervical warts       Cervical warts     Cervical warts       Cervical cancer     Do not know       Point cancer     Do not know       None of the above     Yes       None of the above     Yes       Vinfection     Do not know       Vinfection     Do not know       Yes     No       No     No       No     No       No     Do not know       Yes     No       Infection     Do not know       Section     No       No     No       Set the likelihood of HPV?     Do not know       HPV?     Do not know       Set the likelihood of HPV?     Do not know       HPV?     Do not know		Do not know	176 (38.8)			
No   No     Do not know   Anal cancer     Anal cancer   Genital warts     Cervical cancer   Cervical cancer     Prostate cancer   Prostate cancer     Prostate cancer   Do not know     None of the above   Yes     None of the above   Yes     V infection   Do not know     V infection   Do not know     Yes   Yes     infection?   No     bo not know   Yes     infection?   Do not know     Set the likelihood of HPV?   Do not know     HPV?   Do not know		Yes	134 (29.5)			
Do not know     Anal cancer     Anal cancer     Genital warts     Cervical cancer     Penile cancer     Prostate cancer     Point know     None of the above     Y infection     V infection     V infection     Do not know     None of the above     Yes     None of the above     Yes     No     No     Do not know     No	Can smoking increase the risk of HPV?	No	33 (7.3)			
Anal cancer     Genital warts     Genital warts     Gervical cancer     Prostate cancer     Proste     Prostate     Proter		Do not know	287 (63.2)			
Genital warts Cervical cancer Prostate cancer Prostate cancer Prostate cancer Prostate cancer Prostate cancer No       Ves     Ves       Value     Ves       Void     No       Void     No       Void     No       Value     Ves       Void     No       Void     No       Void     No       Infection?     No       Infection?     No       Infection?     No       Reside     No       Infection?		Anal cancer	136 (12.6)			
Cervical cancer     Cervical cancer       Prostate cancer     Prostate cancer       No     No       No     No       Pronot know     Yes       Infection?     Do not know       Infection?     Do not know       See the likelihood of HPV?     No       HPV?     Do not know		Genital warts	174 (16.1)			
Oropharyngeal cancer     Penile cancer     Do not know     None of the above     Yes     No     Do not know     Yes     Root     No     No     Itansmission?     Do not know     Yes     Infection     Do not know     No     No     No     No     No     See the likelihood of HPV?     Do not know     HPV?     Do not know     Yes     No     No     No     No     No     No     No     No     No     HPV?     Do not know     No     No <		Cervical cancer	286 (26.4)			
Penile cancer     Prostate cancer     Do not know     None of the above     Yes     No     Do not know     Vinfection     Do not know     Yes     No     Do not know     Yes     No     Do not know     Yes     Itansmission?     transmission?     Do not know     Yes     transmission?     Do not know     No     Se the likelihood of HPV?     Do not know     HPV?     Do not know     Se the likelihood of HPV?     Do not know		Oropharyngeal cancer	125 (11.5)			
Prostate cancer     Do not know       Do not know     Yes       No     No       No     Do not know       Yes     No       No     No       On ot know     Yes       No     Do not know       Yes     No       Of HPV?     Do not know       Yes     No       Do not know     Yes       No     Do not know       Yes     No       Do not know     No       Do not know     No       Do not know     No       No     Do not know	HPV can cause (munple responses accepted)	Penile cancer	136 (12.6)			
Do not know None of the above Yes No Do not know Yes No Do not know No Do not know Yes No Do not know No Do not know Do not know		Prostate cancer	70 (6.5)			
None of the above Yes No Do not know Yes No Do not know No Do not know		Do not know	156 (14.4)			
Yes     No     Do not know		None of the above	0 (0)			
No Do not know Yes No Do not know Yes No Do not know Yes No Do not know Yes No Do not know Yes No Do not know Yes No Do not know Do not know Do not know Do not know Do not know		Yes	161 (35.5)			
Do not know Yes No Do not know Yes No Do not know Yes No Do not know Yes No Do not know Yes No Do not know Yes No Do not know Do not know Do not know Do not know Do not know	Can HPV lead to infertility?	No	37 (8.1)			
Yes     No     Do not know     Yes     No     No     No     Do not know     Yes     No     Of HPV?     Do not know     Yes     No     Do not know     Yes     No     Do not know     Yes     No     Do not know     No     Do not know		Do not know	256 (56.4)			
No Do not know Yes No Do not know Yes No Do not know Yes No Do not know Yes No Do not know No Do not know Do not know No Do not know		Yes	363 (78.9)			
Do not know Yes No Do not know Yes No Do not know Yes No Do not know Yes No Do not know Yes No Do not know Do not know Do not know	Condom use will reduce the possibility of HPV infection	No	90 (19.8)			
Yes No Do not know Yes No Do not know Yes No Do not know Yes No No Do not know No Do not know		Do not know	1(0.2)			
No     Do not know     Yes     No     Yes     No     Yes     No     Yes     No     No     No     No     No     No     No     Do not know     No     Do not know		Yes	50 (11 0)			
Do not know Yes No Do not know Yes No Do not know Yes No Do not know Yes No Do not know	Condom use will completely prevent HPV infection	No	403 (88.8)			
Yes     No     Do not know     Yes     No     Do not know     Yes     No     Of HPV?     Do not know     Yes     No     Do not know     Yes     No     Do not know	monorma a ser monord (mondarios and som monormo)	Do not know	1 (0.2)			
of HPV? Do not know <b>Yes</b> No Do not know <b>Yes</b> No Do not know <b>Yes</b> No Do not know Do not know		Yes	53 (11.7)			
Do not know Yes No Do not know Yes No Do not know No Do not know	Can circumcision prevent HPV infection and transmission?	No	156(34.4)			
Yes No Do not know Yes No No Do not know No Do not know		Do not know	245 (54.0)			
No Do not know <b>Yes</b> No No Do not know No Do not know		Yes	366 (80.6)			
Do not know <b>Yes</b> No Do not know <b>Yes</b> No Do not know	Multiple sex partners will increase the risk of infection?	No	2 (0.4)			
of HPV? of HPV? Do not know Yes No Do not know		Do not know	86 (18.9)			
of HPV? No Do not know Yes No Do not know		Yes	162 (35.7)			
Do not know Yes No Do not know		No	81 (17.8)			
Yes No Do not know		Do not know	211 (46.5)			
No Do not know		Yes	118 (26)			
Do not know	Will washing intimal area reduce the risk of HPV?	No	147 (32.4)			
	5	Do not know	189 (41.6)			

6

	TABLE 3: Continued.				
Variable		(%) <i>u</i>	Mean±SD	Min	Max
Is it possible to have asymptomatic HPV infection for many years?	Yes No Do not know	266 (58.6) 9 (2) 179 (39.4)			
Is there any cure for HPV infection?	Y <sub>es</sub> No Do not know	111 (24.4) 102 (22.5) 241 (53.1)			
Can HPV be tested in men?	Y <sub>es</sub> No Do not know	$\begin{array}{c} 232 \ (51.1) \\ 27 \ (5.9) \\ 195 \ (43.0) \end{array}$			
What are the possible screening tests?	<b>PAP smear and HPV testing</b> PAP smear alone HPV testing alone	296 (65.2) 111 (24.4) 47 (10.4)			
When should HPV screening begin?	<b>Between 21 and 29</b> Younger than 21 Older than 29	257 (56.6) 81 (17.8) 116 (25.6)			
SD: standard deviation. The correct answers are in bold.					

Variable		N (%)	Mean ± SD	Min	Max
	Yes	188 (41.4)			
The vaccine is divided into 3 doses over 6 months	No	23 (5.1)			
	Do not know	243 (53.5)			
	Yes	154 (33.9)			
HPV vaccine can be given from the age of 9	No	76 (16.7)			
	Do not know	224 (49.3)			
	Yes	123 (27.1)			
HPV vaccine can prevent oropharyngeal cancer	No	55 (12.1)			
	Do not know	276 (60.8)			
	Yes	154 (57.9)			
HPV vaccine can prevent penile cancer	No	37 (8.1)			
	Do not know	263 (57.9)			
	Yes	90 (19.8)			
HPV vaccine can prevent prostate cancer	No	75 (16.5)			
1 1	Do not know	289 (63.7)			
	Yes	151 (33.3)			
HPV vaccine can prevent anal cancer	No	38 (8.4)			
1	Do not know	265 (58.4)			
	Yes	206 (45.4)			
Is the vaccine more effective if given before first sexual intercourse?	No	58 (12.8)			
0	Do not know	190 (41.9)			
	Yes	58 (12.8)			
Persons who received all doses are fully immunized against cervical cancer?	No	187 (41.2)			
,	Do not know	209 (46.0)			
	Yes	124 (27.3)			
Persons who received all doses are fully immunized against HPV?	No	127 (28)			
, 0	Do not know	203 (44.7)			
	Yes	26 (5.7)			
Is it safe to have unprotected sex after HPV vaccination?	No	306 (67.4)			
1	Do not know	122 (26.9)			
	Yes	143 (31.5)			
Are HPV vaccine side effects limited to mild local reaction?	No	41 (9)			
	Do not know	270 (59.5)			
	Yes	41 (9)			
Can HPV cause permanent health problems?	No	195 (43)			
1 1	Do not know	218 (48)			

TABLE 4: Responses to questions about HPV vaccine knowledge by the study participants.

SD: standard deviation. The correct answers are in bold.

only 3% of their female citizens had received all vaccine doses and achieved full immunization. The authors explained this low rate by emphasizing the lack of knowledge of cervical cancer and HPV vaccination by their surveyed subjects. When the unvaccinated were asked about their will to take the HPV vaccine in the upcoming year, only 33% showed willingness to be vaccinated. This result is lower compared to neighboring countries where 43.2% were willing to receive the vaccine [11]. In contrast to a developed and liberal nation, such as the United States of America, the Center for Diseases Control and Prevention (CDC) declared that, in 2020, 77.1% of adolescents aged between 13 and 17 were vaccinated against HPV with at least one dose of whom 61.4% were fully vaccinated [12]. The difference in the vaccination rate between the United States of America and our population may be attributable to the difference in economic status, HPV and HPV vaccine knowledge, and awareness campaigns.

The ideal time for HPV vaccination should be before an individual's first sexual relationship [13]. In our study, among all participants with a positive history of sexual activity, we found that 5.5% had already taken the HPV vaccination before their first sexual intercourse and that the mean age of vaccination was 19.6 years old for taking at least one HPV vaccine dose. This constitutes a public health issue since, as stated earlier, HPV is associated with multiple malignant diseases, most notably cervical cancer among others [14]. To concur, Lei et al. calculated an 88% decreased risk of cervical cancer in participants who initiated HPV vaccination before the age of 17 compared to those who had never been vaccinated [15]. Concerning vaccination barriers, we have found that 24.7% have stated that they did not get vaccinated since the idea was never recommended by their primary physician in the first place. Being rarely covered by Lebanese health care providers, 13.4% did not take the vaccine because of its' expensive price on the market. Abi

Variable		F	Results	6 mal
Variable		Vaccinated	Non-vaccinated	p value
Sociodemographic characteristics				
Age*		23.48	22.5	0.016t
M.:*	Medical specialty	51 (23.3)	168 (76.7)	0.022
Major*	Non-medical specialty	35 (14.9)	200 (85.1)	0.023c
TT:*	Public	10 (5.6)	169 (94.4)	0.00-
University*	Private	76 (27.6)	199 (72.4)	0.00c
Nationality	Lebanese	84 (19.4)	348 (80.6)	0.227.
Nationality	Other <sup>1</sup>	2 (9.1)	20 (90.9)	0.227c
	Married	46 (16.5)	232 (83.5)	
Family status	Single/divorced	32 (23.4)	105 (76.6)	0.242c
	In a relationship/engaged	8 (20.5)	31 (79.5)	
	Beirut	17 (17.9)	78 (82.1)	
Residence	Mount-Lebanon	58 (20.9)	219 (79.1)	0.298c
	Other <sup>2</sup>	11 (13.4)	71 (86.6)	
Religion*	Christian	64 (22.3)	223 (77.7)	
	Muslim/druze	18 (12.1)	131 (87.9)	0.033c
	Non-religious	4 (22.2)	14 (77.8)	
N.C. 1 1 (* 11 1*	Less than university degree	26 (12.8)	177 (87.2)	0.000
Maternal educational level*	University degree or higher	60 (23.9)	191 (76.1)	0.002c
ר <u>ו ו ו ו ו</u> א	Less than university degree	30 (11.3)	236 (88.7)	0.000
Paternal educational level*	University degree or higher	56 (29.8)	132 (70.2)	0.000c
т 1···1 1 .11 · · *	Less than 5.000.000LBP	67 (17)	328 (83)	0.040
Individual monthly income*	More than 5.000.000LBP	19 (32.2)	40 (67.8)	0.043c
	Medical history			
Company history	No	86 (19.2)	362 (80.8)	0.000.
Cancer history	Yes <sup>3</sup>	0	6 (100)	0.233c
	Yes	2 (28.6)	5 (71.4)	0.510
Family history of cancer related to HPV <sup>4</sup>	No history	84 (18.8)	363 (81.2)	0.512c
	At least once	4 (15.4)	22 (84.6)	
History of STD	Never	82 (19.2)	346 (80.8)	0.633c
	Habits	()		
	No	19 (11.4)	148 (88.6)	
Alcohol drinking*	Frequently	23 (26.1)	65 (73.9)	0.005c
0	Occasionally	44 (22.1)	155 (77.9)	
	Yes	13 (13.5)	83 (86.5)	
Smoking	No	73 (20.4)	285 (79.6)	0.128c

TABLE 5: Associations between HPV vaccine uptake and sociodemographic, medical histo	ory and habits.
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c: Chi-square test; t: student's *T*-test; \*: variables with significant difference; LBP: Lebanese pounds; STD: sexually transmitted diseases. <sup>1</sup>: Syrian, Palestinian, Iraqi, Filipino, Brazilian. <sup>2</sup>: Aakar, North, Baalbeck-Hermel, Beqaa, South, Nabatiyeh. <sup>3</sup>: breast cancer, CML, non-Hodgkin lymphoma, Hodgkin lymphoma (x2), kidney cancer. <sup>4</sup>: cervical cancer, penile cancer, anal cancer, oropharyngeal cancer.

Jaoude et al. stated that 64% of the 228 primary physicians surveyed in the Greater Beirut area considered the price to be the main barrier against HPV vaccination. In addition, the authors concluded that a refund of vaccine costs by healthcare providers can significantly improve vaccination rates [5]. Drawing conclusions from the Australian experience, HPV vaccination rates increased to an astonishing 70% for girls aged between 12 and 13 after reforming their vaccination program and administering all vaccines free of charge [16].

In an effort to better understand the Lebanese vaccination rates, two knowledge scores were computed, an HPV and its vaccination score. We have found that our sample has a mean HPV knowledge score of 13.9 over 28 points (49.64%). Although some of the questions asked may differ, this score can be compared to the 2015 study by Dany et al. that found a mean knowledge score of 52.7%, where scores were calculated for 512 female students attending one of Beirut's universities. These scores reflect female university students' poor-to-moderate knowledge of HPV [7]. In detail, concerning HPV transmission modes, only 12.8% of our responders chose skin-to-skin transmission; however, it is the main transmission route. The majority of our participants knew that condom use during sexual intercourse will only decrease the risk of HPV transmission and will not prevent the risk completely. These results are compatible with the information found in the literature [17]. In addition, only 29.5% knew that smoking can increase the risk of HPV transmission. In their study, Schabath et al. proved that smoking will increase the risk of HPV transmission with an

Variable Age on first sexual intercourse		R	<u> </u>		
		Vaccinated	Nonvaccinated	<i>p</i> value	
		21.7	20.8	0.247t	
Sexual activity	Active	35 (22.6)	120 (77.4)	0.154c	
Sexual activity	Not active	51 (17.1)	248 (82.9)	0.154c	
	Never	13 (28.3)	33 (71.7)		
Condom use in relationship	Always	8 (21.1)	30 (78.9)	0.1946	
	Frequently	9 (25.7)	26 (74.3)	0.184c	
	Not currently sexually active	56 (16.7)	279 (83.3)		
	One	34 (24.3)	106 (75.7)		
Number of partners usually	Two and more	1 (10)	14 (93.3)	0.092c	
	Not sexually active	51 (17.1)	248 (82.9)		
	Man	32 (22.1)	113 (77.9)		
Doute on(a) can dou	Woman	2 (33.3)	3 (75)	0.472c	
Partner(s) gender	Both	1 (25)	4 (66.7)	0.4720	
	Not sexually active	51 (17.1)	248 (82.9)		
	Anal giving or receptor	2 (28.6)	5 (71.4)		
Sorreal professor of	Oral giving or receptor	16 (21.9)	57 (78.1)	0.426 -	
Sexual preference	Vaginal sex	32 (22.9)	108 (77.1)	0.426c	
	Not active	51 (17.1)	248 (82.9)		

TABLE 6:	Bivariate	analysis	of sexual	behavior	questions.
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c: chi-square test; t: Student's T-test.

#### TABLE 7: Bivariate analysis of vaccine uptake and knowledge scores.

Variable	F	t volue		
variable	Vaccinated	Nonvaccinated	<i>p</i> value	
HPV knowledge score*	16.7	13.1	<0.01t	
HPV vaccine knowledge score*	7.43	3.25	<0.01t	

\*: variable with a significant difference; t: Student's T-test.

#### TABLE 8: Factors associated with HPV vaccine uptake (at least one dose) among study participants.

		Adjusted 95% CI of ORa		⊅ value	
		odds ratio (ORa)	Lower	Upper	<i>p</i> value
University	Public (ref) Private	0.228	0.108	0.479	< 0.001
Paternal educational level	University degree or higher (ref) Less than a university degree	1.881	1.073	3.297	0.027
HPV vaccine knowledge score	7 0	1.378	1.260	1.508	< 0.001

CI, confidence interval; OR, odds ratio binary logistic regression was used (n = 454).

OR of 1.19. They also found that this association is only valid for carcinogenic HPV [18]. Concerning the rate of HPV transmission in circumcised patients, 11.7% knew that circumcision can lower HPV transmission rates. In a trial by Yuan et al., comparing HPV transmission between uncircumcised and circumcised participants, a 35% reduction in the risk of HPV transmission was noticed in the latter category. In addition, HPV is known to lead to infertility. In our study, 35.5% answered positively to this question. HPV infections were found to be potential risk factors for infertility. HPV is to be screened and treated in couples with fertility issues [19] As per the CDC, there is currently no approved test for HPV testing in men contrary to females where HPV can be tested, and a PAP smear can lead to diagnosing cervical cancer [20]. Regarding HPV vaccination knowledge scores, our sample scored an average of 4.4 over 12 (36.6%) allowing us to draw the same conclusion as Dany et al. A severe lack of knowledge about HPV and its vaccine exists among the surveyed, and this issue should be addressed [7].

It is known that HPV vaccination can be given to the patients 9 years of age and older. Less than half of our participants knew that information. In addition, concerning the prevention of HPV-related cancers, the results were underwhelming. The results of the HPV vaccine knowledge score are even lower than the HPV knowledge score and this is related to a lack of education on sexually transmitted diseases in general and HPV in particular.

In our final analysis, the logistic multivariate analysis, we found that being in a private university was a predictor of

being vaccinated. This result, as we have already mentioned, is confirmed by the literature and may be related to the financial status of the participants. However, no previously published studies have found the university type to be a predictor of HPV vaccination. In addition, paternal educational level was also found to be a predictor of HPV vaccination. In detail, women with a paternal university degree or higher had an increased probability of being vaccinated. A recent study conducted in Ethiopia has found similar results concerning HPV vaccination and parental educational level [21]. However, we have found a contradictory result in the literature, where Ganczak et al. mentioned in their study conducted in Poland that there is no significant variance in the HPV knowledge between parents from different educational levels [22]. Another contradictory result was noticed in Feiring et al.'s paper published in 2015. In their study, they found that a higher paternal educational level was associated with a decrease in the probability of receiving HPV vaccines. They have also found a significant association between maternal educational level and HPV vaccination likelihood. Additionally, a study conducted in Texas in 2013 has shown that vaccine refusal was more frequent among parents who are more educated [23]. These findings may be explained by the fear of vaccination side effects. Also, the HPV vaccine knowledge score was found to be a predictor of girls' HPV vaccination. These results are in agreement with a published study from Nigeria where an OR of 7.98 was found concerning the rate of HPV vaccination [24]. In our study, being in a medical or nonmedical major was not found to be a predictor of HPV vaccination status contrary to the results found by Dany et al. [7]. Also, HPV vaccination status increased with increasing age [25]. We did not find age to be a significant predictor of HPV vaccination in our study.

Our study allows the collection of large amounts of data via a single survey. This type of survey is cost-effective and time-effective and of high-yield. Concerning weaknesses, our study has some limitations. In our questionnaire, there were little to no open-ended questions and it did not deepen our understanding of the situation, except through prewritten questions. It is considered inflexible. Furthermore, the majority of the questions in our questionnaire targeted a taboo subject. This may induce a response bias, or it may lead to a decrease in the response rate due to the discontinuation of the questionnaire. Also, the response rate cannot be calculated due to the impossibility of knowing the number of people reached. Finally, a selection bias might be present due to the use of a "snowball" method of sampling. We were limited by the online distribution of the questionnaire due to the COVID-19 pandemic. This step may have limited face-to-face questionnaire filling and the ability to clarify litigious questions.

### 5. Conclusion

In summary, HPV and its related diseases, most notably cervical cancer, are an important public health issue that is often overlooked. Passing on to the literature review, we have found an alarming lack of scientific papers discussing

HPV and its vaccine in the Lebanese population. Being part of the HPV vaccination primary target population, we surveyed 454 women enrolled in multiple Lebanese universities and aged between 17 and 30 years. To better understand the burden of HPV on Lebanese society, HPV vaccination prevalence was calculated (18.9%) and was considered low. This prevalence can be partly explained by the computed average low-to-moderate HPV vaccination knowledge score of our sample. In addition, and through our multivariate analysis, we found that having a father with a university degree and being enrolled in a private institution both increase the subject's odds of being vaccinated. We could argue that both of these predictors are associated with better economic status, given that the vaccine is expensive and usually not reimbursed by healthcare providers. To date, no national strategy for HPV vaccination has been put in place. If we were to propose a solution to decrease HPVrelated morbidity and mortality in the Lebanese population, we would suggest state-sponsored vaccination awareness campaigns to increase knowledge scores. Making the vaccine cheap or even free and available in the Lebanese market will undoubtedly boost the vaccination prevalence and is crucial to the well-being of our society.

#### Abbreviations

CDC:	Centers for Disease Control and Prevention
COVID-19:	Coronavirus disease 2019
CI:	Confidence interval
HPV:	Human papillomavirus
INSPECT:	Institut National de Santé Publique,
	D'Epidemiologie Clinique et de Toxicologie
ORa:	Adjusted odds ratio
PAP:	Papanicolaou test
SD:	Standard deviation
URL:	Uniform resource locators.

#### **Data Availability**

The datasets used and/or analyzed during the current study are available from the corresponding author on reasonable request.

#### **Ethical Approval**

The study was reviewed and approved by the ethical review board of INSPECT (IRB Approval No. 2022REC-005-IN-SPECT-01-21).

#### Disclosure

This article was submitted as a preprint [26], available at https://www.researchsquare.com/article/rs-2057410/v1. Joey El Khoury and Rami Halabi are first co-authors.

#### **Conflicts of Interest**

The authors declare no conflicts of interest.

#### **Authors' Contributions**

J.K. and R.H. were involved in sampling and data collection and writing the manuscript. W.K. conceptualized the study and contributed to the literature review. M.H., R.K, and N.S contributed by monitoring the data collection and reviewing the manuscript. All authors read and approved the final manuscript. Joey El Khoury and Rami Halabi have contributed equally to this work.

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