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RESEARCH ARTICLE

Reasons for non-attendance to cervical cancer screening and acceptability of HPV self-sampling among Bruneian women: A cross-sectional study

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Abstract

Objective

Uptake for cervical cancer screening remains well below the 80% target as recommended by Brunei's National Cervical Cancer Prevention and Control plan. We conducted a pilot study to determine the reasons for non-attendance and explore their acceptance of human papillomavirus (HPV) self-sampling as an alternative to the Pap test.

Methods

A cross-sectional study was conducted at a primary healthcare center in Brunei, from January to December 2019. We recruited screening non-attendees, defined as women who were eligible for Pap test but who either never, or did not have one within the recommended screening interval of 3 years. This recruitment was done conveniently among women attending outpatient care and/or child health services at the primary healthcare center. Participants were first asked to complete a self-administered paper-based questionnaire on their reasons for screening non-attendance, and then invited for HPV self-sampling. Among those who agreed to participate in HPV self-sampling, they were asked to complete a second questionnaire on the self-sampling procedure and their samples were tested for highrisk HPV (hr-HPV). Results were analyzed using descriptive and inferential statistics.

Result

We enrolled 174 screening non-attendees, out of which 97 (55.7%) also participated in HPV self-sampling. The main reasons for not attending Pap test screening were fear of bad results (16.1%, n = 28); embarrassment (14.9%, n = 26) and lack of time due to home commitments (10.3%, n = 18). When compared to those who agreed to participate in HPV self-sampling, those who declined were significantly older (p = 0.002) and less likely to agree that they are susceptible to cervical cancer (p = 0.023). They preferred to receive Pap test-

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related information from healthcare workers (59.0%, n = 155), social messaging platforms (28.7%, n = 51) and social media (26.4%, n = 47). HPV self-sampling kits were positively received among the 97 participants, where > 90% agreed on its ease and convenience. Nine (9.3%) tested positive for hr-HPV, out of which eight were non-16/18 HPV genotypes.

Conclusion

Our findings suggest that promoting awareness on cervical cancer, clarifying any misconceptions of Pap test results, and highlighting that the disease is preventable and that early detection through screening can facilitate successful treatment would help increase screening uptake among Bruneian non-attendees. Response to HPV self-sampling was highly positive, suggesting the possibility of implementing this strategy in the local setting. Our high detection of non-16/18 HPV genotypes suggest high prevalence of other hr-HPV genotypes in Brunei. Larger studies should be conducted to further validate our findings.

Introduction

Cervical cancer is highly preventable but still remains one of the most common cancers among women worldwide. Globally, an estimated 604,000 women were diagnosed with cervical cancer, and 342,000 women died from the disease in 2020 [1]. Cervical cancer screening has drastically reduced the incidence of invasive cervical cancer in countries that have implemented such screening programs [2], which traditionally involves the use of the Papanicolaou (Pap) test.

Brunei Darussalam (population 459,500) is a small Southeast Asian country with a predominant Muslim population and a crude birth rate of 15.3 per 1,000 population [3]. Within this region, it has one of the highest age-standardized incidence rate (ASR) for cervical cancer: 20.6 per 100,000 women-years in Brunei when compared to 10.5 and 7.7 per 100,000 women-years in Malaysia and Singapore, respectively [4]. Brunei has initiated an organized cervical screening program since 2009, where married or ever married Bruneian women between 20 and 65 years old were invited to attend cervical cancer screening through periodic mail invitations. Pap test is the only screening test used in the country, and currently, liquid-based cytology is being used since 2012. Women with any positive cytology result, defined as with atypical squamous cells of undetermined significance (ASC-US) or worse, are followed up with colposcopyguided cervical biopsy to diagnose cervical intra-epithelial neoplasia (CIN). The latter refers to premalignant lesions that are mainly caused by infection with certain types of human papillomavirus (HPV) [5], and can be categorized into any one of three stages (CIN1, CIN2, or CIN3) depending on the degree of dysplasia. If untreated, either CIN2 or CIN3 (collectively referred to as CIN2+) can progress to cervical cancer.

Despite this screening service being offered free of charge, the national screening coverage rate remains low at 44% in 2018 (unpublished data). Reasons for screening non-attendance can vary across settings [6, 7], but they can be broadly categorized into two groups: practical and organizational barriers (such as forgot to schedule an appointment, work and childcare commitments) [6], and emotional barriers (such as feeling healthy, lack of time, discomfort associated with gynecologic examination embarrassment, fear of smear test, previous negative experiences and dissatisfaction with their general practitioner) [6, 8–10]. As women who do not attend screening are at increased risk of developing cervical cancer [6], it is thus important to first understand why women chose not to attend screening in the local context.

HPV DNA detection has been recommended by the World Health Organization as the primary screening test for cervical cancer as specific high-risk HPV subtypes (hr-HPV) are known to be a causative agent [1]. In particular, the use of HPV self-sampling kits was suggested to increase screening uptake particularly among screening non-attendees [11, 12], due to its ease of access (where kits could be mailed to women's homes) and also flexibility for women to perform the test by themselves. Previous studies have shown HPV self-sampling to be highly acceptable among screening non-attendees [13, 14]. In addition, HPV self-sampling results exhibit similar sensitivity and specificity compared to those from samples taken by trained professionals [15]. Repeated HPV self-sampling and testing were shown to increase screening uptake [11, 12, 16], and also resulted in at least two-fold higher detection rate of CIN2+ when compared to the Pap test [17, 18]. In the United Kingdom and Australia, early detection through an organized screening program using HPV testing as the primary screening test was shown to reduce cervical cancer morbidity and mortality [19, 20].

With Brunei's relatively high incidence of cervical cancer and low screening uptake, we conducted a pilot study to explore the reasons behind non-attendance and to assess the acceptability of HPV self-sampling as a possible alternative to the Pap test among non-attendees (specifically women who are currently not accessing or attending screening). Study findings could be used to strategize ways to improve screening uptake and provide preliminary evidence towards implementing HPV testing as the primary screening test for cervical cancer in Brunei.

Methods

Study design and data collection

A cross-sectional survey was conducted at the Jubli Perak Sengkurong Health Center (JPSHC), from January to December 2019. JPSHC is a government-funded primary healthcare center located at Brunei-Muara District, where the majority (69.3%) of the country's population resides. This center provides primary health care services to Mukim Sengkurong, a sub-district with about 32,000 people from various socioeconomic backgrounds.

Eligible women attending either the outpatient or child health clinic at JPHSC were conveniently recruited by triage nurses. We defined screening non-attendees as married or ever married women between 20 and 65 years old and have never undergone cervical cancer screening, or did not have one within the recommended screening interval of 3 years. We excluded women who could not comprehend Brunei-Malay or English language, were pregnant, have had total hysterectomy, or with a history of malignancies.

We implemented a two-stage recruitment procedure. In the first stage, participants were first recruited to complete a self-administered questionnaire on the reasons for screening non-attendance (Q1) onsite. Q1 consists of 19 questions on the participant's socio-demographics, reasons for not getting Pap test, attitude and knowledge on cervical cancer, and lastly, preferred sources to acquire information about Pap test. From a prepared list of 16 possible reasons for not attending the screening program, participants were also asked to select one "Major" reason (defined as the main reason) and one or more "Minor" reason (defined as other reasons for not attending the screening program). Responses for questions on their attitude and knowledge on cervical cancer were recorded using a five-point Likert scale, ranging from "Strongly Agree" to "Strongly Disagree".

In the second stage, all participants were given an envelope containing information on HPV self-sampling and an instruction leaflet on the procedure, after completing Q1. Within the following two weeks, they were contacted via telephone by a trained nurse, and those who gave verbal consent were given an appointment at JPSHC. On the day, the nurse first explained the procedure using an instructional video and answered any questions. Participants were

then given a self-sampling kit and asked to perform the procedure in the clinic. After completion, they were asked to complete the second self-administered questionnaire (Q2) on their acceptability of the self-sampling procedure. Q2 consists of 12 five-point Likert scale questions (ranging from "Strongly Agree" to "Strongly Disagree") on their experiences and opinions of the self-sampling kit.

Two separate written consents were requested from participants: one for completing Q1 and another for performing self-sampling and completing Q2.

Questionnaires used

The two questionnaires used (Q1 and Q2) were bilingual self-administered paper-based questionnaires in the two languages commonly used in Brunei (namely, Brunei-Malay and English language). Both questionnaires were adapted from similar studies [21, 22]. They were first translated to Brunei-Malay language by native speakers, and then back-translated to check for inconsistencies in comprehension. Both questionnaires were also pre-tested on six eligible women to assess if the questions could be easily understood. Responses from pre-testing were not included in the analysis.

Self-sample handling and laboratory testing

Swabs collected from the self-sampling kits were sent to an overseas laboratory (at BNH hospital, Thailand) for hr-HPV testing. The self-sampling device used was the Evalyn® brush from Rovers Medical Devices. Upon receipt at the testing laboratory, the dry Evalyn® brush was suspended in SurePath medium from which a sample was obtained for the identification of the presence of hr-HPV using the cobas HPV test (Roche, USA). Cobas HPV tests are automated qualitative *in vitro* tests for the detection of HPV DNA in patient specimens. The tests utilize amplification of target DNA by polymerase chain reaction (PCR) and nucleic acid hybridization for the detection of 14 hr-HPV types (namely genotypes 16, 18, and 12 pooled hr-HPV genotypes 31, 33, 35, 39, 45, 51, 52, 56, 58, 59, 66 and 68) in a single analysis. Results obtained from this test can be categorized into four groups: Negative, HPV-16 positive, HPV-18 positive, and positive for non-16/18 HPV genotypes.

Clinical management of hr-HPV positive participants

We adopted the cytology triage strategy for HPV self-sampling participants [23]. Participants with hr-HPV positive results were invited for an immediate clinic-based cytology triage test. Those found to have negative cytology results were invited for a repeat cytology triage test after six months. Those with second negative results were returned to routine cervical screening recall. Participants with any positive cytology result (defined as with ASC-US or worse) were referred for colposcopic examination.

Statistical analysis

Descriptive statistics was conducted to characterize the socio-demographic characteristics of the study population, their reasons for not attending screening, their attitudes and preference for information access on such screening, as well as responses from Q2. Where appropriate, Mann-Whitney and/or Fisher's exact tests were used to assess significant differences in socio-demographic characteristics, reasons for not attending screening, and attitudes between women who agreed and women declined to join HPV self-sampling. Count responses for questions with the five-point Likert scale were categorized into three categories (agree, neutral, and disagree), and those with missing values were classified as neutral. Statistical analysis was

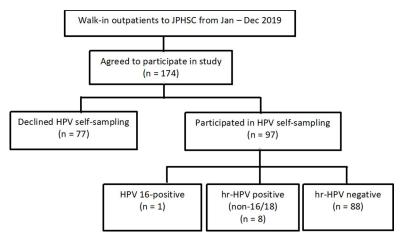


Fig 1. Flowchart of study participants at JPHSC, Brunei (Jan-Dec 2019).

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conducted using R ver. 3.6 [24]. Ethics approval was obtained from the Medical and Health Research and Ethics Committee (MHREC), Ministry of Health, Brunei Darussalam [Reference no. MHREC/MOH/2018/9(2)].

Results

A total of 174 eligible women were enrolled in this study from January to December 2019, out of which 97 (55.7%) also participated in HPV self-sampling (Fig 1). Their median age was 45 years, ranging between 23 and 65 years (Table 1). The participants were mainly of Malay ethnicity (92.5%, n = 161), married (90.2%, n = 157), and had \geq 3 births (60.3%, n = 105). About half of the participants had their last Pap test performed between 4 and 10 years ago (52.3%, n = 91), and have never received the HPV vaccine (54.6%, n = 95). There were significant differences between those who agreed and declined to participate in HPV self-sampling: those who declined were significantly older (p = 0.002) and more likely to have their last Pap test performed \geq 10 years ago (p = 0.031).

Fig 2 and S1 Table show the responses for the major and minor reasons for not attending cervical cancer screening. The top three major reasons reported were that they were "afraid of getting a bad result" (16.1%, n = 28), "feeling embarrassed being examined by a doctor or nurse" (14.9%, n = 26), and "I can't find the time as I'm too busy at home" (10.3%, n = 18). The top three minor reasons were "feeling embarrassed being examined by a doctor or nurse" (20.7%, n = 36), "I can't find time as I'm too busy at work" (20.7%, n = 36), and "afraid of getting a bad result" (20.1%, n = 35).

When comparing top 10 major reasons for not attending screening between those who agreed and declined HPV self-sampling (Table 2), those whose major reason was "feeling embarrassed being examined by a doctor or nurse" were significantly more likely to join self-sampling (p = 0.020). Also, those whose major reason was "afraid of getting a bad result" were significantly more likely to decline self-sampling (p = 0.034). Among those who were employed (n = 98), about a quarter (23.5%, n = 23) reported work-related reasons as their major reason for not attending screening ("I can't find the time as I'm too busy at work" and "Difficult to get permission from employer").

While the responses vary when asked about their health and susceptibility to disease, most agreed on the benefits of undergoing Pap test (92.0%, n = 160), and that cervical cancer is a severe and potentially lethal disease (82.8%, n = 144; Table 3 and S2 Table). Those who agreed

Table 1. Sociodemographic characteristics of the study population, including comparison between groups that agreed and declined to join HPV self-sampling.

Characteristics		Total study population (n = 174)	Joined self-sampling (n = 97)	Declined self-sampling (n = 77)	p-value 0.002*	
Median age in years (IQR)		n (%)	n (%)	n (%)		
		45.0 (15.25)	41.0 (17)	49.0 (14.5)		
Age group (in years)	20-24	3 (1.7)	3 (100)	0 (0.0)	0.018*	
	25–29	20 (11.5)	15 (75.0)	5 (25.0)		
	30-34	14 (8.1)	10 (71.4)	4 (28.6)		
	35–39	28 (16.1)	18 (64.3)	10 (35.7)		
	40-44	18 (10.3)	11 (61.1)	7 (38.9)		
	45-49	27 (15.5)	12 (44.4)	15 (55.5)		
	50-54	35 (20.1)	15 (42.9)	20 (57.1)		
	55-59	14 (8.1)	10 (71.4)	4 (28.6)		
	> 60	13 (7.5)	3 (23.1)	10 (76.9)		
	Missing	2 (1.1)	0 (0.0)	2 (100)		
Race	Malay	161 (92.5)	90 (55.9)	71 (44.1)	0.752	
	Chinese	6 (3.5)	4 (66.7)	2 (33.3)		
	Other	7 (4.0)	3 (42.9)	4 (57.1)		
Education level	Primary school	16 (9.2)	10 (62.5)	6 (37.5)	0.396	
	Secondary school	96 (55.2)	49 (51.0)	47 (49.0)		
	College / University	57 (32.7)	35 (61.4)	22 (38.6)		
	Missing	5 (2.9)	3 (60.0)	2 (40.0)	-	
Marital status	Married	157 (90.2)	91 (58.0)	66 (42.0)	0.15	
	Divorced	8 (4.6)	2 (25.0)	6 (75.0)		
	Widowed	9 (5.2)	4 (44.4)	5 (55.6)		
Occupation	Housewife	64 (36.8)	39 (60.9)	25 (39.1)	0.087	
•	Government employee	67 (38.5)	41 (61.2)	26 (38.8)		
	Private employee	31 (17.8)	13 (41.9)	18 (58.1)		
	Retired	9 (5.2)	3 (33.3)	6 (66.7)		
	Unemployed	1 (0.6)	1 (100)	0 (0.0)		
	Other	2 (1.1)	0 (0.0)	2 (100)		
Monthly household income	< \$500	27 (15.5)	15 (55.6)	12 (44.4)	0.498	
,	\$500 < \$999	27 (15.5)	12 (44.4)	15 (55.6)		
	\$1000-\$1999	40 (23.0)	22 (55.0)	18 (45.0)		
	\$2000-\$2999	19 (10.9)	14 (73.7)	5 (26.3)		
	\$3000-\$5000	24 (13.8)	15 (62.5)	9 (37.5)		
	>\$5000	3 (1.7)	2 (66.7)	1 (33.3)		
	Missing	34 (19.5)	17 (50.0)	17 (50.0)		
Number of births	0	27 (15.5)	17 (63.0)	10 (37.0)	0.103	
	1	19 (10.9)	14 (73.7)	5 (26.3)		
	2	22 (12.1)	14 (66.7)	7 (33.3)		
	3 or more	105 (60.3)	51 (48.6)	54 (51.4)		
	Missing	2 (1.2)	1 (50.0)	1 (50.0)		
Last Pap test done	Never	41 (23.6)	29 (70.7)	12 (29.3)	0.031	
up toot done	4–10 years	91 (52.3)	48 (52.7)	43 (47.3)	- 0.001	
	> 10 years	36 (20.7)	15 (41.7)	21 (58.3)		
	Missing	6 (3.4)	5 (83.3)	1 (16.7)		

(Continued)

Table 1. (Continued)

Characteristics		Total study population (n = 174)	Joined self-sampling (n = 97)	Declined self-sampling (n = 77)	p-value
		n (%)	n (%)	n (%)	
HPV vaccination status	Unvaccinated	95 (54.6)	50 (52.6)	45 (47.4)	0.483
	Fully vaccinated	27 (15.5)	16 (59.3)	11 (40.7)	
	Partly vaccinated	41 (23.6)	26 (63.4)	15 (36.6)	
	Missing	11 (6.3)	5 (45.5)	6 (54.5)	

IQR = Interquartile range.

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that they are more susceptible to develop cervical cancer were significantly more likely to join HPV self-sampling (p = 0.023).

Most participants would like to obtain more information about cervical cancer screening (85.6%, n = 149). The top preferred information sources were healthcare workers (59.8%, n = 104), social messaging platforms (28.7%, n = 50) and social media (26.4%, n = 46; S1 Fig).

Among those who participated in self-sampling (55.7%, n = 97), their responses on Q2 (Table 4) were mostly positive. A majority agreed that the instructions were clear (94.8%, n = 92), that it was easy to perform the swab (93.8%, n = 91), and that it was more convenient than the Pap test (91.7%, n = 89). They also reported their confidence in correctly getting the sample (92.8%, n = 90), would prefer to use this method next time (94.8%, n = 92), and would recommend this method to other women (93.8%, n = 91). Notably, 54.6% (n = 53) still prefer a proper Pap test for their subsequent check-up.

Among the 97 samples taken, nine (9.3%) tested positive for hr-HPV: one was positive for HPV 16 and eight were positive for non-HPV 16/18 HPV genotype. The HPV 16 positive case was found to have high-grade ASC-US (ASCUS-H) on the initial follow up smear, but found negative after subsequent follow up cervical biopsy. Among the other 8 non-HPV 16/18 HPV genotype positive cases: 2 had negative follow up smears, 4 were reported to have ASC-US on their initial follow up smears, but had subsequent negative follow up smears, 1 was reported to

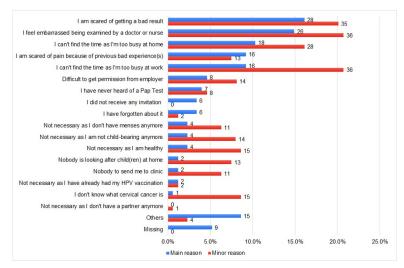


Fig 2. Responses on their major and minor reasons for not attending cervical cancer screening, among non-attendees at JPSHC (Jan-Dec 2019). The x-axis indicates the percentage, and the number next to each bar indicates the actual number of responses.

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Table 2. The top ten major reasons of not attending cervical cancer screening at JPSHC, Brunei (Jan-Dec 2019), with comparison between those who agreed and declined HPV self-sampling. Responses from 146 participants (83.9% of the total study population) were included.

No.	Top ten major reasons of screening non-attendance	Total	Joined HPV self-sampling	Declined HPV self-sampling	p-value
		n (%)	n (%)	n (%)	
1	I feel embarrassed being examined by a doctor or nurse	26 (14.9)	20 (76.9)	6 (23.1)	0.020*
2	I am scared of pain because of previous bad experience(s)	16 (9.2)	8 (50.0)	8 (50.0)	0.825
3	I am scared of getting a bad result	28 (16.1)	10 (35.7)	18 (64.3)	0.034*
4	I can't find the time as I'm too busy at home	18 (10.3)	12 (66.7)	6 (33.3)	0.463
5	I can't find the time as I'm too busy at work	16 (9.2)	10 (62.5)	6 (37.5)	0.759
6	Difficult to get permission from employer	8 (4.6)	5 (62.5)	3 (37.5)	1
7	I have never heard of a Pap Test	7 (4.0)	4 (57.1)	3 (42.9)	1
8	I have forgotten about it	6 (3.4)	5 (83.3)	1 (16.7)	0.229
9	I did not receive any invitation	6 (3.4)	5 (83.3)	1 (16.7)	0.229
10	Others	15 (8.6)	7 (46.7)	8 (53.3)	0.492

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have low-grade squamous intraepithelial lesion (LSIL) on her initial follow up smear, and had a subsequent negative follow up smear, and 1 was found to have CIN 3 with glandular involvement on the follow up smear and cervical biopsy, and has received treatment with a cone biopsy where excision of CIN 3 was confirmed. No significant differences were observed when comparing the sociodemographic characteristics between those who tested hr-HPV positive and those who tested hr-HPV negative (S3 Table).

Discussion

Our study findings highlight three important points to consider for improving cervical cancer screening uptake and detection. First, our findings suggest that it is necessary to provide accurate information among Bruneian women on cervical cancer, the importance of screening and addressing any misconceptions about the Pap test. Important facts to relay include the slow development from pre-cancerous changes to cervical cancer, that pre-cancerous changes are highly treatable, and that screening will help in early detection and thus facilitate successful treatment.

Table 3. Attitudes towards cervical cancer screening among non-attendees at JPSHC, Brunei (Jan-Dec 2019), between those who agreed and declined HPV self-sampling.

No.	Attitude questions		Total study population (n = 174)	Joined HPV self-sampling (n = 97)	Declined HPV self-sampling (n = 77) n (%)	p-value
			n (%)	n (%)		
1	I believe I am healthy and free of any diseases	Agree	62 (35.6)	34 (54.8)	28 (45.2)	0.984
		Neutral/ Disagree	112 (64.3)	63 (56.3)	49 (43.7)	
2	Having Pap test taken is beneficial for my health	Agree	160 (92.0)	91 (56.9)	69 (43.1)	0.464
		Neutral/ Disagree	14 (8.0)	6 (42.9)	8 (57.1)	
3	Like any women, I am susceptible to develop cervical cancer	Agree	110 (63.2)	69 (62.7)	41 (37.3)	0.023*
		Neutral/ Disagree	64 (36.8)	28 (43.8)	36 (56.2)	
4	Cervical cancer can be severe and may lead to death	Agree	144 (82.8)	79 (54.9)	65 (45.1)	0.754
		Neutral/ Disagree	30 (17.2)	18 (60.0)	12 (40.0)	

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Table 4. Acceptability of HPV self-sampling among non-attendees who participated in HPV self-sampling at JPSHC, Brunei (Jan-Dec 2019). Responses from all Q2 respondents (n = 97) were included.

No.	Self-sampling Questions	Agree	Neutral	Disagree	
		n (%)	n (%)	n (%)	
1	I thought the instructions were clear	92 (94.8)	0 (0.0)	5 (5.2)	
2	It was easy to do the swab	91 (93.8)	1 (1.0)	5 (5.2)	
3	Taking the sample with the swab was painful	12 (12.4)	8 (8.2)	77 (79.4)	
4	Taking the sample was uncomfortable to do	11 (11.4)	4 (4.1)	82 (84.5)	
5	I felt embarrassed doing the self-sampling	9 (9.3)	1 (1.0)	87 (89.7)	
6	It was convenient to do without having to undergo a Pap Test	89 (91.7)	3 (3.1)	5 (5.2)	
7	I am confident I did it correctly	90 (92.8)	4 (4.1)	3 (3.1)	
8	I want to use this method next time	92 (94.8)	2 (2.1)	3 (3.1)	
9	I prefer to do this at home	64 (66.0)	15 (15.5)	18 (18.5)	
10	I would recommend this method to other women	91 (93.8)	3 (3.1)	3 (3.1)	
11	I trust that the result of this self-sampling will be accurate	74 (76.3)	19 (19.6)	4 (4.1)	
12	I would like to attend for a proper Pap test in clinic next time	53 (54.6)	23 (23.7)	21 (21.7)	

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In our study, emotional barriers (fear of unfavorable test results and embarrassment) were the most common major reasons of screening non-attendance. Also, about two-thirds of our participants cited "I am scared of getting a bad result" as their main barrier also declined to take part in HPV self-sampling, possibly due to relating abnormal Pap test results to cervical cancer diagnoses [25]. Although such barriers may play a large role at the onset of screening program, its role diminishes over time with increasing education on the benefits of screening [26]. Educational interventions could also benefit the small group of women who cited menopause, cessation of child-bearing and having had HPV vaccination as reasons for not attending screening (Fig 2 and S1 Table). We also observed that those who agree that they are susceptible to cervical cancer were significantly more likely to participate in HPV self-sampling. This suggests that perceived susceptibility could be an important factor for self-sampling participation, whereby those who do not perceive themselves as susceptible were less likely to engage in preventative behaviors [27-29]. Perceived susceptibility can be increased through education to improve their beliefs on the importance of screening [30]. We suggest that such information could be more effectively disseminated as simple health messages endorsed by Brunei's Ministry of Health via website and social media platforms.

Second, we observed high acceptability of HPV self-sampling among our participants. Meta-analyses have indicated strong acceptance of and preference for self-sampling over clinician sampling [31], mainly due to logistical reasons [32]. More than half of our participants who joined HPV self-sampling cited embarrassment or lack of time due to home and/or work commitments as their major reason of not attending Pap test screening. This suggest that providing flexibility to accommodate women's screening method preference [33], such as the option of self-sampling [34], could improve screening uptake.

Thirdly, most of the detected hr-HPV genotypes in our study were non-16/18, with only one out of nine participants tested positive for HPV 16. Although our sample size is small, this result suggests that it may not be accurate to assume HPV 16 or 18 as common hr-HPV genotypes in Brunei, even though this is true in the global context [35]. Other studies have detected a significant percentage of non-16/18 hr-HPV genotypes, suggesting the presence of region-specific heterogeneity in the HPV genotype distribution [35–39]. Also, variation in HPV distribution among different ethnic groups has been reported in an American study [40]. Our finding has potential implication on Brunei's national school-based HPV vaccination program

[41] which currently provides vaccines which do not confer protection against non-16/18 genotypes. Larger population-based studies to understand the distribution of HPV genotypes among Bruneian women will be crucial to determine the efficacy or impact of the current vaccines.

One notable point for the local context relates to the presence of cultural barriers. Being a predominantly Islamic society, religious and cultural modesty could be a contributing factor for embarrassment among Muslim women [42]. Also, having premarital sex is a taboo in Brunei and is generally not openly discussed [43]. This could prevent any unmarried but sexually active women from participating in the screening program. It should be emphasized that only married or ever married females were included in this study; included because it is part of the eligibility criteria for the national cervical cancer screening program in Brunei.

A major limitation for this study is that non-attendees from only one health center were recruited, thus our findings are not representative of the adult female population in Brunei. Secondly, our findings should also be interpreted with caution due to the small sample size and the non-probability sampling approach used. This study was initially conceived as a pilot study due to resource and logistics limitations. There were two reasons for choosing JPSHC as our study site: It is the third largest primary government healthcare center in the country, and that it serves a sub-district with a sizable percentage of residents in the middle- to low-income groups. However, even at this pilot stage, our study findings could encourage stakeholders to conduct similar and larger studies, using random sampling approach. Lastly, as this study relied on the self-reported history of previous Pap test attendance, we might have missed recruiting those who may have forgotten their last Pap test date.

In conclusion, our findings indicate the need to further promote knowledge on cervical cancer, the benefits of screening and clarifying any misconceptions of Pap test results. Reasons of cervical cancer screening non-attendance were mainly related to emotional and logistical factors. As we found high acceptance towards HPV self-sampling, this could be adopted as an alternative for women who refrain from Pap test. Our high detection of non-16/18 HPV genotypes suggest high prevalence of other hr-HPV genotypes in Brunei. Future larger studies involving more Bruneian women should be done to verify our results. Follow-up studies should also be conducted to consider HPV testing as the suitable method for cervical cancer screening.

Supporting information

S1 Table. Responses on their major and minor reasons for not attending cervical cancer screening among non-attendees at JPSHC, Brunei (Jan-Dec 2019). (DOCX)

S2 Table. Attitudes towards cervical cancer screening among non-attendees at JPSHC, Brunei (Jan-Dec 2019). Responses from the total study population (n = 174) were included. (DOCX)

S3 Table. Socio demographic characteristics and comparison between screening non-attendees who tested positive and negative for hr-HPV at JPSHC, Brunei (Jan-Dec 2019). (DOCX)

S1 Fig. Preferred sources of information about cervical cancer among non-attendees at JPSHC, Brunei (Jan–Dec 2019). The x-axis indicates the percentage, and the number next to each bar indicates the number of responses. Multiple responses were allowed and responses from the total study population (n = 174) were included. (DOCX)

S1 Questionnaire. Reasons why women do not participate in the national cervical cancer screening program.

(DOCX)

S2 Questionnaire. Acceptability of self sampling for HPV testing among non-attendees of cervical screening program in Brunei Darussalam.

(DOCX)

S1 Dataset. Minimal dataset.

(XLSX)

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References

- World Health Organization. WHO guideline for screening and treatment of cervical pre-cancer lesions for cervical cancer prevention (2nd edition). 2021. Available: https://www.who.int/publications/i/item/ 9789240030824
- Gustafsson L, Pontén J, Zack M, Adami H-O. International incidence rates of invasive cervical cancer after introduction of cytological screening. Cancer Causes Control. 1997; 8: 755–763. https://doi.org/10.1023/a:1018435522475 PMID: 9328198
- 3. Ministry of Health Brunei Darussalam. Health Information Booklet 2017 (Second Revision). 2019 [cited 11 Feb 2020]. Available: http://moh.gov.bn/SitePages/Health Information Booklet.aspx
- Arbyn M, Weiderpass E, Bruni L, de Sanjosé S, Saraiya M, Ferlay J, et al. Estimates of incidence and mortality of cervical cancer in 2018: a worldwide analysis. Lancet Glob Heal. 2020; 8: e191–e203. https://doi.org/10.1016/S2214-109X(19)30482-6 PMID: 31812369
- WHO. Guidelines for screening and treatment of precancerous lesions for cervical cancer prevention. WHO Guidel. 2013.

- Bosgraaf RP, Ketelaars PJW, Verhoef VMJ, Massuger LFAG, Meijer CJLM, Melchers WJG, et al. Reasons for non-attendance to cervical screening and preferences for HPV self-sampling in Dutch women. Prev Med (Baltim). 2014. https://doi.org/10.1016/j.ypmed.2014.04.011 PMID: 24736093
- Virtanen A, Nieminen P, Niironen M, Luostarinen T, Anttila A. Self-sampling experiences among nonattendees to cervical screening. Gynecol Oncol. 2014. https://doi.org/10.1016/j.ygyno.2014.09.019
 PMID: 25284037
- 8. Knops-Dullens T, de Vries N, de Vries H. Reasons for non-attendance in cervical cancer screening programmes: an application of the Integrated Model for Behavioural Change. Eur J Cancer Prev. 2007; 16. Available: https://journals.lww.com/eurjcancerprev/Fulltext/2007/10000/Reasons_for_non_attendance_in_cervical_cancer.9.aspx https://doi.org/10.1097/01.cej.0000236250.71113.7c PMID: 17923815
- Waller J, Bartoszek M, Marlow L, Wardle J. Barriers to cervical cancer screening attendance in England: a population-based survey. J Med Screen. 2009; 16: 199–204. https://doi.org/10.1258/jms.2009.009073 PMID: 20054095
- Oscarsson MG, Benzein EG, Wijma BE. Reasons for non-attendance at cervical screening as reported by non-attendees in Sweden. J Psychosom Obstet Gynecol. 2008; 29: 23–31. https://doi.org/10.1080/01674820701504619 PMID: 18266164
- Yeh PT, Kennedy CE, Vuyst H de, Narasimhan M. Self-sampling for human papillomavirus (HPV) testing: a systematic review and meta-analysis. BMJ Glob Heal. 2019; 4: e001351. https://doi.org/10.1136/bmigh-2018-001351 PMID: 31179035
- CS R, DR W, D G. Self-collected HPV testing improves participation in cervical cancer screening: a systematic review and meta-analysis. Can J Public Health. 2013; 104. https://doi.org/10.1007/BF03405681 PMID: 23618210
- Maza M, Melendez M, Masch R, Alfaro K, Chacon A, Gonzalez E, et al. Acceptability of self-sampling and human papillomavirus testing among non-attenders of cervical cancer screening programs in El Salvador. Prev Med (Baltim). 2018. https://doi.org/10.1016/j.ypmed.2018.06.017 PMID: 29958860
- Sancho-Garnier H, Tamalet C, Halfon P, Leandri FX, Retraite L Le, Djoufelkit K, et al. HPV self-sampling or the Pap-smear: A randomized study among cervical screening nonattenders from lower socioeconomic groups in France. Int J Cancer. 2013. https://doi.org/10.1002/ijc.28283 PMID: 23712523
- Chrysostomou AC, Stylianou DC, Constantinidou A, Kostrikis LG. Cervical cancer screening programs in Europe: The transition towards HPV vaccination and population-based HPV testing. Viruses. 2018; 10. https://doi.org/10.3390/v10120729 PMID: 30572620
- 16. Arrossi S, Thouyaret L, Herrero R, Campanera A, Magdaleno A, Cuberli M, et al. Effect of self-collection of HPV DNA offered by community health workers at home visits on uptake of screening for cervical cancer (the EMA study): A population-based cluster-randomised trial. Lancet Glob Heal. 2015; 3: e85–e94. https://doi.org/10.1016/S2214-109X(14)70354-7 PMID: 25617202
- Gustavsson I, Aarnio R, Berggrund M, Hedlund-Lindberg J, Strand A-S, Sanner K, et al. Randomised study shows that repeated self-sampling and HPV test has more than two-fold higher detection rate of women with CIN2+ histology than Pap smear cytology. Br J Cancer. 2018; 118: 896–904. https://doi.org/10.1038/bjc.2017.485 PMID: 29438367
- Arrossi S, Paolino M, Laudi R, Gago J, Campanera A, Marín O, et al. Programmatic human papillomavirus testing in cervical cancer prevention in the Jujuy Demonstration Project in Argentina: a population-based, before-and-after retrospective cohort study. Lancet Glob Heal. 2019. https://doi.org/10.1016/S2214-109X(19)30048-8 PMID: 31097279
- Landy R, Pesola F, Castañón A, Sasieni P. Impact of cervical screening on cervical cancer mortality: Estimation using stage-specific results from a nested case-control study. Br J Cancer. 2016; 115: 1140–1146. https://doi.org/10.1038/bjc.2016.290 PMID: 27632376
- Canfell K, Sitas F, Beral V. Cervical cancer in Australia and the United Kingdom: Comparison of screening policy and uptake, and cancer incidence and mortality. Med J Aust. 2006; 185: 482–486. https://doi.org/10.5694/j.1326-5377.2006.tb00661.x PMID: 17137451
- Bayu H, Berhe Y, Mulat A, Alemu A. Cervical Cancer Screening Service Uptake and Associated Factors among Age Eligible Women in Mekelle Zone, Northern Ethiopia, 2015: A Community Based Study Using Health Belief Model. PLoS One. 2016; 11: e0149908. https://doi.org/10.1371/journal.pone.0149908 PMID: 26963098
- Sultana F, Mullins R, English DR, Simpson JA, Drennan KT, Heley S, et al. Women's experience with home-based self-sampling for human papillomavirus testing. BMC Cancer. 2015; 15: 849. https://doi. org/10.1186/s12885-015-1804-x PMID: 26536865
- 23. The Hague: Health Council of the Netherlands. Health Council of the Netherlands Population screening for cervical cancer. 2011; 2011/07E. Available: https://www.gezondheidsraad.nl/sites/default/files/201107E PopulationSCC 0.pdf

- R Core Team. R: A language and environment for statistical computing. Vienna, Austria; 2021. Available: http://www.r-project.org/
- Fylan F. Screening for cervical cancer: a review of women's attitudes, knowledge, and behaviour. Br J Gen Pract. 1998; 48: 1509–14. PMID: 10024713
- 26. Teng FF, Mitchell SM, Sekikubo M, Biryabarema C, Byamugisha JK, Steinberg M, et al. Understanding the role of embarrassment in gynaecological screening: A qualitative study from the ASPIRE cervical cancer screening project in Uganda. BMJ Open. 2014. https://doi.org/10.1136/bmjopen-2014-004783 PMID: 24727360
- Agurto I, Arrossi S, White S, Coffey P, Dzuba I, Bingham A, et al. Involving the community in cervical cancer prevention programs. Int J Gynaecol Obstet. 2005; 89 Suppl 2: S38–45. https://doi.org/10.1016/ j.ijgo.2005.01.015 PMID: 15823265
- 28. Shirazi Zadeh Mehraban S, Namdar A, Naghizadeh MM. Assessment of Preventive Behavior for Cervical Cancer with the Health Belief Model. Asian Pac J Cancer Prev. 2018; 19: 2155–2163. https://doi.org/10.22034/APJCP.2018.19.8.2155 PMID: 30139219
- Tapera R, Manyala E, Erick P, Maswabi TM, Tumoyagae T, Letsholo B, et al. Knowledge and Attitudes towards Cervical Cancer Screening amongst University of Botswana Female Students. Asian Pac J Cancer Prev. 2017; 18: 2445–2450. https://doi.org/10.22034/APJCP.2017.18.9.2445 PMID: 28952274
- Khademolhosseini F, Noroozi A, Tahmasebi R. The Effect of Health Belief Model-Based Education through Telegram Instant Messaging Services on Pap smear performance. Asian Pac J Cancer Prev. 2017; 18: 2221–2226. https://doi.org/10.22034/APJCP.2017.18.8.2221 PMID: 28843259
- Nelson EJ, Maynard BR, Loux T, Fatla J, Gordon R, Arnold LD. The acceptability of self-sampled screening for HPV DNA: A systematic review and meta-analysis. Sexually Transmitted Infections. 2017. https://doi.org/10.1136/sextrans-2016-052609 PMID: 28100761
- 32. Lorenzi NPC, Termini L, Longatto Filho A, Tacla M, de Aguiar LM, Beldi MC, et al. Age-related acceptability of vaginal self-sampling in cervical cancer screening at two university hospitals: a pilot cross-sectional study. BMC Public Health. 2019. https://doi.org/10.1186/s12889-019-7292-1 PMID: 31319838
- **33.** Bansil P, Wittet S, Lim JL, Winkler JL, Paul P, Jeronimo J. Acceptability of self-collection sampling for HPV-DNA testing in low-resource settings: a mixed methods approach. BMC Public Health. 2014; 14: 596. https://doi.org/10.1186/1471-2458-14-596 PMID: 24927941
- Jeronimo J, Perkins RB, Scalici J, Pierce JY. Should Self-Sampling Be an Option for Women in the United States? J Low Genit Tract Dis. 2019; 23: 54. https://doi.org/10.1097/LGT.0000000000000453 PMID: 30586018
- **35.** Albawardi A, Quddus MR, Al Awar S, Almarzooqi S. Frequency of rare and multi viral high-risk HPV types infection in cervical high grade squamous intraepithelial lesions in a non-native dominant middle eastern country: A polymerase chain reaction-based pilot study. Diagn Pathol. 2018. https://doi.org/10.1186/s13000-018-0716-x PMID: 29945637
- 36. Zeng Z, Yang H, Li Z, He X, Griffith CC, Chen X, et al. Prevalence and Genotype Distribution of HPV Infection in China: Analysis of 51,345 HPV Genotyping Results from China's Largest CAP Certified Laboratory. J Cancer. 2016. https://doi.org/10.7150/jca.14971 PMID: 27326245
- Khoo SP, Bhoo-Pathy N, Yap SH, Anwar Shafii MK, Hairizan Nasir N, Belinson J, et al. Prevalence and sociodemographic correlates of cervicovaginal human papillomavirus (HPV) carriage in a cross-sectional, multiethnic, community-based female Asian population. Sex Transm Infect. 2018. https://doi.org/10.1136/sextrans-2017-053320 PMID: 29180538
- 38. Peng RR, Li HM, Chang H, Li JH, Wang AL, Chen XS. Prevalence and genotype distribution of cervical human papillomavirus infection among female sex workers in Asia: A systematic literature review and meta-analysis. Sexual Health. 2012. https://doi.org/10.1071/SH11066 PMID: 22498154
- Al-Awadhi R, Chehadeh W, Jaragh M, Al-Shaheen A, Sharma P, Kapila K. Distribution of human papillomavirus among women with abnormal cervical cytology in Kuwait. Diagn Cytopathol. 2013. https://doi.org/10.1002/dc.21778 PMID: 21987449
- Keller MJ, Burk RD, Massad LS, Eltoum IE, Hessol NA, Anastos K, et al. Racial differences in human papilloma virus types amongst United States women with HIV and cervical precancer. AIDS. 2018. https://doi.org/10.1097/QAD.0000000000000005 PMID: 30234608
- Chaw LL, Lim STW, Md Yussof SR. Human Papillomavirus vaccine coverage among female students in Brunei Darussalam: results from the first 4 years of the national school-based vaccination programme. Heliyon. 2019; 5. https://doi.org/10.1016/j.heliyon.2019.e02588 PMID: 31692704
- **42.** Ali S, Skirton H, Clark MT, Donaldson C. Integrative review of cervical cancer screening in Western Asian and Middle Eastern Arab countries. Nurs Heal Sci. 2017; 19: 414–426. https://doi.org/10.1111/nhs.12374 PMID: 29058371

43. Haji Tahamit N. Cultural and Religious Barriers to Setting Up Sex and Relationship Education in a Muslim Country: A Case Study of Brunei Darussalam. University of Leeds; 2015 Aug. Available: https://etheses.whiterose.ac.uk/15306/