Effectiveness of Visual Inspection with Acetic Acid versus Papanicolaou Smear for Cervical Cancer Screening at Kirtipur Hospital

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Aims: The aim of this study was to compare the accuracy of detecting precancerous cervical cells, via a visual inspection of the cervix and an application of acetic acid (VIA) with the Papanicolaou (Pap) smear, to screen for cervical cancer in resource poor countries.

Methods: The study involved 189 women between the ages of 30-60 years, who attended the Gynaecological Out Patient Department at Kirtipur hospital from 1st December 2014 to 31st March 2015. A VIA and Pap smear was done in all women, with cervical biopsies conducted in VIA positive women.

Results: The majority of these women were in the age range of 30-39 years, non-smokers, housewives, multipara and non-users of contraception. Of the tested group 8 (4.2%) had a history of post coital bleeding, 64 (21.2%) presented with lower abdominal pain, 64 (33.9%) presented with suspected abnormal vaginal discharge and 34 (17.98%) with dysfunctional uterine bleeding. Out of 189 women the VIA was positive in 5 (2.62%) and a Pap smear was positive in 2 (1.1%). By comparison to a Pap smear the VIA has an accuracy of 100%, specificity 98.4%, positive predictive value 40% and negative predictive value 100%.

Conclusions: A visual inspection of the cervix with acetic acid performed by a qualified physician is a suitable alternative for precancerous cell screening in low resource settings.

Keywords: acetic acid; cervical cancer; Pap smear; Papanicolaou smear; visual inspection.

INTRODUCTION
Cervical cancer remains the second most common cancer in women worldwide and the leading cause of cancer related death. Of the approximately 500,000 new cases reported annually almost 300,000 women will die with 88% of those occurring in developing countries. It is estimated that more than 10,000 new cases of invasive cancer arise in Nepal each year and that 26,000-45,000 women currently have an undiagnosed precancerous lesion at this moment.

Most developed countries have shown a dramatic reduction in the incidence and death rate from cervical cancer following the implementation of organized screening and awareness programs. It has been estimated that only 5% of women in developing countries have been screened for cervical dysplasia in the past 5 years compared with 85% in developed countries.

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In a developing country like Nepal it is difficult to implement new organized screening programs such as liquid based cytology, HPV DNA testing or automated Pap testing. Conventional Papanicolaou (Pap) smear testing requires good infrastructure, specialized personnel and multiple visits by the woman for follow up and/or treatment, it can become time consuming and expensive. Whilst a visual inspection with administration of acetic acid (VIA) is a cost effective, simple and real-time screening with accuracy that is comparable to a good quality Pap smear. The cervix is easily accessible and has a longstanding precancerous stage known as intraepithelial lesion (CIN) and unlike many cancers, cervical cancer can be prevented with appropriate screening and treatment. VIA could be a possible alternative screening tool for early detection of cervical cancer in Nepal where cervical cancer is the most frequent cause of cancer death among the female population.

METHODS
This is a comparative study conducted among 189 women, who attended the Gynaecological Outpatient Clinic at Kirtipur Hospital from 1 December 2014
to 31 March 2015. The study was ethically approved by IRC (Institutional Review Committee) of phect-Nepal. In this study, women between the ages of 30-60 years were included with a broad cross section of those who were unmarried, pregnant, had active vaginal bleeding, had a recent Pap smear, had a frank growth on the cervix, had a hysterectomy and had a previous history of treatment for cancerous lesions were included but patients with a history of an allergic reaction to acetic acid were excluded. All relevant obstetric and gynaecological history were taken as well as information such as smoking habits, number of sexual partners, any abnormal vaginal discharge, postcoital bleeding, menstrual irregularities and age of marriage were also recorded. All women involved in this study were educated about cancer of the cervix, and all procedures to be performed were explained with written informed consent given by each.

All women were given a general physical examination and, after proper positioning, a local vulvar examination was performed. Cusco’s vaginal speculum was inserted and fixed to view the cervix. Any visible abnormal discharge was noted, the squamo-columnar junction was visualized and with the hooked end of Ayre’s spatula the squamo-columnar area was scraped gently around the circumference and the collected material was transferred to a glass slide that was put into a container with 95% alcohol. Following the Pap smear, abnormal discharge such as blood and mucus from the cervix was removed by normal saline soaked cotton swabs, then a solution of 5% acetic acid was applied to the cervix using a cotton swab. The cervix was visualized after one minute using an adequate light source and the appearance of any distinct acetowhite opaque areas at the transformation zone touching the squamo-columnar junction was considered to be a positive result. The women with positive results were educated about colposcopic examination and cervical biopsy. Biopsies were taken using punch biopsy forceps from the white and opaque areas within the transformation zone, the specimen were then transferred inside a container in formalin and sent to the pathological laboratory at Kathmandu Model Hospital. Results were collected then recorded for analysis using the Chi-square statistical test.

**RESULTS**

During the study period 189 women attending the Gynaecological Out Patient Department of Kirtipur Hospital were enrolled, a majority of the women were of the age group 30-39 years, non-smoker, housewives, multipara and non-user of contraception. Only 8 (4.2%) women had a history of postcoital bleeding, 64 (21.2%) women had lower abdominal pain, 64 (33.9%) women presented with suspected abnormal vaginal discharge and 34 (17.98%) women with dysfunctional uterine bleeding.

| Table 1. Age and parity of the study population. |
|-----------------|-----------------|-----------------|
| **Age in years** | **Number** | **Percentage** |
| 30-39            | 90            | 47.6           |
| 40-49            | 74            | 39.1           |
| 50-60            | 25            | 13.3           |
| **Parity**       |               |                |
| Nullipara        | 8             | 4.2            |
| P1-4             | 171           | 90.5           |
| >=P5             | 10            | 5.3            |

| Table 2. Relation between Pap smear and VIA. |
|-----------------|-----------------|-----------------|
| **VIA**         | **Pap positive**| **Pap negative**| **Total**|
| Positive        | 2              | 3              | 5 | p-0.001 |
| Negative        | 0              | 184            | 184 |
| Total           | 2              | 187            | 189 |

Out of five VIA positive cases two were also Pap positive. The table above shows the probability of having a Pap positive if the VIA is positive as 0.001. Sensitivity of the VIA and Pap test were 100%, whereas specificity was 98.4%, positive predictive value was 40% and negative predictive value was 100%. Biopsies were taken in all VIA positive women. Out of 5 VIA positive women 2 (40%) women were found to be positive for intraepithelial cervical neoplasia with a biopsy.

**DISCUSSION**

Cervical cancer is the leading cause of morbidity and mortality among women worldwide. With a successful screening programme 80% of incidences and mortality due to cervical cancer have declined in developed countries. Screening programs have been introduced into developing countries since the early 1980’s yet have failed to reduce the mortality rate. WHO, in 2002, estimated that only 5% of women in developing countries are screened appropriately. Likely reasons for failure in screening programs include lack of funding, insufficient access in rural areas, where most of the population...
in developing countries reside, lack of awareness/education as to the need for screening and poor follow up. Thus visual inspection using acetic acid (VIA) has emerged as a promising, cost effective, non-cytological based alternative for economically underprivileged geographical regions. The molecular basis is that acetic acid causes dehydration of cells and surface coagulation of cellular proteins thereby reducing the transparency of the cervical epithelium. These changes are more pronounced in abnormal epithelium due to greater nuclear density and consequently a higher concentration of proteins. A positive test is based on the detection of well-defined acetowhite epithelium one minute after the application of 3-5% acetic acid at the transformation zone.

A total of 189 women aged 30-60 years were enrolled in this study. Satyanarayan et al completed a study in the same age group but our study shows precancerous lesions in women with complaints of suspected abnormal vaginal discharge and dysfunctional uterine bleeding. Laddad et al also found precancerous lesions in 70% of VIA positive women with similar complaints. It is well known that early marriage is a high risk factor for cervical cancer, precancerous lesions were seen in women with a social history of early marriage (<18 years) in this study also. Bhattacharya et al also has reported that incidences of intraepithelial lesions were more common in women whose marriage age was 18 or less. Some studies reported high parity as a high risk factor for cervical cancer but in our study precancerous lesions were found in para 2, possibly due to early marriage.

In our study VIA and Pap smears were done in all women and cervical biopsies were taken only in cases that were VIA positive, similar studies were done by Begum et al. Unlike our study Musta et al performed cervical biopsies in all women with abnormal colposcopic findings whilst Tayyeb et al took biopsies not only from women who screened positive but also from women who had an abnormal cervix in colposcopy and advanced lesions but screened negative.

The VIA positive rate in our study was 2.62% this result was in agreement with the report of Dhaubhadel et al that showed the VIA rate as 2.8%. But the result was in disagreement with the findings of Okonkwo and Bharani et al who reported 8.4%, and 26% respectively. These variations in VIA positive rates could be due to differing criteria for the screening of cervical cancer. In the present study the sensitivity of the VIA test was 100% as there were no cases with a dysplastic lesion which was not picked up by VIA. Similar results were shown by Bharani et al and Sheesa et al in their studies. The specificity of VIA in our study was 98.4% which is similar to the findings 98.7% reported by Harsini et al. However this result was in disagreement with the findings of Sheesa and Shammati who have reported lower VIA specificity of 45% and 66.7% respectively. The high rate of sensitivity and specificity of the VIA test indicates that it is also a useful screening tool for cervical cancer. The positive predictive value of VIA was 40% in this study which was higher than 32% reported by Bhattacharya et al. But other studies showed higher positive predictive values, of 73% and 84.6% respectively, than this study. Our study shows that there is a chance of over treatment but in our context most women were lost to follow up and later presented with the late stages of cervical cancer, thus the “see and treat” method is preferable to preventing cervical cancer in resource poor environments. The present study shows the negative predictive value of VIA was 100%, this result was in agreement with the results of Mahamud et al of 98.6%. Unlike our study, Vahedra et al showed lower negative predictive value of 85.7% but since VIA gives immediate results and has a high negative predictive value, women with a negative VIA can be assured immediately that she is precancerous lesion free. Our study shows Pap smears are an effective screening tool but at the same time VIA is a more appropriate screening method for cervical cancer in developing countries. WHO also has recommended VIA as an alternative to cytology to identify patients at risk of cervical cancer.

CONCLUSIONS
VIA is a quick, safe, easy to perform, cost effective screening process that does not require specialized knowledge or instruction and results are available immediately, thus it has a role in screening program of cervical cancer in low resource countries like Nepal.

Conflict of Interest
The authors report no conflicts of interest in this work. No violation of human rights and safety.

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REFERENCES


