CERVICO-VAGINAL SWABS TO PREDICT PROM: A CASE STUDY

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ABSTRACT: BACKGROUND: The presence of lower genital tract infections poses a threat to the health of a woman. During pregnancy, these infections may also have a direct effect on the fetus or may indirectly predispose to fetal damage secondary to premature labour or premature rupture of membranes. Early detection is thus important because maternal physiological alterations and the presence of a fetus may hamper the diagnosis and the management of such infections. Antepartum treatment of lower genital tract infections or bacterial colonization has been found to reduce the incidence of preterm births. OBJECTIVES: To study the impact of education and socio-economic background, to assess diagnostic efficacy in pregnancy, to study the effect of intervention on the outcome of pregnancy and finally, to study the appropriate trimester for taking a cervicovaginal swab in pregnant women. MATERIALS AND METHODS -Our case study was a prospective study involving 100 pregnant women following up in the outpatient department of a community hospital from 2008-2010 with respect to their cervicovaginal microbial growth in each trimester. Patients were divided into culture positive and negative groups and treated accordingly RESULTS - In our study, we found that streptococcus, candida and staphylococcus were the most commonly prevalent among all the microorganisms isolated. Out of 100 women, 27 were swab positive in the 1st trimester with an increasing trend of 1% in swab positivity in subsequent trimesters. Among swab positive cases, $51.85\%(1^{st})$, $50\%(2^{nd})$ and $51.72\%(3^{rd})$ landed up in preterm labour with or without PROM.

CONCLUSION – There was a very high positive correlation between the incidence of a positive swab culture and preterm events. We found that the best time for a cervicovaginal swab was during the 3rd trimester. Our study had a high negative predictive value i.e. absence of growth were best in predicting that cases would not go in to preterm labour

INTRODUCTION: Preterm labour is defined as the presence of uterine contractions of sufficient frequency and intensity to effect progressive effacement and dilation of the cervix prior to term gestation (between 20-37 weeks). Preterm births occur in approximately 12% of pregnancies and is the leading cause of neonatal mortality today ^(1,2). The exact mechanism of preterm labour is yet unknown but is postulated to be commonly due to decidual haemorrhage (abruption, mechanical factors such as uterine over distension as in multiple gestation or polyhydramnios), cervical incompetence (trauma/cone biopsy), cervical inflammation (due to vaginosis/trichomonas), uterine distortion (Mullerian anomalies, fibroid), maternal inflammation or fever, hormonal changes or utero-placental insufficiency (hypertension, diabetes, drug abuse etc.) ^(1,2).

Intrauterine infections are responsible for 40% cases of preterm labour that may be overt or subclinical whereas the incidence of positive cultures of amniotic fluid during preterm labour varies from 10-40%, the average being 40%.

Premature rupture of membranes (PROM) is the rupture of membranes prior to the onset of labour beyond 37 completed weeks of gestation. Preterm premature rupture of membranes (PPROM) is the rupture prior to 37 weeks of gestation. Early PROM is mostly due to inflammation and/or infection of the membranes. Clinical factors associated with PPROM are low socio-economic status, low BMI, tobacco use, history of previous preterm labour, urinary tract infections, history of vaginal bleed during pregnancy or cerclage or amniocentesis (3).PROM occurs in 10% of pregnancies.

The frequency of infections in preterm births is inversely related to the gestational age ⁽⁴⁾. Endotoxins released by microorganisms and the decidual reaction may stimulate uterine contractions and weaken fetal membranes respectively ⁽⁵⁾. Bacterial vaginosis has been linked to preterm birth raising the risk by a factor of 1.5. Extremes of age ^(4,7), maternal height, weight ⁽⁸⁾ and decrease in interval of pregnancies increase the incidence of preterm births⁽⁹⁾.

Proper intervention with appropriate antibiotics is necessary in those with proven infections. They have two effects i.e. first to increase latency (time till delivery) and secondly to reduce the occurrence of neonatal sepsis, intraventricular haemorrhage and perinatal mortality and chorioamnionitis.

The aim of our study was to study the impact of education, socio-economic status, the effect of intervention in cases with positive cervico-vaginal microbiology on final outcome of pregnancy and to study the appropriate trimester for taking cervico-vaginal swabs in pregnant women.

MATERIALS AND METHODS: Our study was carried out in a community hospital from the year 2008-2010. It was a prospective study involving 100 pregnant women with regular follow up in all three trimesters and they were divided into two groups: culture positive and culture negative according to the swab culture reports. Those with positive reports were treated according to the antibiotic sensitivity test.

The following women were included in the study: -

- 1. Those with known last menstrual periods (LMP)
- 2. Presenting from the 1st trimester
- 3. Having singleton pregnancy

The following women were excluded from the study: -

- 1. Those with demonstrable maternal cause of preterm labour and PROM
- 2. Those who refused consent or follow up or defaulters
- 3. Those with history of liver disease and/or on concurrent intake of hepatotoxic drugs
- 4. Those with suspected or confirmed fetal congenital malformations

Data collection was done during the OPD visits. Detailed history and thorough clinical examination with special consideration to per speculum examination was done. Cervico-vaginal swabs were taken in each trimester after informed consent along with routine ANC investigations. Patients were treated as per swab culture results. Finally outcomes were compared with respect to the trimester in which they were intervened.

RESULTS:

Table 1. prevalence of cervico-vaginal infections in various trimesters

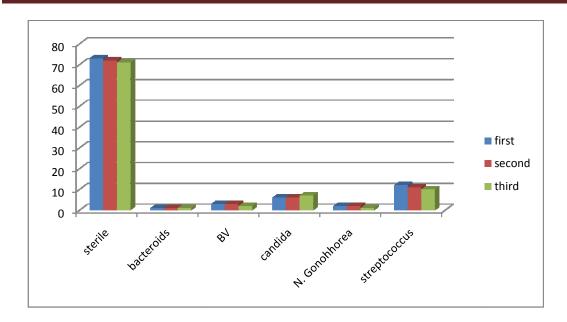


Table 2: Swab results with final outcome in first trimester

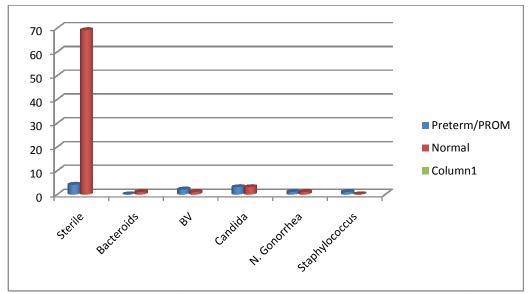


Table 3. Swab results with final outcome in second trimester

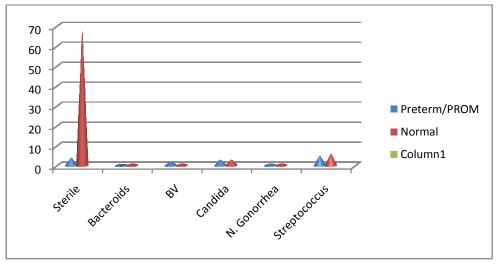


Table 4. Swab results and final outcome in the third trimester

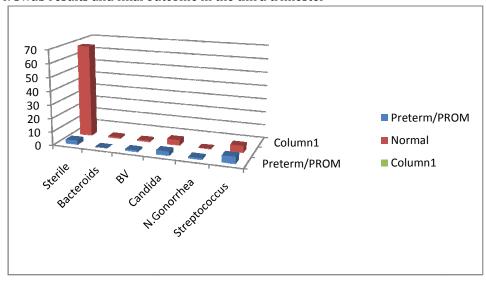


Table 5. Socioeconomic status versus third trimester cervico-vaginal swabs

	Socioeconomic Status		
Swab Results	Average	Below	Total
Positive	13	16	29
Negative	51	20	71
Total	64	36	100

P value - 0.011.

DISCUSSION: Of all the swab positive cases in all three trimesters, streptococcus, candida and staphylococcus were the most commonly prevalent micro-organisms among all the micro-organisms isolated so far in cervico-vaginal cultures in our study.

The mean age of the pregnant women in our study was 24.51 yrs (min – 18 yrs and max – 38 yrs). It has been found that at the age less than 18 yrs and more than 35 yrs, increases the risk of preterm deliveries^(4,7).

Out of total 100, 27 women were swab positive in the first trimester (i.e. 27%) and there was an increasing trend of 1% in swab positivity in subsequent trimesters as a result of introduction of new infections as pregnancy advanced; new infection rate being less in second trimester than third (i.e. 39.28 versus 48.2%).

Of all the organisms, streptococcus (mainly GBS, S. pyogenes), candida and staphylococcus (including coagulase negative) were the most commonly prevalent microorganisms isolated in the cultures in decreasing order of frequency.

Among the swab positive cases in our study, the following percentage landed in preterm labour with or without premature rupture of membranes i.e. 51.58% (first), 50.0% (second), 51.72% (third). The trend of infection rate ultimately causing preterm events in our study is supported by the fact given by Lamont, et al $^{(10)}$ i.e. earlier in pregnancy at which abnormal genital tract colonization is detected, the greater is the risk of late miscarriage and preterm birth.

All culture swab positive cases in our study were treated according to their antibiotic susceptibility and finally the results were analysed. In our study, cure rate when compared with final outcome of pregnancy was found to be best if provided in second trimester amongst all three (i.e. 48.1%, 50%, 48.27% in first, second and third trimester respectively).

Among the 100 women only 10% were formally educated and 11% of total sample size in our study had significant past history. However, neither of the two parameters had any statistical significance on the swab culture results in our study. In a few previous studies by Moreau et al $^{(11)}$ and Philip M. Blumenshine et al $^{(12)}$ mentioned the importance of previous obstetric history as well as the impact of maternal as well as paternal education as a risk factor for preterm labour. The exact mechanism is unclear but it is postulated that it could be due to an unfavourable uterine environment in the former and the lack of awareness and improper nutrition in the latter.

Our study did have a significant P value with respect to socioeconomic status and swab positivity in the third trimester. Out of the 36 women below poverty line (measured by the Kuppuswamy scale), 16 were swab positive i.e.44.44% (as compared to the 13 in the remaining 64 women i.e. 20.31%) i.e. 55.17% of the swab positive women were below poverty line.

48.1%, 82.1% and 65.5% women were symptomatic in the first, second and third trimesters respectively and 77.7%, 75% and 72.4% had demonstrable signs in the respective trimester. Interestingly, both symptoms and signs were significant except in the early trimester. In our study, vaginal discharge in the first trimester was poorly related with swab positivity unless patient complained of persistence of symptoms in the subsequent trimesters.

All culture swab positive women were treated according to their antibiotic susceptibility and results were analyzed. Prior studies have revealed that antibiotic treatment results in significantly greater prolongation of pregnancy as compared with placebo in small studies⁽¹³⁾ but negative results in studies involving a large number of subjects⁽¹⁴⁾. In our study, the cure rate compared to the final outcome was best when provided in the second trimester. Shennan AH et al⁽¹⁵⁾ has stated that unnecessary use of antibiotics with no objective evidence of infection

in spurious preterm labour at late gestations may do more harm than good. Very little difference in the cure rates in subsequent trimesters was seen in our study since intervention had been given to all cases.

Cervicovaginal swab culture has been found to be the most important diagnostic tool to isolate the causative microorganism. Even other studies (16) have revealed its significance in predicting PROM, PPROM and preterm delivery. In fact when compared to a wet mount and cytology methods, it ranks at the top as stated in the International Journal of Tropical Medicine(17-147). Proper culture techniques have been considered the gold standard technique while subjection the selected positive cases to other more accurate tests like real time polymerase chain reaction and optical immunoassay (18-148). In our study, the sensitivity of this method was 77.77% in the first and second trimester and 83.33% in the third trimester. The specificity was 84.14% in the first trimester and 82.92% in the second and third trimester. The negative predictive value ranged from 94-96%. Thus it is safe to say that culture techniques were the best to predict cases that were not going to land up in preterm labour or premature rupture of membranes.

CONCLUSION: Prevalence of cervico-vaginal infections in our study was 27% (first trimester), 28% (second trimester), 29% (third trimester) during the pregnancy. Streptococcus, Candida and Staphylococcus were the leading micro-organisms in various trimesters.

The mean age at pregnancy was 24.5 years. The ratio of the uneducated versus the educated was 9:1 and it did not affect swab culture results significantly. Amongst all the parameters, socio-economic status proved to be influencing swab culture with a significant P value of 0.011.

Persistent vaginal discharge (with or without itching) in subsequent trimesters and bleeding per vaginum were the two most common clinical presentation in women who had positive swab cultures.

The Diagnostic efficacy of cervico-vaginal swabs was $80\% \pm 3\%$ in terms of sensitivity and $83\% \pm 1\%$ in terms of specificity. Absence of growth on the cervicovaginal swabs were the best to predict cases that would not have preterm events as proved by the negative predictive value.

Medical intervention was helpful in reducing the preterm events as demonstrable by conversion of culture positive cases to negative cases in our study. Though a pregnant woman is prone for new infections as pregnancy advances but they need to be treated as soon as infection is diagnosed. So, this study concludes that a pregnant woman should be treated with antibiotics depending on the culture sensitivity results.

Third trimester was found to be best time for taking a cervico-vaginal swab in a pregnant woman. However swab may be taken at any time if a pregnant woman complaint of progressively increasing vaginal discharge or at any time she experiences bleeding per vagina in the present pregnancy, as this was found to be statistically significant in our study.

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