

Screening of preterm labor in Yazd city: transvaginal ultrasound assessment of the length of cervix in the second trimester

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Abstract

Background: Spontaneous preterm labor is one of the common obstetrics problems causing several physical, psychological and economical outcomes. Although due to these outcomes and the efficacy of cares for decreasing them, preterm labor screening is cost-effective and it is still one of the challenging issues in obstetrics.

Objective: In this study preterm labor screening by using cervical transvaginal sonography was evaluated.

Materials and Methods: This observational cohort study was performed in Yazd, Iran. Samples were selected from pregnant women at gestational age of 21-24 weeks who had single live fetus and referred to the obstetrics clinics of two selected hospitals in Yazd. Gestational age was estimated based on the sonography of the first trimester and cervical length measured by transvaginal sonography. Data analysis was done by using t and χ^2 test as well as ANOVA. Statistical significant level was considered as $p < 0.05$.

Results: From 450 participants, 47 cases had preterm labor and 6 cases had positive funneling. Mean age of women with term labor was 26.09 ± 4.13 years and that of women with preterm labor was 26.7 ± 3.51 years ($p = 0.334$). Duration of pregnancy and cervical length significantly differed between women with and without funneling ($p = 0.001$). The sensitivity and specificity of screening based on cervical length of 25mm were 55.5% (50.9-60.1%) and 93.6% (91.2-96%) respectively.

Conclusion: Based on the results of the present study, transvaginal ultrasound assessment of cervical length in low risk women has an acceptable reliability for screening of preterm labor.

Key words: Early detection, Obstetric labor, Ultrasonography, Cervix uteri, Iran.

Introduction

Preterm labor is one of the common obstetrics problems with an annual worldwide incidence rate of 13,000,000 cases (1). It is the major cause of prenatal mortality and is usually associated with recognized maternal and neonatal outcomes (2-7). Moreover, preterm labor imposes psychological and socio-economical pressures on both neonate's parents and the society (8, 9). Spontaneous preterm delivery (SPTD) accounts for approximately 72% of preterm deliveries and several factors can increase the risk of this type of preterm delivery. Some of them are early threatened abortion in the current pregnancy, genetic factors, demographic features, behavior features such as cigarette smoking, low maternal weight gain during pregnancy,

consumption of illicit drugs, pregnancy in extremes of reproductive life, short stature, vitamin C deficiency, prolonged walking and standing, exhausting activities and psycho-physical stresses (6, 10-17). Early diagnosis of high risk pregnancies for preterm labor and its outcomes has significant role in decreasing the incidence and mortality rates (5).

According to the previous studies, the risk scoring system for predicting preterm birth developed by Papiernnik and its modified version by Creasy have not been efficient in the diagnosis of several cases leading to preterm labor (2, 17). There are different techniques for the diagnosis of preterm labor (18-21). Khani *et al* have used β -HCG level in cervicovaginal secretions as an indicator for the risk of preterm labor. They studied 354 high risk pregnant women in 24-28 weeks of gestation and have determined cervicovaginal

β -HCG level of 30mu as cut off point. They asserted that this factor can be beneficial in predicting preterm labor (22). Considering the positive relationship between short cervical length and increased rate of preterm labor incidence, measurement of cervical length can be suggested as an efficient way for predicting preterm labor (3, 4, 21, 23-25).

In comparison to other techniques, in transvaginal ultrasound assessment the quality and details of cervical canal can be observed better due to the closeness of the probe to the cervix (5, 25). Although the cost effectiveness and non-invasiveness of transvaginal sonography have been proved in several studies, in most of them further studies have been recommended (3, 5, 18, 26, 27). For example, Mahshidian *et al* have used cervical sonography parameters for predicting preterm labor in high risk women. In the mentioned study, cervical length, funneling and glands were compared between two groups of women with and without preterm labor. In their observational study, on 200 pregnant women with one of the risk factors of preterm labor they concluded that sensitivity and specificity of cervical length <18mm for preterm labor in gestational age of 35th wk or less are 25% and 99% respectively. The same values for cervical funneling were found to be 50% and 91.8% respectively. Finally, the researchers suggested further studies with greater sample size (21).

Considering the results of the mentioned study and also genetic and environmental differences of societies, evaluation of the results of studies in other countries for their efficacy in our country seems to be necessary (28). Therefore, the present study was performed with the aim of evaluating the efficacy of measuring cervical length and funneling by transvaginal sonography at gestational age of 21-24 wk in screening of preterm labor in Yazd city, Iran.

Materials and methods

This study was an observational prospective cohort project in Yazd city. The permission was obtained from the Vice Chancellor of Treatment and Ethics Committee, Yazd University of Medical Sciences. In the obstetrics clinics of Shahid Sadooghi and Madar Hospitals in Yazd, participants were selected from the aimed

population. Inclusion criteria were having gestational age of 21-24 wk, single pregnancy and live fetus.

Those with special condition disturbing the results like induced preterm labor due to severe preeclampsia, HELLP syndrome, severe IUGR (intra uterine growth restriction), abruption, fetal distress were excluded. Sequential sampling was performed and sample size, based on Carley *et al* study and considering 80% response rate had been determined 450 pregnant women (29). After considering the conditions of women referred to the clinics based on the inclusion criteria, the aims and process of study were explained to them and they were asked to give their consent for participating in the study. It should be mentioned that in studying the inclusion criteria, gestational age was estimated based on the last menstrual period (LMP) and its agreement with the result of sonography in the first trimester.

In the next stage, data related to age, job, educational level, gestational age, history of previous preterm labor, cerclage and curettage obtained in face to face interview and also extracted from the patients' medical files, were recorded in the previously prepared forms. Then participants underwent transvaginal sonography at 21-24 wk (one time) and their cervical length or the presence of funneling was evaluated. Transvaginal sonography was performed after bladder emptying in dorsal lithotomy position and by 7.5 MHZ transducer.

In order to observe sagittal cervical profile, probe was placed in anterior vaginal fornix and for observing internal and external cervical os, it was slightly moved and the distance between the echodensity of external os and V-shaped region of internal os was measured. Cases with the extrusion of fetal membranes into the cervical canal in a Y shape were considered as positive funneling. After recording data and performing transvaginal sonography, follow up for determining the exact time of delivery was started and time of delivery was recorded for all subjects. The study had no cost for participants and the secrecy of their data was considered.

Statistical analysis

Data analysis was done through SPSS11.5 and STATA 8 software packages by using t

and χ^2 tests as well as ANOVA. In order to estimate sensitivity (SN), specificity (SP), positive predictive value (PPV) and negative predictive value (NPV) and their confidence range, SPSS software was used. Statistical significant level was considered as $p < 0.05$.

Results

A total of 450 women from 508 eligible participants were scheduled in this study. Mean age of participants was 26.16 ± 4.07 years with the range of 15-45 years. From all, 53 cases had history of previous curettage and 42 cases reported history of preterm labor. In regard to the educational level, 101 cases were illiterate, 277 cases had high school Diploma or lower and 70 cases had higher degrees. In total, 404 women did not work out, 26 had low physical activity, 16 had difficult jobs and 4 women did not report their jobs. Considering gestational age < 37 wk as a criterion for preterm labor, 47 women had preterm labor and according to the results of transvaginal sonography, 6 cases were positive for funneling.

Mean age of subjects with term labor was 26.09 ± 4.13 years and that of those with spontaneous preterm labor was 26.7 ± 3.51 years that shows no significant difference ($p = 0.334$). Mean cervical length of whole participants was 35.25 ± 6.87 mm and it showed

significant difference ($p = 0.001$) in the two groups of term (36.53 ± 5.81 mm) and preterm (24.31 ± 5.33 mm). Statistical analysis showed significant relationship between the cervical length and duration of pregnancy ($p = 0.0001$). Table I shows duration of pregnancy, the number of preterm labors and age distribution of women based on cervical length classification.

As it is seen, mean cervical length in women with history of preterm labor is 28.76 ± 4.09 mm that shows significant difference as compared to 36.8 ± 6.7 mm in those without history of preterm labor ($p = 0.0001$). According to the results, two groups of term and preterm labor had no significant difference in regard to the educational level ($p = 0.18$), but they had significant difference in regard to job ($p = 0.001$). Mean pregnancy duration and mean cervical length in women with positive funneling were respectively 32 ± 2.5 wk and 15.0 ± 2.1 mm, while the same values for women with negative funneling were respectively 38.8 ± 1.3 wk and 35.5 ± 6.5 mm that is evident of significant difference between the two groups ($p = 0.001$). Table II shows the results of evaluation of SPTD screening based on the cervical length measured through transvaginal sonography in women without history of preterm labor and at the gestational age of 21-24 wk.

Table I. Descriptive statistics of duration of pregnancy, the number of preterm and term labors, and age categories regarding cervical length

Cervical length (mm)	Count	Duration of pregnancy (mean \pm SD) weeks	Number of term labors		Number of preterm labors	
			>20 years old	≤ 20 years old	>20 years old	≤ 20 years old
11- 15	6	33.16 ± 3.6	0	1	0	5
16- 20	10	35.75 ± 1.4	0	2	1	7
21- 25	30	36.98 ± 1.1	4	11	0	15
26- 30	58	38.1 ± 1.08	4	46	1	7
≥ 31	346	39.2 ± 1.1	25	310	0	11

Table II. Evaluation of SPTD screening in women without history of preterm labor and at the gestational age of 21-24 weeks

Cervical length (mm)	Sensitivity (%)		Specificity (%)		PPV (%)		NPV (%)	
	Point	CI	point	CI	point	CI	point	CI
≤ 15	11.1	8.2-14	99.7	98.8-100	75	71-79	93.5	91.1-95.9
≤ 20	16.6	13.2-20	96	94.2-97.8	58	53.5-62.5	94.7	92.5-96.9
≤ 25	55.5	50.9-60.1	93.6	91.2-96	51	46.4-55.6	98.8	97.5-100
≤ 30	94.4	92.2-96.6	83.9	80.4-87.4	30.6	26.4-34.8	99.3	98.4-100

SPTD: Spontaneous preterm delivery.
PPV: Positive predictive Value.

CI: Confidence Interval.
NPV: Negative predictive Value.

Discussion

The results of the present study showed significant relationship of cervical length at gestational age of 21-24wk with duration of pregnancy and preterm labor risk ($p=0.001$). Mean cervical length in participants with preterm labor was estimated to be 24.3mm which is approximately close to the estimated value in Imas *et al*, Berghella *et al* and Heath *et al* studies, but differs with 32mm reported in Farshchian study (24, 30-31). This finding is evident of the importance of cervical lengths of other than 25mm. The difference between the results of our study with Farshchian study can be attributed to the difference in the techniques of measuring cervical length.

In the present study compared to Mahshidian *et al* study, the percentage of SPTD cases was higher (approximately 10% vs. 3.5%), while percentage of positive funneling cases (1.3% approximately) was lower (10). In the mentioned study, participants were high risk women for preterm labor that can explain higher percent of funneling in them, but in regard to the reason of lower incidence of preterm labor in this group, it is necessary to pay attention to the borderline pregnancy durations for determining preterm labor in the two studies. According to the results of other studies mentioned in the introduction, the results of both Mahshidian *et al* study and the present study seem to be logic (21).

Based on the present study, funneling has significant role in decreasing cervical length and increasing the incidence of SPTD ($p=0.001$). This factor showed severe statistical relationship with preterm labor before 35th week of gestation in Mahshidian *et al* study too, but its relationship with preterm labor before 37th week of gestation has been unclear (21). This may be due to higher rate of preterm labor in lower gestational ages in the mentioned study as compared to the present study. Comparison of sample size in the two studies makes the consistency of the results of the present study more acceptable. Considering lower mean cervical length in women with history of SPTD compared to those without this history and significant relationship of cervical length with preterm labor incidence, the significance of difference in mean cervical length between the two groups of with and without history of SPTD

can be well explained. Even though, attention to the disturbing factors such as causes of previous preterm labors can be of a great help in any decision making.

Maternal educational level and age had no effect on cervical length in the two groups of with and without preterm labor. This finding can guaranty the results of the present study in relation to the disturbing effects resulted from these factors. In spite of this, participants with preterm labor had jobs with significantly higher risk. Even though, small number of participants with difficult jobs in the present study prevented the possibility of exact conclusion and this issue can be investigated in future studies.

Although in table II, 5-mm cervical length classification has been presented for more exact evaluation of the results, according to the results of other studies, cervical length of 25mm can be considered as a cut off point for the screening of SPTD. In this level, half of preterm labors can be predicted and approximately 95% of normal individuals can be identified. Comparison of the results of the present study (with cutoff point of 20mm) with the results of Mahshidian *et al* study (with the cutoff point of 18mm) shows high agreement of the two studies (21). According to this, the reliability of SPTD screening based on cervical length is acceptable. Although the results of the present study are in line with the results of previous studies and confirm the efficacy of cervical length measurement in predicting preterm labor, simultaneous presence of affecting factors have not been analyzed statistically and it can be mentioned as one of the shortcomings of this study.

Conclusion

According to the present study, the incidence rate of spontaneous preterm labor before the 37th week of gestation is approximately 10% and it reaches to approximately 3-4% before the 35th week. Cervical length measured in transvaginal sonography has acceptable consistency for screening and early diagnosis of spontaneous preterm deliveries in low risk women. Even so, more comprehensive studies with considering simultaneous effects of risk factors are required for suggesting this screening program into the health care system of the country.

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Conflict of interest

There is no conflict of interest in this article.

References

1. Mokuolu OA, Suleiman B, Adesiyun O, Adeniyi A. Prevalence and determinants of pre-term deliveries in the University of Ilorin Teaching Hospital, Ilorin, Nigeria. *Pediatr Rep* 2010; 2 :e3.
2. Mukherjee GG, Buckshee K. Preterm Labor. New Delhi, INDIA: Jaypee Brothers Medical Publishers (P) Ltd; 1998.
3. Kwasan S, Paisarntuntiwong R, Charoenchainont P. Cervical length measurement by transvaginal sonography in preterm pregnant women for prediction of preterm birth. *J Med Assoc Thai* 2005; 88: 48-55.
4. Honest H, Bachmann LM, Coomarasamy A, Gupta JK, Kleijnen J, Khan KS. Accuracy of cervical transvaginal sonography in predicting preterm birth: a systematic review. *Ultrasound Obstet Gynecol* 2003; 22: 305-322.
5. Greco E, Lange A, Ushakov F, Calvo JR, Nicolaides KH. Prediction of spontaneous preterm delivery from endocervical length at 11-13 weeks. *Prenat Diagn* 2011; 31: 84-89.
6. Golestan M, Fallah R, Akhavan KS. Prediction of preterm delivery in the second trimester. *Iran J Reprod Med* 2008; 6: 205-208.
7. de Carvalho MH, Bittar RE, Brizot Mde L, Bicudo C, Zugaib M. Prediction of preterm delivery in the second trimester. *Obstet Gynecol* 2005; 105: 532-536.
8. Rashidi M, Ghorbani R, Moradan S, Tavan P. [Comparison of Administration of Magnesium Sulfate and Nifedipine in Prevention of Preterm Delivery]. *Iran J Obstet Gynecol Infertil* 2010; 13: 1-6. (In Persian)
9. Lotfalizadeh M, Mohammadzadeh A, Kamandi S, Bagheri S. [prevalence and risk factor of preterm labor in Emam Reza Hospital (1381-1382)]. *Iran J Obstet Gynecol Infertil* 2005; 8: 93-100. (In Persian)
10. Dikamba MN. Determinants of preterm delivery and low birth weight in five rural health zones of Equateur Province, Democratic Republic of Congo. New Orleans: Tulane University; 2010.
11. Reagan PB, Salsberry PJ. Race and ethnic differences in determinants of preterm birth in the USA: broadening the social context. *Soc Sci Med* 2005; 60: 2217-2228.
12. Robinson J, Norwitz E, Lockwood Ch, Barss V. Risk factors for preterm labor and delivery. Available at: www.uptodate.com/contents/risk-factors-for-preterm-labor-and-delivery.
13. Lawn JE, Gravett MG, Nunes TM, Rubens CE, Stanton C. Global report on preterm birth and stillbirth (1 of 7): definitions, description of the burden and opportunities to improve data. *BMC Pregnancy Childbirth* 2010; 10: S1.
14. Silveira MF, Victora CG, Barros AJ, Santos IS, Matijasevich A, Barros FC. Determinants of preterm birth: Pelotas, Rio Grande do Sul State, Brazil, 2004 birth cohort. *Escola Nacional de Saude Publica* 2010; 26: 185-194.
15. Cunningham F, Leveno J, Bloom L, Hauth J, Rouse D, Spong C. Preterm Birth William's Obstetrics. 23th Ed. United States: McGraw-Hill; 2010.
16. Khalajinia Z, Jandaghi G. Maternal risk factors associated with preterm delivery in Qom province of Iran in 2008. *Sci Res Essays* 2012; 7: 51-54.
17. Khakbazan Z, Granmayeh M, Taghizadeh J, Haghani H. [Association of occupational factors and preterm labor]. *Hayat Research Journal* 2007; 13: 5-14. (In Persian)
18. Iams JD. Prediction and early detection of preterm labor. *Obstet Gynecol* 2003; 10: 402-412.
19. Elder G, Lamont R, Romero R. chapter 3. Preterm labor. Edinburgh: Churchill Livingstone; 1997.
20. Roghaei M, Sabahi R, Ghasemi M. [The diagnostic value of dilatation and interleukin 6 and 8 in cervicovaginal fluid to prediction of preterm labor]. *J Isfahan Med School* 2009; 27: 127-33. (In Persian)
21. Mahshidian M, Marsosi V, Ziaee S, Asghari JM. [Sonographic cervical parameters in predicting spontaneous birth in high-risk pregnant women]. *Tehran Univ Med J* 2011; 68: 583-589. (In Persian)
22. Khani S, Khalilian A. [Preterm delivery prediction by the measurement of cervicovaginal Beta human chorionic Gonadotropin (B-HCG)]. *Razi J Med Sci* 2004; 11: 749-756. (In Persian)
23. Palma-Dias RS, Fonseca MM, Stein NR, Schmidt AP, Magalhaes JA. Relation of cervical length at 22-24 weeks of gestation to demographic characteristics and obstetric history. *Braz J Med Biol Res* 2004; 37: 737-744.
24. Heath VC, Southall TR, Souka AP, Elisseou A, Nicolaides KH. Cervical length at 23 weeks of gestation: prediction of spontaneous preterm delivery. *Ultrasound Obstet Gynecol* 1998; 12: 312-317.
25. Mella MT, Berghella V. Prediction of preterm birth: cervical sonography. *Semin Perinatol* 2009; 33: 317-324.
26. Conde-Agudelo A, Romero R, Hassan SS, Yeo L. Transvaginal sonographic cervical length for the prediction of spontaneous preterm birth in twin pregnancies: a systematic review and metaanalysis. *Am J Obstet Gynecol* 2010; 203: 128 e1-12.
27. Mukherji J, Anant M, Ghosh S, Bhattacharyya SK, Hazra A, Kamilya GS. Normative data of cervical length in singleton pregnancy in women attending a tertiary care hospital in eastern India. *Indian J Med Res* 2011; 133: 492-496.
28. Behrman RE, Butler AS. Prematurity at Birth: Determinants, Consequences, and Geographic Variation. Preterm Birth: Causes, Consequences,

- and Prevention. Washington (DC), USA: National Academies Press; 2007.
29. Carley S, Dosman S, Jones SR, Harrison M. Simple nomograms to calculate sample size in diagnostic studies. *EMJ* 2005; 22: 180-181.
30. Berghella V, Bega G, Tolosa JE, Berghella M. Ultrasound assessment of the cervix. *Clin Obstet Gynecol* 2003; 46: 947-962.
31. Iams JD, Goldenberg RL, Meis PJ, Mercer BM, Moawad A, Das A, et al. The length of the cervix and the risk of spontaneous premature delivery. National Institute of Child Health and Human Development Maternal Fetal Medicine Unit Network. *New Eng J Med* 1996; 334: 567-572.