# RESEARCH NOTE Open Access

# Does short inter-pregnancy interval predicts the risk of preterm birth in Northern Ethiopia?



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# **Abstract**

**Objective:** The study aimed to assess the effect of inter pregnancy interval on preterm birth in Northern Ethiopia: prospective cohort study.

**Result:** This study showed that, total incidence of premature birth was 10.4%. Among mothers with short inter pregnancy interval the incidence of preterm birth was 39 (25.9%). Whereas, among mothers who had recommended inter pregnancy interval was 9 (2.9%). Short inter-pregnancy interval [adjusted hazard ratio (AHR): 6.85, 95% confidence interval (CI) 3.07–15.31], antenatal care (ANC) visit 1–3 times (AHR: 2.24, 95% CI 1.04–4.85), complication during pregnancy (AHR: 3.16, 95% CI 1.58–6.33) and birth defect (AHR: 8.01, 95% CI 2.56–25.07) were predictors of premature birth

Keywords: Inter pregnancy interval, Incidence, Preterm birth, Cohort study, Northern Ethiopia

#### Introduction

Preterm birth (PTB) is defined as a delivery which occurs at less than 37 completed weeks of gestation. It is classified as extremely preterm (<28 weeks), very preterm (28 to <32 weeks) and moderate to late preterm (32 to <37 weeks) [1]. Globally, 15 million babies are born too early every year, which are more than 1 in 10 babies. Approximately 1 million children die each year due to complications of preterm birth. Moreover, many survivors face a lifetime of disability, learning incapacities, visual and hearing problems [2].

Systemic review done in 2007 showed that, 9.6% of all births were preterm, which translates to about 12.9 million births definable as preterm. Approximately 85% of this burden was concentrated in Africa and Asia, where 10.9 million births were preterm [3]. Based on UNICEF report among the neonates delivered in Ethiopia 10% of them are preterm [4]. Different studies conducted in Ethiopia revealed that the incidence of preterm birth

ranges from 4.4 to 25.9% [5–9]. And it also accounts 28% of newborn deaths in Ethiopia [10].

Other studies indicates that history of abortion, still birth, preterm birth, Premature rupture of fetal membrane, history of bleeding during the pregnancy, pregnancy induced hypertension, HIV infection, substance intake, history of low birth weight and short inter-pregnancy interval (IPI) were the factors leading to preterm birth [4–8].

Furthermore, women with short interval between pregnancies are at increased risk of preterm birth. However, whether this association is confounded by other risk factors, including various aspects of socioeconomic status, ethnicity, demographics, and lifestyle is unclear [11]. Therefore this study aimed to assess the effect of interpregnancy interval and preterm birth in Ethiopia. This will help health providers to have adequate knowledge and intervene early in the antenatal care visit and also might have an input to programmers and policy makers.

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#### Main text

# Methods and material

# Study setting, design and sample size

Health facility based prospective cohort study design was conducted in Tigray Region, Northern Ethiopia, located



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at distance of 1087 km from Addis Ababa, the capital city of Ethiopia. Source population was child bearing mothers who are attending delivery in north western zone health facilities of Tigray.

The sample size of 480 pregnant women was calculated using Epi-info-Calc. statistical software with the assumption of 95% CI, power of 90%, ratio of unexposed to exposed was 2:1 and percentage of exposed among preterm birth of 8.5% (14), odds ratio of 2.7, Loss to follow-up rate was estimated to be 10%.

Four hospitals and eight health centers were selected randomly from North West zone health facilities. Study participants were proportionally allocated to each health facilities based on eligible pregnant mothers for ANC. Lastly, participants were included through systematic sampling method until the desired sample achieved.

# Data collection procedure and quality assurance

Data was collected using standardized, structured and face to face interviewer questionnaire and card review. Data quality was managed by trained three Bachelors degree of science holders fluent in the local language (Tigrigna). Pre-test, daily supervision, spot checking and reviewing completed questionnaire was conducted.

# **Explanatory measurements**

Gestational age: was estimated as the interval in completed weeks from the last normal menstrual period (LNMP) to the child's date of birth. When there are extra days it is counted to the near lowest gestational age but if the mother didn't remember or recall her LNMP; gestational age was estimated from ultra sound result through reviewing the mothers chart.

Inter-pregnancy interval: was documented as the time interval between delivery of the first pregnancy and the conception of the subsequent pregnancy.

Short inter-pregnancy interval: mothers with interpregnancy interval from 0 to 24 months.

Recommended inter-pregnancy interval: mothers with inter-pregnancy interval from 24 to 36 months or longer.

#### **Outcome measures**

Censor: mothers with normal weight, low birth weight, still birth, and congenital anomaly.

Event: preterm birth.

# Statistical analysis

Data was entered into Epi Data 3.5.1 statistical software and analyzed using SPSS version  $20.0^{\text{TM}}$  package. The results were presented in tables and texts using descriptive statistics such as mean, standard deviation, frequency and percentages.

The relationship between the event and explanatory variables was analyzed using Cox bivariate proportional regression model. Before fitting the covariate into the model proportional hazard assumption was checked using Log (-Log) S (t) plots.

In order to identify independent predictors a multivariate Cox-proportional adjusted model was fitted with those variables p-value  $\leq$  0.25 in bivariate cox proportional regression analysis. Crude and adjusted hazard ratios with their 95% Confidence interval (CI) were estimated and p-value less than 0.05 were used to declare the presence of significant association.

#### Results

#### Socio demographic characteristics of participants

Four hundred eighty pregnant women were followed with 20 (4.17%) lost in follow-up until delivery. The mean age of mothers was 30 (SD  $\pm$  5.92) ranging from 15 to a maximum of 48 years and above half 260 (56.5%) were between 25 and 34 years. Almost all or 450(97.8%) of the participants were married. Two-fifth 164 (35.7%) of the study participants had no formal education. About maternal occupation four out of ten participants 205(44.6%) were housewife and above half 273 (59.3%) of the participants come from Urban (Table 1).

Table 1 Socio-demographic characteristic of pregnant mothers in Tigray, Northern Ethiopia, 2018 (N = 460)

| Categories             | Frequency | Percentage |  |
|------------------------|-----------|------------|--|
| Age in completed years |           |            |  |
| 15-24                  | 75        | 16.3       |  |
| 25-34                  | 260       | 56.5       |  |
| 35–48                  | 125       | 27.2       |  |
| Marital status         |           |            |  |
| Married                | 450       | 97.8       |  |
| Unmarried              | 4         | 0.9        |  |
| Divorced/widowed       | 6         | 1.3        |  |
| Educational level      |           |            |  |
| None                   | 164       | 35.7       |  |
| Primary                | 82        | 17.8       |  |
| Secondary              | 135       | 29.3       |  |
| College and above      | 79        | 17.2       |  |
| Occupation             |           |            |  |
| House wife             | 205       | 44.6       |  |
| Farmer                 | 125       | 27.2       |  |
| Employed               | 130       | 28.3       |  |
| Residence              |           |            |  |
| Rural                  | 187       | 40.7       |  |
| Urban                  | 273       | 59.3       |  |

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# Past-obstetrics and current pregnancy characteristics

Among the cohort women two-third 308 (67.0%) of them had recommended inter pregnancy interval and three-fourth 346 (75.2%) of them had < 4 pregnancy with two-third 312 (67.8%) of them were multiparous. One out of ten (n=418) women experience perinatal death in their preceding pregnancy. Around three-fourth 332 (72.2%) of the participants had planned pregnancy. Majority 445 (96.7%) of the pregnant women had at least one ANC checkup and four hundred thirteen (89.8%) of the pregnant women had maternal obstetrical complication during their current pregnancy. Eight out of ten (n = 375) of the study participants had hemoglobin level above 11 g/dl and eighty percent (n = 374) participants had delivered through spontaneous vaginal delivery for their current baby with a majority, 281 (61.1%) of them gave male baby by sex (Table 2).

## Incidence and predictors of premature birth

The total incidence of premature birth in this study was 10.4%. Incidence among mothers exposed for short inter pregnancy interval was 39 (25.9%) and this indicates that around 3 out of 10 mothers who are exposed for short inter pregnancy interval have premature baby. But the incidence among mothers with recommended inter-pregnancy interval was 9 (2.9%).

The predictors for premature birth were mothers with short inter-pregnancy interval had around seven times (AHR: 6.85, 95% CI 3.07–15.31) hazard of bearing premature birth than those with recommended inter-pregnancy interval; also mothers with 1-3 times ANC visit had two times (AHR: 2.24, 95% CI 1.04-4.85) likelihood to have premature birth than those who have greater than or equal to 4 visits and pregnant mother who had any problem during her current pregnancy had 3 times (AHR: 3.16, 95% CI 1.58-6.33) more hazard to give premature birth than their counter parts. And those pregnant who gave birth a newborn with birth defect was eight times (AHR: 8.01, 95% CI 2.56–25.07) more hazardous than those with normal birth. However, maternal residence, planned pregnancy, history of perinatal death and maternal hemoglobin level were significant at bi-variable but in significant in multivariable regression analysis (Table 3).

#### Discussion

This study showed the incidence of spontaneous premature birth was decreasing from the previous studies done in Ethiopia. Short inter-pregnancy interval, number of ANC visit, newborn birth defect and presence of complication during pregnancy were the predictors of preterm birth.

Table 2 Past-obstetrics and current pregnancy characteristics of pregnant mothers in Tigray, Northern Ethiopia, 2018 (N=460)

| (N=460)  |           |            |  |  |  |
|--|-----------|------------|--|--|--|
| Category   | Frequency | Percentage |  |  |  |
| Inter pregnancy interval                             |           |            |  |  |  |
| Recommended  | 308       | 67.0       |  |  |  |
| Short  | 152       | 33.0       |  |  |  |
| Gravidity  |           |            |  |  |  |
| ≤4 pregnancy   | 346       | 75.2       |  |  |  |
| ≥ 5 pregnancy  | 114       | 24.8       |  |  |  |
| Parity   |           |            |  |  |  |
| Primipara  | 148       | 32.2       |  |  |  |
| Multipara  | 312       | 67.8       |  |  |  |
| History of perinatal death                           |           |            |  |  |  |
| No   | 418       | 90.9       |  |  |  |
| Yes  | 42        | 9.1        |  |  |  |
| Planned pregnancy                                    |           |            |  |  |  |
| No   | 128       | 27.8       |  |  |  |
| Yes  | 332       | 72.2       |  |  |  |
| ANC follow up  |           |            |  |  |  |
| No   | 15        | 3.3        |  |  |  |
| Yes  | 445       | 96.7       |  |  |  |
| Number of ANC visits                                 |           |            |  |  |  |
| ≥ 4 times  | 225       | 57.3       |  |  |  |
| 1–3 times  | 190       | 42.7       |  |  |  |
| Initiation of ANC visit                              |           |            |  |  |  |
| Within 16 weeks                                      | 22        | 4.9        |  |  |  |
| 24–28 weeks  | 244       | 54.8       |  |  |  |
| 28–32 weeks  | 169       | 38.0       |  |  |  |
| 34–36 weeks  | 10        | 2.2        |  |  |  |
| Current pregnancy maternal complication <sup>a</sup> |           |            |  |  |  |
| No   | 413       | 89.8       |  |  |  |
| Yes  | 47        | 10.2       |  |  |  |
| Maternal hemoglobin level (g/dl)                     |           |            |  |  |  |
| ≥11  | 375       | 81.5       |  |  |  |
| <11  | 85        | 18.5       |  |  |  |
| Maternal Rh factor                                   |           |            |  |  |  |
| Positive   | 449       | 97.6       |  |  |  |
| Negative   | 11        | 2.4        |  |  |  |
| Maternal and fetal intra partum complication b       |           |            |  |  |  |
| No   | 414       | 90.0       |  |  |  |
| Yes  | 46        | 10.0       |  |  |  |
| Mode of delivery                                     | 40        | 10.0       |  |  |  |
| Cesarean section                                     | 45        | 9.8        |  |  |  |
| Instrumental   | 18        | 3.9        |  |  |  |
| Induction/augmentation                               | 23        | 5.9        |  |  |  |
| Spontaneous vaginal delivery                         | 23<br>374 | 81.3       |  |  |  |
| Sex of newborn                                       | J/4       | 01.0       |  |  |  |
| Male Sex of Newborn                                  | 281       | 61.1       |  |  |  |
|  |           |            |  |  |  |
| Female   | 179       | 38.9       |  |  |  |

<sup>&</sup>lt;sup>a</sup> Current pregnancy maternal problem: APH, preeclampsia, anemia, uterine rupture/scar dehiscence

<sup>&</sup>lt;sup>b</sup> Maternal and fetal Intra partum complication: PPH, obstructed labor, prolonged labor, uterine rupture/scar dehiscence

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Table 3 Bi-variable and multivariable cox-regression analysis for predictors of premature birth, Northern Ethiopia, 2018 (N = 460)

| Characteristics            | N         | Outcome n (%) |            | Hazard ratio, 95% CI | Hazard ratio, 95% CI |  |
|----------------------------|-----------|---------------|------------|----------------------|----------------------|--|
|                            |           | Event         | Censored   | Crude                | Adjusted             |  |
| IPI                        |           |               |            |                      |                      |  |
| Short                      | 152       | 39 (25.9)     | 113 (74.1) | 9.5 (4.6–19.6)       | 6.85 (3.07–15.31)    |  |
| Recommended                | 308       | 9 (2.9)       | 299 (97.1) | 1                    |                      |  |
| Residence                  |           |               |            |                      |                      |  |
| Rural                      | 187       | 30 (16)       | 157 (84)   | 2.5 (1.37-4.41)      | 1.2 (0.61-2.35)      |  |
| Urban                      | 273       | 18 (6.6)      | 255 (93.4) | 1                    |                      |  |
| Planned pregnancy          |           |               |            |                      |                      |  |
| No                         | 128       | 22 (17.2)     | 106 (82.8) | 2.26 (1.28-3.99)     | 1.11 (0.57-2.13)     |  |
| Yes                        | 332       | 26 (7.8)      | 306 (92.2) | 1                    |                      |  |
| Number of ANC visits       |           |               |            |                      |                      |  |
| ≥ 4                        | 255       | 11 (4.3)      | 244 (95.7) | 1                    |                      |  |
| 1-3                        | 190       | 31 (16.3)     | 159 (83.7) | 4.0 (2.01-7.97)      | 2.24 (1.04-4.85)     |  |
| Complication in current p  | oregnancy |               |            |                      |                      |  |
| No                         | 413       | 33 (8.0)      | 380 (92)   | 1                    |                      |  |
| Yes                        | 47        | 15 (31.9)     | 32 (68.1)  | 4.78 (2.59-8.79)     | 3.16 (1.58–6.33)     |  |
| History of perinatal death | ١         |               |            |                      |                      |  |
| No                         | 418       | 40 (9.6)      | 378 (91.4) | 2.16 (1.01-4.61)     | 0.85 (0.36-2.03)     |  |
| Yes                        | 42        | 8 (19)        | 34 (81)    | 1                    |                      |  |
| Newborn birth defect       |           |               |            |                      |                      |  |
| No                         | 452       | 43 (9.5)      | 409 (90.5) | 1                    |                      |  |
| Yes                        | 8         | 5 (62.5)      | 3 (37.5)   | 12.97 (5.11–32.87)   | 8.01 (2.56–25.07)    |  |
| Mother hemoglobin leve     | el (g/dl) |               |            |                      |                      |  |
| ≥11                        | 375       | 32 (8.5)      | 343 (91.5) | 2.33 (1.28-4.25)     | 0.91 (0.46-1.79)     |  |
| <11                        | 85        | 16 (18.8)     | 69 (81.2)  | 1                    |                      |  |
| Maternal age (years)       |           |               |            |                      |                      |  |
| 15-24                      | 75        | 9 (12)        | 66 (88)    | 1.68 (0.67-4.23)     |                      |  |
| 25-34                      | 260       | 30 (11.5)     | 230 (88.5) | 1.64 (0.78-3.46)     |                      |  |
| 35–48                      | 125       | 9 (7.2)       | 116 (92.8) | 1                    |                      |  |

The finding of this study revealed that incidence of premature birth was 10.4% which is almost similar with the study done in Amhara Regional state, Ethiopia in which the preterm birth was 11.6% [8]. However, this result was higher than the study done in Mexico (7.4%) [11] and Southern India (5.4%) [12]. This inconsistency of the finding might be due to the socioeconomic difference and awareness on family planning use. And this result was again lower than the studies done in different parts of Ethiopia such as Addis Ababa (16.15%), Gondar (14.3%) and Jimma (25.9%) [5-7]. This dissimilarity could be due to the success of different programs introduced by federal Ministry of Health to improve health service quality delivered to pregnant women including pre-pregnancy and pregnancy health care in the last decades.

Other result indicated that, short inter-pregnancy interval increases the occurrence of premature birth,

which is similar with the study done in Ethiopia [7], Pakistan [13]. The result from Pakistan indicated that mothers with inter-pregnancy interval less than a year were at a higher frequency of preterm birth as compared to mothers with recommended inter-pregnancy interval. The reason could be short inter-pregnancy interval mothers cannot recover from the biological stresses imposed by the preceding pregnancy resulting in diminution of macronutrient supplementation in maternal body, folate depletion, cervical insufficiency, vertical transmission of infections, incomplete healing of uterine scar and abnormal remodeling of endometrial blood vessels, anemia and increasing the risks of certain other factors achieving pregnancy outcomes [14–16]. So promoting and encouraging the recommended inter-pregnancy interval among couples will reduce the incidence of preterm birth.

In addition, number of ANC visit were another predictor of incidence of preterm birth which was also

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determined in the studies done in Debremarkos and Gondar Ethiopia [6, 8], Ghana [17] and China [18]. This could be due to the importance of recommended ANC follow up and proper visit on detection and early treatment of the problem or any complications. When pregnant mothers come to health facility with any problem or complication that can lead to preterm birth, she might get early treatment so as to prevent the occurrences of preterm birth and other related adverse obstetric outcomes. Thus, mothers should have to follow the recommended ANC visits.

The other predictor in this study was presence of complication during pregnancy (APH, preeclampsia and anemia) which is similar with studies done in Ethiopia [5–8], Ghana [14], China [15], Mexico [11] and southern India [12]. Explicitly, hypertension during pregnancy negatively impacts placental blood flow and leads to poor fetal growth and obstetric emergencies, which increase the risk of having a preterm birth. Therefore early identification, detection and management are crucial for pregnant mothers during their recommended ANC visits.

Having birth defect was the other predictor of preterm birth which is comparable with the study done in Ethiopia [7], New-York United States [19]. The causes of most birth defects and the reasons why these birth defects might contribute to being preterm birth remain unknown. While it is likely that the most common of these defects result from the interface of genetic and environmental risk factors, the identification of specific modifiable risk factors continues to be an important research and public health priority. The possible pathways that could explain the association between preterm birth and birth defects were have a shared risk factors (e.g., maternal smoking, obesity) to cause both birth defects and preterm birth. On the other side, a particular risk factor (e.g., valproic acid, insufficient peri-conceptional folic acid) could lead to a spina bifida, and the presence of that birth defect could then be part of the underlying mechanism causing in a preterm delivery [20]. This suggests that clinical interventions aimed at preventing birth defects may have added benefits in averting the incidences of preterm birth and low birth weight.

#### Conclusion

This study indicates that incidence of premature birth is decreasing when compared with studies done in other parts of Ethiopia. Our findings, in conjunction with those of other studies, strongly suggest that a short inter pregnancy interval is a causal factor for preterm birth. Number of ANC visit, complication during pregnancy and being birth defect are also the predictors for premature birth.

Therefore, the evaluation of the outcomes of primary prevention programs based on this putative causal factor would resolve the matter and health professionals should target to improve maternal education and counseling on recommended ANC visits and the adverse effect of shorter IPIs on subsequent birth, better access to contraception to increase intervals between births and therefore reduce the risk of preterm birth.

Enhancing social marketing approaches emphasized the need to seek early and focused antenatal care to those non-booked mothers is also mandatory. Better management of obstetric complications and research to elucidate the mechanisms by which they cause preterm birth, offers a practical approach of reducing the high preterm birth rates.

# Limitation of the study

There are some limitations on this study. There was lost to follow-up of participants and it was not possible to enroll mothers who had difficult to estimate their gestational age that is unknown last normal menstrual period (LNMP) and missed the ultra sound result in her chart.

#### **Abbreviations**

ANC: antenatal care; IPI: inter pregnancy interval; PTB: preterm birth.

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#### Authors' contributions

BH and MB: conceptualization, designing and writing—original draft. MWA: performed statistical analysis and review the manuscript. HGW: conception, designing and reviewing the manuscript. All authors read and approved the final manuscript

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#### Availability of data and materials

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

#### Ethics approval and consent to participate

The study was approved by Mekelle University Institutional Review Board (ERC1287/2017). Written informed consent was obtained from each study participant and for those participants under the age of 18 years written informed consent was received from their legal guardians.

#### Consent for publication

Not applicable.

#### Competing interests

The authors declare that they have no competing interests.

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