Predictors of mortality among newborns admitted with perinatal asphyxia at public hospitals in Ethiopia: A Prospective cohort study

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Abstract

Introduction:

Perinatal asphyxia is a fetus or the newborn’s health problem caused; due to altered breathing or inadequate inhalation and exhalation resulting in reduced oxygen perfusion to certain body tissues and organs. Irrespective of the increased progress in health care towards newborns and implementations in reductions in under-five, infant, and neonatal mortality in the past ten years, perinatal asphyxia remains as the most common severe newborn health challenge causing a high number of mortalities and morbidity and is a major public issue.

Methods

A prospective cohort longitudinal study was implemented among the predetermined 573 samples of newborns admitted with Perinatal asphyxia at public hospitals in Southern Ethiopia from 1st March 2018 to 28th February 2020. Data entry was conducted using Epi data version 3.02 statistical software and exported to SPSS Version 25 statistical software for analysis. The perinatal survival time was determined using Kaplan Meier survival curve together with a log-rank test. Variables that had a P-values less than 0.05 in the multivariable cox proportional hazard model were declared as statistically significant predictors of mortality.

Results

The cumulative proportion of survival among the newborns admitted with perinatal asphyxia was 95.21% (95%CI:91.00,97.48), 92.82% (95%CI:87.95,95.77), 92.02%(95%CI:86.84,95.22) and 90.78% (95%CI:84.82,94.48) at the end of first, second, third and fourth follow-up days respectively with the overall mean survival time of 6.55(95%CI:6.33,6.77). Cord prolapse (AHR:6.5;95%CI:1.18,36.01), history of PIH (AHR:25.4;95%CI:3.68,175.0), maternal iron deficiency anemia (AHR:5.9;95%CI:1.19,29.5) and having convulsion of the newborn (AHR:10.23;95%CI:2.24,46.54) were statistically significant in multivariable cox proportional hazard model.

Conclusion

The risk of death among newborns with perinatal asphyxia was high during the early follow-up periods after admission to the hospital and the mortality risk decreased at the later follow up periods and cord prolapse, history of PIH, maternal history of iron deficiency anemia and newborns history of convulsion were the independent predictors of mortality.

Introduction
Perinatal asphyxia is a complicated newborn health problem and applies a high contribution to the increased proportion of newborn mortality (1). It is a leading cause of morbidity and mortality in newborn babies globally, with higher case fatality rates and consequent complications in developing countries due to poor health facilities (2, 3). Globally, around 2,500,000 child deaths were reported in the early 28 days of age (neonatal age). These accounts for nearly 47% of under-five mortality and 54% of all under-five deaths occur during this age in developing countries (4).

However, greater than 2/3rd of newborns can be saved through established maternal and newborn health intervention programs. Though, most of the observed deaths have occurred at home delivered newborns (5). Nearly 3.6 million (3%) of all infants suffer from a certain level of perinatal asphyxia. Among this 840,000 (23%) will die and approximately a similar proportion of newborns develop life-threatening health problems in developing countries (6, 7).

Globally, around 25% of all newborn mortality is caused by perinatal asphyxia (8). In Ethiopia, in the year 2015, perinatal asphyxia contributed to 31.6% of newborn deaths, followed by prematurity and neonatal sepsis, which accounted for 21.8% and 18.5% respectively (9).

Irrespective of the increased advancements in perinatal care and implementations in reductions of under-five, infant and neonatal mortality in the past decades (4, 10, 11), perinatal asphyxia remains a severe newborn health problem. This leads to a high number of mortality and morbidity and is a major common public health issue, commonly in developing countries like Ethiopia (12).

Even though Ethiopia reached its child mortality reduction goal 2 years earlier, the neonatal mortality rate remained high. One of the major causes of newborn deaths was intrapartum-related complications of which birth asphyxia accounts for 25% (13, 14).

Moreover, a very limited number of studies were conducted in Ethiopia to identify information for intervention regarding the death due to perinatal asphyxia. Therefore; this study was planned to estimate the time to death and its predictors among newborns with perinatal asphyxia at governmental hospitals in Southern Ethiopia.

**Methods**

**Study design, setting, period and populations**

A prospective cohort longitudinal study was employed at Sawla General Hospital, Arba Minch General Hospital and Chencha district Hospital from first of March 2018 to 28th of February 2020. Among those hospitals, over four thousand newborns were delivered per year and more than 612 newborns were admitted to the neonatal intensive care unit (NICU) at each hospital.\(^{15}\)

Follow up was started at diagnosis of perinatal asphyxia immediately after birth for those delivered at the hospital and at admission to the hospital for those delivered from 1st March 2018 and the follow-up...
period was closed on 28th February 2020. Follow-up was initiated immediately at the diagnosis of PNA and followed until seventh day of life. The follow-up was closed if the newborn was died, discharge with recovery, lost to follow-up from treatment, transferred to another institution and follow-up time ended without the event happening.

In this study, a newborn that withdrew treatment, discharged with recovery, transferred to another institution, and who did not yet develop the event at the end of the follow-up period was operationally defined as Censored. Sample size was estimated by Open Epi 3.02 statistical software using double population proportion formula in considering the assumptions; 95% CI, 80% power, exposed to unexposed ratio: 1, percent of unexposed with outcome (Not having history of premature rupture of membrane (PROM)): 50%, percent of exposed with outcome (Having history of PROM): 62%, AHR: 1.67 and considering 10% for non-response, the sample size became 573. Sample size was allocated to each hospital proportionally based on the number of the admitted cases and consecutive sampling method was applied (Figure 1).

**Study variables**

The dependent variable was time to perinatal mortality and the independent variables were classified as socio-demographic factors (sex of the newborn, maternal age, marital status, a religion of the mother, maternal educational status, maternal occupational status, family size, place of residence, distance between home and hospital and estimated monthly income), obstetrics related characteristics (number of antenatal care (ANC) visits, gravidity, parity, number of pervaginal examinations, history of meconium-stained amniotic uid, the onset of labor, history of antepartum hemorrhage, history of obstructed labor, history of premature rupture of membrane, history of prolonged rupture of membrane, cord prolapse, presentation of the fetus, mode of delivery and gestational age), newborn related factors (cry immediately at birth, history of convulsion or spasm and birth weight) and maternal medical related characteristics (history of PIH, maternal iron deficiency anemia, maternal diabetes mellitus, and maternal HIV status).

**Operational definitions**

Perinatal Asphyxia: is a diagnosis when the newborn’s fifth minute APGAR score less than 7 OR complete absent respiratory effort immediately at birth.

Maternal Anemia: The hemoglobin level of a pregnant woman or early delivery mother less than 11gm/dl.

Premature rupture of the membrane: a rupture (breaking open) of the membranes (amniotic sac) before labor begins.

Prolonged rupture of membrane: a rupture of membranes lasting longer than 18-24 hours (i.e., between the time of rupture and time of delivery).

Convulsion: newborn who experience an episode of rigidity and uncontrolled jerky motions that generally last a minute or two along with altered consciousness.
**Data collection procedure, quality control and analysis**

Structured checklist was used to collect the data. Data extraction tool was carefully designed to improve data quality. In addition, both data collectors and supervisors were trained. Pretest was conducted (5% of the population). Sensitivity analysis was conducted. The maternal hemoglobin test results were obtained from a laboratory report which was prepared for this research purpose. The hemoglobin level was adjusted for altitude according to criteria set by WHO (World health organization).

Epi Data version 3.02 was used to enter the data, code the data, edit the data and clean the data. Finally, the data entered into Epi Data were exported to SPSS version 25 for statistical analysis. The Kaplan Meier survival curve, together with a log-rank test, was used to estimate the survival time and the time which had higher risk of death. Variables that had a p-value <0.05 in bivariate analysis were considered as candidates for multivariable analysis and variables which had a p-value <0.05 in multivariable cox proportional hazard model were considered as statistically significant.

**Ethical consideration**

Ethical clearance was obtained from Arba Minch University, college of medicine and health sciences ethical review board. All participants provided an informed consent. Mothers were informed about the objective and significance of the study prior to the data collection. Appropriate measures were applied to ensure the confidentiality of the data.

**Results**

**Socio-demographic characteristics**

In this study a total of 573 newborns were involved, of which 351 (61.3%) of them were males. In considering maternal age maximum of the mothers (70.7%) were categorized as under 20–34 years of age and the smallest amount (13.6%) were mothers having age less than 20 years old. Regarding the marital status and religious status of the mothers, 51 (8.9%) of the mothers were never married and the maximum number of the mothers (43.5%) were orthodox in their religion followed by protestants (31.9%).

Equal numbers of mothers were both unable to read and write and college and above, each accounted for 11% of the whole mothers. Among the mothers of the newborns, 273 (47.6%), 192 (33.5%), and 108 (18.8%) of the mothers had less than four, four to six, and greater than six family sizes. More than three fourths (76.4%) of the mothers were urban residents and around 93 (16.2%) of the mothers had more than 50km distance between their home and the hospital. Among the dead newborns, 33.3% of mothers were urban residents and 66.7% were rural residents.

In considering the maternal khat chewing and alcohol consumption habits, 18 (7.9%) of the mothers had a habit of khat chewing and similar proportions (7.9%) of the mothers had a habit of alcohol consumption. In addition, one newborn was died among the khat chewer mothers, which accounts for
6.7% of the dead newborns. Similarly, one newborn with perinatal asphyxia was died among the mothers who had a history of alcohol intake, which accounts for 6.7% of the dead newborns with perinatal asphyxia (Table 1).
| Variables                     | Category     | Status of the newborn |
|-------------------------------|--------------|-----------------------|
|                               |              | Died      | Survived |
|                               |              | n(%)  | n(%)  |
|                               |              | Died  | Survived |
| Sex                           | Male         | 18(60%) | 324(61.4%) |
|                               | Female       | 18(40%) | 204(38.6%) |
| Maternal age                  | < 20         | 9(20%)  | 69(13.1%)  |
|                               | 20–34        | 15(33.3%) | 390(73.9%) |
|                               | > 35         | 21(46.7%) | 69(13.1%)  |
| Marital status                | Not married  | 30(66.7%) | 21(4.0%)   |
|                               | Married      | 15(33.3%) | 507(96.0%) |
| Religion                      | Orthodox     | 21(46.7%) | 228(43.2%) |
|                               | Muslim       | 6(13.3%) | 105(19.9%) |
|                               | Protestant   | 18(40.0%) | 165(31.3%) |
|                               | Others       | 0(0.00%) | 30(5.7%)   |
| Educational status of the mother | Unable to read and write | 9(20.0%) | 54(10.2%) |
|                               | Able to read and write | 12(26.7%) | 105(19.9%) |
|                               | Grade 1–8    | 9(20.0%) | 207(39.2%) |
|                               | Grade 9–12   | 9(20.0%) | 105(19.9%) |
|                               | College and above | 6(13.3%) | 57(10.8%)  |
| Occupational status of the mother | House wife | 9(20.0%) | 42(8.0%)   |
|                               | Self-employee | 12(26.7%) | 120(22.7%) |
|                               | Farmers      | 9(20.0%) | 198(37.5%) |
|                               | Merchant     | 9(20.0%) | 117(22.2%) |
|                               | Civil servant | 6(13.3%) | 51(9.7%)   |
| Family size                   | < 4          | 9(20.0%) | 264(50.0%) |
|                               | 4–6          | 15(33.3%) | 177(33.5%) |
|                               | > 6          | 21(46.7%) | 87(16.5%)  |
| Place of residence            | Urban        | 15(33.3%) | 423(80.1%) |
|                                   | Rural          | 30(66.7%) | 105(19.9%) |
|-----------------------------------|---------------|-----------|------------|
| Distance b/n home and hospital    | < 10km        | 9(20.0%)  | 357(67.6%) |
|                                   | 10-50km       | 12(26.7%) | 102(19.3%) |
|                                   | > 50km        | 69(13.1%) | 24(53.3%)  |
| Estimated monthly income (ETB)    | < 1399        | 18(40.0%) | 36(6.8%)   |
|                                   | 1400–1999     | 9(20.0%)  | 114(21.6%) |
|                                   | 2000–2599     | 9(20.0%)  | 189(35.8%) |
|                                   | > 2600        | 9(20.0%)  | 189(35.8%) |

**Obstetric related characteristics**

In this study, 81(14.1%) of the mothers had no antenatal visits. In addition, 207(36.1%), 87(15.2%), 78(13.6%), and 120(20.9%) of the mothers had one, two, three, and four antenatal visits, respectively. In considering gravidity and parity, 177(30.9%) of the mothers were primigravids for this birth and nearly one fourth (25.7%) of the mothers were primiparous.

Regarding the number of pervaginal examinations, 351(61.3%) of the mothers had one to three pervaginal examinations. Nearly one fths (19.9%) of the newborns had a meconium-stained amniotic fluid. Among the respondents, 465(81.2%), 60(10.5%), 114(19.9%), 129(22.5%) and 120(2.9%) of the newborn's mother faced the spontaneous onset of labor, obstructed labor, prolonged labor, PROM, and prolonged rupture of membranes, respectively.

Nearly one fths (21.5%) of the newborns were delivered at the health center. Among those who were delivered from the health center, 33(71.3%) were died. In considering complications during delivery, 102(17.8%) of the newborns with perinatal asphyxia had cord prolapse, and 129(22.5%) of the newborns present with breech presentation. In addition, 408(71.2%), 117(20.4%), and 48(8.4) of the newborns were delivered by SVD and assisted instrumental and cesarean sections, respectively (Table 2).
Table 2
Obstetric related characteristics of the newborns with perinatal asphyxia

| Variables                              | Category             | Status of the newborn | Died      | Survived  |
|----------------------------------------|----------------------|-----------------------|-----------|-----------|
|                                        | n(%)                 | n(%)                  | Died      | Survived  |
|                                        |                      |                       | n         | n         |
| Number of ANC visits                   | No                   | 18(60.0%)             | 54(10.2%) |           |
|                                        | One                  | 9(20.0%)              | 198(37.5%)|           |
|                                        | Two                  | 3(6.7%)               | 84(15.9%) |           |
|                                        | Three                | 3(6.7%)               | 75(14.2%) |           |
|                                        | Four and above       | 3(6.7%)               | 117(22.2%)|           |
| Gravidity                              | Primigravida         | 6(13.3%)              | 171(32.4%)|           |
|                                        | Multigravida         | 39(86.7%)             | 357(67.6%)|           |
| Parity                                 | Primipara            | 3(6.7%)               | 144(27.3%)|           |
|                                        | 2–4 birth            | 15(33.3%)             | 159(30.1%)|           |
|                                        | Five and above births| 27(60.0%)             | 225(42.6%)|           |
| Number of pervaginal examinations      | 1–3                  | 21(46.7%)             | 330(62.5%)|           |
|                                        | Four and above       | 24(53.3%)             | 198(37.5%)|           |
| History of Meconium stained amniotic fluid | Yes                  | 36(80.0%)             | 78(14.8%) |           |
|                                        | No                   | 9(20.0%)              | 450(85.2%)|           |
| Onset of labor                         | Spontaneous          | 36(80.0%)             | 429(81.3%)|           |
|                                        | Induced              | 9(20.0%)              | 99(18.8%) |           |
| Antepartum hemorrhage                  | Yes                  | 12(26.7%)             | 120(22.7%)|           |
|                                        | No                   | 33(73.3%)             | 408(77.3%)|           |
| Obstructed labor                       | Yes                  | 18(60.0%)             | 33(6.3%)  |           |
|                                        | No                   | 18(40.0%)             | 495(93.8%)|           |
| Duration of labor (hrs)                | Less than 18         | 33(73.3%)             | 81(15.3%) |           |
|                                        | Greater than 18      | 12(26.7%)             | 447(84.7%)|           |
| Premature rupture of membrane          | Yes                  | 30(66.7%)             | 99(18.8%) |           |
|                                        | No                   | 15(33.3%)             | 429(81.3%)|           |
| Prolonged rupture of membrane          | Yes                  | 33(73.3%)             | 87(16.5%) |           |
|                                  | No     | 12(26.7%) | 441(83.5%) |
|----------------------------------|--------|-----------|------------|
| Cord prolapse                    | Yes    | 33(73.3%) | 63(13.1%)  |
|                                  | No     | 12(26.7%) | 459(86.7%) |
| Presentation                     | Cephalic | 15(33.3%) | 429(81.3%) |
|                                  | Breech | 30(66.7%) | 99(18.8%)  |
| Mode of delivery                 | Spontaneous vaginal delivery | 18(40.0%) | 390(73.9%) |
| Mode of delivery                 | Assisted instrumental | 21(46.7%) | 96(18.2%)  |
| Mode of delivery                 | Cesarean section | 6(13.3%) | 42(8.0%)   |
| Gestational age(week)            | <37    | 12(26.7%) | 279(52.8%) |
| Gestational age(week)            | 37–42 | 12(26.7%) | 198(37.5%) |
| Gestational age(week)            | >42    | 21(46.75) | 51(9.7%)   |

**Newborn related characteristics and Medical related characteristics of the mother**

Nearly half (50.8%) of the newborns were delivered at gestational age, less than 37 weeks and 72(12.6%) of them were post terms (gestational age more than 42 weeks). In this study, a total of 573 newborns had a first minute APGAR score of three or less. All of the newborns were resuscitated at birth and 471(82.2%) of the newborns did not cry during birth. Considering the birth weight, 186(32.5%) of the newborns had a birth weight less than 2500 grams, of which 30 newborns died, which accounts for 66.7% of the dead newborns. Twenty-one (3.7%) of the admitted newborns had at least one spot of convulsion or spasm.

Pregnancy induced hypertension (PIH), maternal iron deficiency anemia; maternal diabetes mellitus (DM) and maternal HIV status were the identified health problems affecting the survival status of newborns with perinatal asphyxia. In this study, 165(28.8%), 105(18.3%), 18(3.1%) and 30(5.2%) of the mothers had a history of diagnosed PIH, iron deficiency anemia, DM and HIV respectively (Table 3).
## Table 3
Newborn related and maternal medical disorders affecting the survival status of the newborns with perinatal asphyxia

| Variable                           | Category | Status of the newborn | Died n(%) | Survived n(%) |
|------------------------------------|----------|-----------------------|-----------|---------------|
| Cry immediately at birth           | Yes      | 3(6.7%)               | 99(18.8%) |
|                                   | No       | 42(93.3%)             | 429(81.3%)|
| History of convulsion or spasm     | Yes      | 12(26.7%)             | 9(1.7%)   |
|                                   | No       | 33(73.3%)             | 519(98.3%)|
| Birth weight (gram)                | < 2500   | 30(66.7%)             | 156(29.5%)|
|                                   | ≥ 2500   | 15(33.3%)             | 372(70.5%)|
| Pregnancy induced hypertension     | Yes      | 36(80.0%)             | 129(24.4%)|
|                                   | No       | 9(20.0%)              | 399(75.6%)|
| Maternal Iron deficiency anemia    | Yes      | 24(53.3%)             | 81(15.3%) |
|                                   | No       | 21(46.7%)             | 447(84.7%)|
| Maternal Diabetes mellitus         | Yes      | 3(6.7%)               | 15(2.8%)  |
|                                   | No       | 42(93.3%)             | 513(97.2%)|
| HIV status                         | Positive | 3(6.7%)               | 27(5.1%)  |
|                                   | Negative | 42(93.3%)             | 501(94.9%)|

### The survival status of newborns with perinatal asphyxia

The cumulative proportion of survival among the newborns admitted with perinatal asphyxia was 95.21% (95%CI: 91.00, 97.48) at the end of the first follow-up day. In addition, it was 92.82% (95%CI: 87.95, 95.77), 92.02% (95%CI: 86.84, 95.22) and 90.78% (95%CI: 84.82, 94.48) at the end of the second, third, and fourth follow-up days respectively. As we have seen from Fig. 2, there was a rapid decline of survival on the first day and it became slow in the corresponding follow-up days. The overall mean survival time was 6.55 (95%CI: 6.33, 6.77) (Fig. 2).

Among newborns admitted with perinatal asphyxia 27 (4.71%) died in the first follow-up day, which is 60% of the observed deaths within the study period. Similarly, the proportion of death at the second and third follow-up days was 2.09% and 0.52% respectively. There was no observed death after the fourth follow-up. In this study, 45(7.85%) of the newborns with perinatal asphyxia died and 531 (92.15%) were recovered (Fig. 3).
Log rank estimate of the covariates of variables

The Kaplan Meier survival curve together with the log-rank test estimates the chi square and p-value of each variable. Distance between home and hospital, referral status of the newborn, meconium-stained amniotic fluid, obstructed labor, premature rupture of membrane, prolonged rupture of membrane, cord prolapse, presentation, place of delivery, mode of delivery, history of convulsion or spasm, birth weight, pregnancy-induced hypertension, and iron deficiency anemia were candidate variables for multivariable analysis in cox proportional hazard model (Table 4).
Table 4

The log rank estimate of the variables determining the survival status among newborns admitted with perinatal asphyxia

| Variables                                           | Log rank estimate | Chi square (X²) | P-value |
|-----------------------------------------------------|-------------------|-----------------|---------|
| Sex                                                 | 0.003             | 0.959           |
| Maternal age                                        | 1.365             | 0.81            |
| Marital status                                      | 3.85              | 0.64            |
| Religion                                            | 1.48              | 0.68            |
| Educational status of the mother                    | 2.97              | 0.56            |
| Occupational status of the mother                   | 3.73              | 0.44            |
| Family size                                         | 2.21              | 0.310           |
| Place of residence                                  | 1.79              | 0.32            |
| Distance between home and hospital                  | 20.05             | 0.0001          |
| Referral status                                     | 26.02             | 0.0001          |
| Estimated monthly income (ETB)                      | 1.89              | 0.53            |
| Maternal Khat chewing status                        | 0.089             | 0.765           |
| Alcohol intake                                      | 0.031             | 0.86            |
| Number of ANC visits                                | 3.09              | 0.74            |
| Gravidity                                           | 2.31              | 0.129           |
| Parity                                              | 3.27              | 0.194           |
| Number of pervaginal examinations                   | 1.43              | 0.23            |
| Meconium stained amniotic fluid                     | 37.04             | 0.0001          |
| Onset of labor                                      | 0.005             | 0.94            |
| Antepartum hemorrhage                               | 0.134             | 0.715           |
| Obstructed labor                                    | 43.79             | 0.0001          |
| Duration of labor                                   | 5.43              | 0.631           |
| Premature rupture of membrane                       | 17.9              | 0.0001          |
| Prolonged rupture of membrane                       | 28.35             | 0.0001          |
| Cord prolapse                                       | 35.44             | 0.0001          |
| Variables                          | Log rank estimate |
|-----------------------------------|-------------------|
| Presentation                      | 18.40             |
| Place of delivery                 | 26.02             |
| Mode of delivery                  | 8.36              |
| Gestational age                   | 0.48              |
| Cry immediately at birth          | 1.38              |
| History of convulsion or spasm    | 26.32             |
| Birth weight(gram)                | 8.94              |
| Pregnancy induced hypertension    | 21.12             |
| Maternal Iron deficiency anemia   | 13.15             |
| Maternal Diabetes mellitus        | 0.733             |
| Maternal HIV status               | 0.07              |

The mean survival time was different among the covariates of each predictor. The mean survival time was higher among the newborns that had no cord prolapse as compared with those who had cord prolapse during delivery. Similarly, the average survival time was higher among newborns born with a mother who had no pregnancy-induced hypertension and no diagnosed iron deficiency anemia as compared with those with pregnancy-induced hypertension and iron deficiency anemia respectively (Table 5).
Table 5
The mean survival time estimate of the newborns with perinatal asphyxia among the covariates of predictors

| Variables                      | Category | Mean survival time (95%CI) |
|--------------------------------|----------|---------------------------|
| Cord prolapse                  | Yes      | 5.19(4.31, 6.07)          |
|                                | No       | 6.85(6.70, 6.99)          |
| Pregnancy induced hypertension | Yes      | 5.76(5.14, 6.38)          |
|                                | No       | 6.87(6.73, 7.01)          |
| Iron deficiency anemia         | Yes      | 5.72(4.94, 6.50)          |
|                                | No       | 6.74(6.56, 6.93)          |
| Convulsion or spasm            | Yes      | 3.04(2.07, 4.02)          |
|                                | No       | 6.65(6.45, 6.85)          |

Predictors of mortality among newborns with perinatal asphyxia

In this study, cord prolapse, pregnancy induced hypertension, iron deficiency anemia of the mother and having a history of convulsion or spasm of the newborn were statistically significant in the multivariable cox regression model.

Newborns having cord prolapse during delivery had six times higher risk of mortality as compared with those who had no cord prolapse (AHR: 6.5; 95%CI: 1.18, 36.01). The risk of mortality among newborns with perinatal asphyxia and delivered with mothers who had a history of pregnancy induced hypertension was 25 times higher as compared with those who had no pregnancy induced hypertension (AHR: 25.4; 95%CI: 3.68, 175.0).

Newborns with perinatal asphyxia and delivered with mothers with iron deficiency anemia had five times higher risk mortality as compared with those mothers who had no iron deficiency anemia (AHR: 5.9; 95%CI: 1.19, 29.5). Newborns admitted with perinatal asphyxia and had history of convulsion or spasm had 10 times higher risk of mortality as compared with those who had no history of convulsion of spasm (AHR: 10.23; 95%CI: 2.24, 46.54)(Table 6).
Table 6
Predictors of mortality among newborns admitted with perinatal asphyxia.

| Variables                        | Category                          | Status | COR(95%CI) | AOR(95%CI) |
|----------------------------------|-----------------------------------|--------|------------|------------|
|                                  | Died                              | Survived |           |            |
| Distance b/n home and hospital   | < 10km                            | 3       | 119        | 1          | 1          |
|                                  | 10-50km                           | 4       | 34         | 4.26(0.95, 19.04) | 0.24(0.01, 5.36) |
|                                  | > 50km                            | 23      | 8          | 11.53(3.04, 43.65)* | 0.18(0.007, 5.01) |
| Referral status                  | Yes                               | 11      | 30         | 10.52(3.34, 33.05)* | 0.65(0.02, 16.62) |
|                                  | No                                | 4       | 146        | 1          | 1          |
| Meconium stained amniotic fluid  | Yes                               | 12      | 26         | 16.95(4.78, 60.12)* | 1.46(0.17, 12.24) |
|                                  | No                                | 3       | 150        | 1          | 1          |
| Obstructed labor                 | Yes                               | 9       | 11         | 13.76(4.89, 38.69)* | 0.48(0.02, 7.87)  |
|                                  | No                                | 6       | 165        | 1          | 1          |
| Premature ROM                    | Yes                               | 10      | 33         | 7.08(2.42, 20.72)* | 1.54(0.20, 11.90) |
|                                  | No                                | 5       | 143        | 1          | 1          |
| Prolonged ROM                    | Yes                               | 11      | 29         | 11.34(3.60, 35.70)* | 0.50(0.02, 10.84) |
|                                  | No                                | 4       | 147        | 1          | 1          |
| Cord prolapse                    | Yes                               | 11      | 23         | 13.68(4.35, 43.02)* | 6.5(1.18, 36.01)** |
|                                  | No                                | 4       | 153        | 1          | 1          |
| Presentation                     | Cephalic                          | 5       | 143        | 1          | 1          |
|                                  | Breech                            | 10      | 33         | 7.22(2.46, 21.14)* | 4.1(0.91, 18.54)  |
| Place of delivery                | Health center                     | 11      | 30         | 10.52(3.34, 33.05)* | 6.4(0.73, 6.82)   |
|                                  | Hospital                          | 4       | 146        | 1          | 1          |
| Birth weight(gram)               | < 2500                            | 10      | 52         | 2.09(1.22, 3.58)* | 2.99(0.73, 12.13) |

* p < 0.05; ** p < 0.01
| Variables                          | Category                       | Status | COR(95%CI)          | AOR(95%CI)          |
|-----------------------------------|--------------------------------|--------|---------------------|---------------------|
|                                   |                                | Died   | Survived           | Died               |
| Died                             | > 2500                         | 5      | 124                | 1                   | 1                   |
| Died                             | Pregnancy induced hypertension | Yes    | 12                 | 43                 | 3.23(1.71, 6.09)*   | 25.4(3.68, 175.0)** |
| Died                             | Maternal Iron deficiency anemia| Yes    | 8                  | 27                 | 2.28(1.37, 3.79)*   | 5.9(1.19, 29.5)**   |
| Died                             | Mode of delivery               | SVD    | 6                  | 130                | 1                   | 1                   |
| Died                             | Assisted instrumental          | 7      | 32                 | 4.2(1.4, 12.65)*   | 0.87(0.15, 5.12)    |
| Died                             | CS                              | 2      | 14                 | 2.96(0.59, 14.70)  | 5.86(0.84, 40.77)   |
| Died                             | Convulsion                      | Yes    | 4                  | 3                  | 3.27(1.84, 5.82)*   | 10.23(2.24, 46.54)** |
| Died                             | No                              | 11     | 173                | 1                   | 1                   |

Key note: * indicates variables which have p-value < 0.25 and ** indicates variables which have p-value < 0.05

**Discussion**

This study assesses the predictors of mortality among newborns admitted with perinatal asphyxia at public hospitals in Southern Ethiopia and it showed there was a high proportion of mortality at the early admission periods especially at the first day and gradually declines as the follow-up period has been increased.

Newborns having cord prolapse during delivery had six times higher risk of mortality as compared with those who had no cord prolapse (AHR: 6.5; 95%CI: 1.18, 36.01). This study finding is similar with the study conducted in Karachi, Pakistan (2). The principal causes of perinatal asphyxia in this context was thought to be cord compression and umbilical arterial vasospasm which prevents venous and arterial blood flow to and from the fetus. In addition; it can predispose other factors that lead the newborn to die such as assisted ventilation requirement, low cord pH, meconium aspiration, hyaline membrane disease, convulsion, neonatal encephalopathy, and cerebral palsy (16, 17).

Consistent with the study conducted at Dilla University referral hospital, Southern Ethiopia, Tigray regional state, Ethiopia and tertiary care center in Ahmedabad, Gujarat, India and Ayder comprehensive specialized hospital, Northern Ethiopia (18, 19, 20, 21), the risk of mortality among newborns with
perinatal asphyxia and delivered with mothers who had a history of pregnancy induced hypertension was 25 times higher as compared with those who had no pregnancy-induced hypertension (AHR: 25.4; 95%CI: 3.68, 175.0). This might be related to the effect of diminished uteroplacental blood flow and placental ischemia, which can be due to pregnancy-induced hypertension which reduces blood flow to the fetus (22, 23). In addition; pregnancy-induced hypertension (PIH) has an effect on the reduction of blood supply, nutrients, and oxygen to the fetus at intrauterine life, finally which ends up in intrauterine growth restriction. This condition can contribute to newborn mortality with perinatal asphyxia (24).

Newborns with perinatal asphyxia and delivered by mothers who have iron deficiency anemia had five times higher risk mortality as compared with the counterparts who had no iron deficiency anemia (AHR: 5.9; 95%CI: 1.19, 29.5). This study finding is concise with the study done at Dilla University referral hospital, Southern Ethiopia, Southern Nations Nationalities and Peoples Regional State of Ethiopia and Jimma Zone, Southwest Ethiopia (16, 25, 26).

Newborns admitted with perinatal asphyxia and had a history of convulsion of spasm had a 10 times higher risk of mortality as compared with those who had no history of convulsion of spasm (AHR: 10.23; 95%CI: 2.24, 46.54). The possible reason might be convulsion that may cause the newborn to cease breathing (apnea). If this interruption in breathing persists, it can result in a decline in oxygen saturation in the blood to a life-threatening level.

Irrespective of the study conducted at Southern Nepal, Ayder comprehensive specialized hospital, Dilla University referral hospital, primiparity, place of delivery, multi-parity, low birth weight, mode of delivery and premature rupture of the membrane was not statistically significant predictors of mortality (18, 27, 28).

Limitation of the study

Since the study was conducted among the newborns delivered at public hospitals, it cannot be generalized for the newborns delivered at home, health centers and health posts. In addition, this study did not assess the complications secondary to perinatal asphyxia.

Conclusion

The survival status of newborns admitted with perinatal asphyxia was low at the early follow up periods after admission to the hospital and the risk of mortality was decreased at the later follow up periods and having cord prolapse of the newborn during delivery, maternal history of pregnancy-induced hypertension, maternal history of iron deficiency anemia and newborn history of convulsion or spasm were the independent predictors of mortality.

Abbreviations

ANC Antenatal Care
APGAR Appearance, pulse, Grimace, Activity and Respiratory effort

APH Antepartum hemorrhage

DM Diabetes mellitus

PIH Pregnancy induced hypertension

WHO World Health Organization

**Declarations**

**Consent for publication**

Not applicable

**Availability of data and materials**

The data sets generated and/or analyzed are available with a reasonable request through the corresponding author.

**Competing interest**

I declare that the authors have no any conflict of interest.

**Funding**

Not applicable

**Author's contribution**

All authors made a significant contribution to the work reported, whether that is in the conception, study design, execution, acquisition of data, analysis and interpretation, or in all these areas; took part in drafting, revising or critically reviewing the article; gave final approval of the version to be published; have agreed on the journal to which the article has been submitted; and agree to be accountable for all aspects of the work.

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Figures
Figure 1

The schematic presentation of sampling procedure for the study on predictors of mortality among newborns presented with perinatal asphyxia at public hospitals at Southern Ethiopia.
Figure 2

The Kaplan Meier estimate of the survival function among newborns admitted with perinatal asphyxia
Figure 3

The Kaplan Meier estimate of hazard of death among newborns admitted with perinatal asphyxia