1	Exploring the factors associated with birth asphyxia among the new-born
2	infants at a rural hospital in Bangladesh
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4	Running Title: Birth asphyxia among the new-born infants
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29 30	Abstract:
50	AU311 all.

Background: Birth asphyxia is a critical problem to address within the context of public 32 health. There are many reasons a baby may not be able to take in enough oxygen before, 33 during, or just after birth. **Objective:** The objective of this research is to determine the risk 34 factors that are associated with birth asphyxia in the Matlab, Hospital Bangladesh. Methods: 35 This is a case-control study. Cases were selected from the hospital records. A group of 94 36 mothers who gave live births resulting in asphyxiated new-borns during 24th October 2006 to 37 20th August 2008 at the labour ward of Matlab hospital, Bangladesh. Results: Result 38 indicates the largest group of mothers with Asphyxiated new-born between <25 years old, 39 40 was 61 (35.5%) while in the control group it was 111 (64.5%). Foetal presentation was associated with birth asphyxia in new-born (OR= 15.21; 95% CI: 1.841-125.67; p= 0.001). 41 The study shows that infant with birth weight <2500g had 3.49 times greater risk of 42 developing birth asphyxia compared to the infant with birth weight >2500g (CI: 1.79-6.78, p 43 value = 0.00). Conclusions: A future community based study with larger sample size is 44 necessary to find the temporal relationship between Birth asphyxia and cord around 45 the neck, Gestational age at birth less than 37 weeks and prolong first stage of 46 labour. 47

48 Keywords: Birth asphyxia, newborns, Bangladesh, rural, hospital

49 Background

Birth asphyxia is a critical problem to address within the context of public health. According 50 to The World Health Organization (WHO) Birth asphyxia is defined as "failure to initiate and 51 sustain breathing at birth" and based on Apgar score as an Apgar score of <7 at one minute of 52 life¹. Each year, globally, it results in many neonatal deaths and neurological disability 53 especially in low and middle-income countries after infections and preterm births. It is the 54 third major cause of neonatal death. Globally, birth asphyxia is estimated to account for 23% 55 of the 4 million neonatal deaths² and 26% of the 3.2 million stillbirths each year^{3,4}. According 56 57 to WHO (World Health Organization) about 3% from 120 million infants born every year in developing countries suffers from birth asphyxia and estimated 900,000 die each year⁵. 58

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One of the present challenges is the lack of a gold standard for accurately defining birth
asphyxia. Because of same reason the incidence of birth asphyxia is difficult to quantify.
Definitions of birth asphyxia designed for use in hospital-based settings require evaluation of
neonatal umbilical cord pH, Apgar scores, neurological clinical status, and markers of

multisystem organ function,⁶ and are not feasible for community settings⁷. In our study we 64 have used Apgar score to determine the condition of Birth asphyxia. Risk factor of birth 65 asphyxia in Hospital-based and home-based may be similar. Birth asphyxia is a major 66 neonatal health problem in Bangladesh and it is more common in rural areas. Because in rural 67 68 areas most of births are attended by untrained birth and antenatal risk factors are not identified. This is because the pregnant mothers are not empowered to seek healthcare 69 services⁸. Therefore risk factors are not diagnosed in time. When these deliveries are 70 conducted at home by untrained personnel- results into perinatal asphyxia in most of the 71 situations. Birth asphyxia is the main causes of neonatal mortality especially in low income 72 countries and there is dearth of publications on the incidence and risk factors of severe birth 73 asphyxia in Bangladesh. On this backdrop, the purpose of this research is to explore the 74 factors responsible for birth asphyxia in the Matlab Hospital, Bangladesh. 75

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77 Methods and Data

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This is a retrospective case –control research design. The study populations were the mothers who gave live birth at Matlab hospital. In a series of 282 mothers were selected, 94 mothers were gave live births resulting in asphyxiated new-born and other 188 mother gave birth to healthy new-born. Apgar score was used to diagnose birth asphyxia in the present study. Apgar score of less than < 7 at 1 minute of birth were included. Apgar scores less than 7 were considered as cases and Apgar scores 7 or more at 1 minutes of birth were in control group.

Selection of Cases: Cases were selected from the hospital records. A group of 94 mothers
who gave live births resulting in asphyxiated new-borns during 24th October 2006 to 20th
August 2008 at the labour ward of Matlab hospital, Bangladesh.

Selection of Control: Control are the group of 188 mothers who gave live birth resulting in non-asphyxiated new-borns at same hospital at the same period of time of cases and whose Apgar score more than 7 at 1 minute of birth. To avoid misclassification bias, controls were selected from the groups of new-born who's Apgar score more than 7 at one minute of birth. Two cases of control group compared to one case of study group by selecting the sample case which is prior and after to a sample case in the study group.

Sample Size determination: The sample size was determined by using Epi-info version 6. Probability that if the two samples differ this reflects a true difference in the two populations (confidence level or $1-\alpha$) = 0.05 Probability that if the two populations differ, the two samples will show a 'significant' difference (power or $1-\beta$) = 75

98 Proportion of controls with exposure = 0.5; Odd Ratio= 2; Control: Case= 2:1

99 By using Epi-info version, I have found my cases that was 94;

- 100 So the control was $94 \ge 188$;
- 101 And total sample size was 188+92=282
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Inclusion criteria- (i) The mother who gave live birth having Apgar score was less than < 7 at
1 minute of birth (ii) The mothers with 28 weeks gestation or more (iii) Mother who deliver
at Matlab hospital.

106 Exclusion criteria- (i) Children with congenital anomalies and genetic disorders were107 excluded from the study.

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Data were collected by reviewing the Hospital records of all the cases and the selected 109 controls by a nurse who works in the Matlab Hospital. Additionally, the delivery records of 110 expectant mothers resulting in the live births have been also included to this study. The nurse 111 was trained properly before data collection and was blinded about the objectives. The data 112 was recorded to the specific data collection form, systematically designed for computer 113 coding and analysis. Information obtained from the cases and controls were coded and 114 entered twice in to the microcomputer to avoid mistake. Data was analysed using Statistical 115 Package for Social Sciences (SPSS) ver. 16. Quality and reliability of collected data were re-116 117 checked. Following the WHO and Bangladesh Medical Research Council (BMRC) guidelines of ethical consideration, and the informed consent was taken before the data 118 119 collection process.

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121 **Results**

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Table 1 shows the general characteristics of the subjects of case and control groups including 123 the OR, with the association of Birth asphyxia. This result indicates the largest group of 124 mother with Asphyxiated new-born between <25 years old, was 61 (35.5%) while in the 125 control group it was 111 (64.5%). On the other hand, the second largest group age between 126 26-35 years old were in case group of 21(33.3%) and control group were 42 (66.7\%). The 127 third largest group below 20 years old or lower being in case group were 11 (26.8%) and 128 control group were 30 (73.2%). The smallest group found in oldest mother with ages 36 years 129 old or higher at 1(20%) in case group and control group were 4(80%). However maternal age 130 below 25 years (OR= 1.49; 95% CI: 0.70-3.19; p= 0.29) and age between 26-35 years (OR= 131

1.27; 95% CI: 0.54-3; p= 0.57) was not statistically significant association between maternal 132 age and asphyxia in new-born. Having one to three children in the family had OR of 0.80 and 133 more than three children had OR of 0.59 which were not statistically significant (95% CI: 134 0.48-1.32; p= 0.39) for one to three children and (95% CI: 0.11-3.03; p= 0.52) for more than 135 three children. From the statistical significance test result, it suggests that maternal Height 136 less than 145 cm had 2.22 times more risk of getting birth asphyxia on her neonates 137 compared with mother more than 145cm height which was statistically significant (OR= 138 2.22; 95% CI: 1.14-4.31; p= 0.01). 139

140 [Table 1 is here]

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Table 2 shows, 41.5% and 18.1% of new born both cases and controls had birth weight less 143 than 2500 gm while 58.5% and 82% of new born both cases and controls had birth weight 144 more than 2500 gm. The statistical significant analysis result suggests that low birth weight 145 was associated with birth asphyxia in new-born compared with new born more than 2500gm 146 (OR= 3.21; 95% CI: 1.84-5.58; p= 0.00). It is found that gestational age 37-41 weeks (OR= 147 0.45; 95% CI: 0.20-1; p= 0.057) and >42 weeks (OR= 0.897; 95.0% CI: 0.32-2.48; p= 0.83) 148 149 are not statistically significant in compared with gestational age <37 weeks at births. The head presentation of foetal are common in both mothers with asphyxiated new-born and non-150 asphyxiated new-born which consist of 92.0% and 99.4% of the sample. Whereas the 151 breech/malpresentation of foetal are not common and only consist of 8.0% and 0.6% of the 152 sample respectively. The logistic test result shows that foetal presentation was associated 153 with birth asphyxia in new-born (OR= 15.21; 95% CI: 1.841-125.67; p= 0.001). 154

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156 [Table 2 is here]

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Table 3 shows that about 28.4% of mother with asphyxiated new-born had Bad obstetric history during delivery while 71.6% mother with asphyxiated new-born had no Bad obstetric history. About 11.4% of mother with asphyxiated new-born had the history of Premature Labour while 88.6% mother with asphyxiated new-born had not such history. In contrast to mothers with non-asphyxiated new-born, 3.9% had the history of Premature Labour while 96.1% had not such history. The test result suggests that history of Premature Labour was

significantly associated with Birth Asphyxia in new born (OR = 0.319; 95% CI: 1.149-8.533; P = 0.020). Also the results that the time taken in the Prolong second stage of delivery was associated with birth asphyxia in new-born (OR = 4.943; 95% CI= 1.661-14.71; P= 0.002) and , it suggests that history of Meconium Stained Liquor were significantly associated with Birth Asphyxia in new born (OR = 5.372; 95% CI: 2.479-11.64; p = 0.00).

- 170 171
- 172 [Table 3 is here]
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Table 4 shows there are about 46.8% of mothers with asphyxiated new-born who complete 175 176 the minimum of four visits in antenatal care program and 34% of mothers who complete the three visits of antenatal care program and 19.1% of mother complete 0-2 visit of antenatal 177 care program. Similar result found in mother with non-asphyxiated new-born, about 54.3% 178 complete the program, 26.6% mothers complete the three visits of antenatal care program and 179 180 19.1% of mother who complete 0-2 visit of antenatal care program. No statistical significant association was found between 0-2 visits of antenatal care and Birth Asphyxia compared to 181 182 mother who visits total 4 antenatal visits of antenatal care program (OR= 1.15, 95% CI: .60-2.25, p= 0.66). However, Birth Asphyxia was not significantly associated with mothers who 183 184 have completed the 3 visits of antenatal care program (OR= 1.48, 95% CI: .84-2.61, p=.17). The result also indicates that cytromis induction is not associated with birth asphyxia in new-185 born (OR = 1.11, 95% CI: 0.32-3.82, p = 0.863). 186

187 **[Table 4 is here]**

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Table 5 shows that the case and control status was fitted as dependent variable in to an 189 Unconditional multiple logistic 190 regression model with new born weight (<2500gm,>2500gm), malpresentation, Meconium stained liquor, Prolong second stage of 191 delivery and Height of the mother (<145cm ,>145 cm). The OR corrected for confounding 192 193 factors were significantly associated with new born weight, malpresentation, Meconium stained liquor, Prolong second stage of delivery and Height of the mother. The infant with 194 195 birth weight <2500g had 3.49 times greater risk of developing birth asphyxia compared to the infant with birth weight >2500g (CI: 1.79-6.78, p value = 0.00). The malpresentation had 196 197 14.5 times greater risk of developing Birth asphyxia compared to the infant who's position

198 was normal during delivery (CI: 1.59-132.62 p value = 0.018). Mother with history of Meconium stained liquor had 6.8 times greater risk of developing birth asphyxia of the 199 200 infants compared to the mother who had not such type of history (CI: 2.82-16.40, p value = 0.00). The result in time study in prolong second stage of delivery shows that Prolong 2^{nd} 201 Stage had 4.9 times greater risk of developing birth asphyxia of the infants compared to the 202 mother who had not such type of history (CI: 1.45-16.92, p value = 0.010). The study showed 203 204 that mothers height <145cm had 2.1 times greater risk of developing birth asphyxia compared to the mothers height >145cm (CI: 1.13-3.89, p value = 0.018). 205

206 [Table 5 is here]

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208 **Discussion:**

In this Hospital based case-control study revealed that about 60% of male new-born had Birth 209 asphyxia in both the cases and controls. Only 3% of both cases and controls had more than 210 three children and rest had no children or up to three children that reflects well accepted 211 212 family planning in the society. Several studies showed association between parity and birth asphyxia¹⁻³. This research revealed that there are no association between birth asphyxia and 213 parity and similar to the another study that obtained in India⁴. This study could not find any 214 significant relationship between birth asphyxia and gestational age. And this study is similar 215 to the another study conducted at Jordan⁵. Again, this study could not find any association 216 217 with mother age, mother weight and mother BMI but maternal weight was about to be associated. However, wide OR (95% CI: 0.99-2.75) indicated that with larger sample size 218 significant association could have been ascertained. A Retrospective study conducted at 219 Nepal have seen that maternal age between 18-35 years was associated with Birth asphyxia⁹. 220 221 This study could not find any significant relationship between birth asphyxia and gestational age. And this study is similar to the another study conducted at Jordan⁵. Moreover, there is no 222 significant association found with mother age, mother weight and mother BMI but maternal 223 weight was found to be associated. However, wide OR (95% CI: 0.99-2.75) indicated that 224 with larger sample size significant association could have been ascertained. A Retrospective 225 study conducted at Nepal have seen that maternal age between 18-35 years was associated 226 with Birth asphyxia⁹. 227

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Infant with birth weight <2500g had 3.48 times greater risk of developing birth asphyxia
compared to the infant with birth weight >2500gm. This finding is consistent with the

findings of another study conducted at Nigeria and done over three years period, showed that 231 infants with low birth weight play a significant role in occurrence of asphyxia¹⁰. A 232 Retrospective case-control study conducted at Phramongkutklao Hospital, Thailand have seen 233 that fetal birth weight less than 2,500 grams had 2.5 times greater risk of developing birth 234 asphyxia compared to the fetal birth weight more than 2500 grams¹¹. But different from 235 another Hospital based study which was carried out in Jordan⁵. The study showed that 236 mothers height <145cm had 2.1 times greater risk of developing birth asphyxia compared to 237 the mothers' height >145cm. This is similar to a study conducted in Nepal¹². 238

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Antenatal care from a trained provider is important to monitor the status of a pregnancy and 240 to reduce the risk that is associated with the pregnancy and at delivery for the mother and 241 child. The study findings show that majority of the mother had received four visits of 242 antenatal care and only 19% of both cases and controls had received 0-2 visits of antenatal 243 care and it showed that there was no significant relationship between birth asphyxia and 244 antenatal care and the findings are contrast to the findings in the Jordan study⁵. Around 29% 245 of both cases and controls had bad obstetric history and this study also shows that, it is not 246 associated with birth asphyxia in new-born but this result is different from multiple hospital 247 based studies ^{4,13,14}. In this study malpresentation is associated with birth asphyxia in new-248 born and this study is similar to another Hospital based study which was conducted at 249 Mulago Hospital, Kampala, Uganda have seen that malpresentation had 6.32 times greater 250 risk of developing Birth asphyxia compare to the normal presentation¹⁵. There was no 251 significant difference in the premature rupture of membrane of both the case and control 252 group and it is not statistically significant but this study is different from other authors^{1,13-16}. 253 Prolonged rupture of membrane may be associated with intrauterine infection resulting in 254 birth asphyxia^{1.} 255

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Meconium should always be considered a marker for foetal distress therefore there was a 257 significant effect on the Apgar score of neonates¹⁷. Meconium-staining of the amniotic fluid 258 is present in 9-14 per cent of all deliveries at the time of delivery¹⁸. In this study, Meconium 259 was strongly associated with Birth asphyxia in new-born and this study is similar to the other 260 study¹⁹⁻²¹.A population based retrospective case-control study conducted at Sweden have 261 seen that Meconium release had 4.1 times greater risk of developing birth asphyxia compared 262 to the mother who did not have the history of Meconium release during delivery²². Coiling of 263 the umbilical cord around the fetal neck is a common complication of labour, said to occur 264

about once in every five deliveries²³. In this study the association was borderline (p=0.05). 265 Study conducted at London have found association between birth asphyxia and Cord around 266 the neck^{24.} Prolong second stage of labour is associated with birth asphyxia in new-born and 267 this study is similar to another Hospital based cohort study which was conducted at Maulana 268 Azad Medical College in New Delhi¹⁰. A retrospective study over a 3-year period from 1989 269 to 1991 was performed at a tertiary level, referral hospital have found association between 270 Birth asphyxia and prolonged second stage of labour⁹. 271

Limitations: 272

Potential limitation of the study could not be ignored. Firstly, the research relied on the 273 Hospital records where history was taken from the attendant. The nurse who collected data 274 was blinded of the objectives so; bias is unlikely in this regard. Secondly there was small 275 sample size and as it is hospital-based study. 276 No. of the second s

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278 **Conclusion and Recommendation**

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To Reduce Birth asphyxia requires the use of appropriate obstetric monitoring of pregnancy 280 and labour for risk factors of Birth asphyxia and health education of the general population. 281 Pregnancy related issues should be included in school curricula for adolescents. There is an 282 immediate need for organization of regular workshops and seminars for TBAS, employees of 283 public and private health institutions emphasising the need for early identification and prompt 284 referral of complicated pregnancies and labour to appropriate health care institutions of 285 efficient ambulance services and good network of roads. Further community-based study 286 with larger sample size is necessary to find the temporal relationship between Birth 287 asphyxia and cord around the neck, Gestational age at birth less than 37 weeks and 288 prolong first stage of labour. More Hospital based prospective case-control study 289 290 should be encouraged with larger sample size to identify the possible risk factors for Birth asphyxia. Implement the project to solve the problem from the factor found in 291 this study and do the research to appraise the project of birth asphyxia in new-born 292 problem solving project. 293

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Conflicting Interest: The authors declared that they are no conflicting interests. 296

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Highlights

- 1. Foetal presentation was associated with birth asphyxia in new-born babies.
- 2. Meconium was strongly associated with birth asphyxia
- 3. Infant with birth weight <2500g had 3.48 times greater risk of developing birth asphyxia compared to the infant with birth weight >2500gm
- 4. There is no any significant relationship between birth asphyxia and gestational age
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Table-1. Frequency distribution of Socio demographic and maternal factor by case and

control including OR and p-value

Categories	Cases	Controls	OR (95% CI)	p-value
	(n= 94)	(n= 188)		
	No (%)	No (%)		
Maternal age				
(Years)			•	
<25	45(48.0%)	92(49.0%)	1	
26-35	42(45.0%)	79(42.0%)	1.1(0.65-1.8)	0.75
>35	7(7.4%)	17(9.0%)	0.84(0.32-2.18)	0.72
Parity				
No parity	47(50.0%)	83(44.1%)	1	
1-3	45(47.9%)	99(52.7%)	0.80(0.48-1.32)	0.39
>3	2(2.1%)	6(3.2%)	0.59(0.11-3.03)	0.52
BMI				
<25	73(83%)	155(87.0%)	1	
>25	15(17%)	23(13%)	1.38(0.68-2.8)	0.36
Mother Height				
<145cm	21(24%)	22(12.4%)	2.22(1.14-4.31)	0.018*
>145cm	67(76.1%)	156(87.6%)	1	
Mother weight				
<50kg	55(59.8%)	86(47.3%)	1.65(0.99-2.75)	0.051*
>50kg	37(40.2%)	96(52.7%)	1	
Education	$\langle \cdot \rangle$			
No education	21(22.3%)	29(15.4%)	1	
Primary	25(26.6%)	58(31%)	0.59(0.28-1.23)	0.21
Secondary	48(51%)	101(53.7%)	0.65(0.34-1.26)	0.16
Asset index				
1(Poorest)	13(14.1%)	21(11.2%)	1	
2	20(21.7%)	35(18.6%)	0.92(0.38-2.23)	0.85
3	18(19.6%)	38(20.2%)	0.76(0.31-1.86)	0.55
4	16(17.4%)	40(21.3%)	0.64(0.26-1.59)	0.34
5(Richest)	25(27.2%)	54(28.7%)	0.74(0.32-1.73)	0.49

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p-value					
Categories	Cases $(n=94)$	Controls (n= 188)	OR (95% CI)	p-value	
	No (%)	No (%)			
Birth weight of					
infants (g)					
<2500gm	39(41.5%)	34(18.1%)	3.21(1.84-5.58)	0.00*	
>2500gm	55(58.5%)	154(81.9%)	1		
Gestational age					
at birth (weeks)			T P .		
<37 weeks	13(13.8%)	14(7.4%)	1		
37-42 weeks	66(70.0%)	156(83%)	0.45(0.20-1)	0.057*	
>42weeks	15(16.0%)	18(9.6%)	0.89(0.32-2.48)	0.83	
Sex of infants		Ň, Ń			
Male	55(58.5%)	110(58.5%)			
Female	39(41.5%)	789(41.5%)	1(0.6-1.6)	1	
Mal-					
presentation		$\sim \bigcirc \checkmark$			
Present	7(8.0%)	1(0.6%)	15.21(1.84-	0.001*	
Absent	81(92%)	176(99.4%)	125.67)	1	
Cord around					
the neck	\square				
Present	20(22.7%)	24(13.5%)	1.89(0.97-3.64)	0.056*	
Absent	68(77.3%)	154(86.5%)		1	
	$\langle \rangle$				
	\checkmark				

368	Table 2. Frequency	distribution of Foe	tal factors by case and	d control including OR and
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 Table 3. Frequency distribution of deliveries related factor by case and control
 including OR and p-value

Categories	Cases (n= 94) No (%)	Controls (n=188) No (%)	OR (95% CI)	p value
Bad obstetric history:				
- Present	25 (28.4)	25 (28.4)	0.96	0.89
- Absent	63 (71.6)	63 (71.6)	(0.547-1.69)	1
Leaking Membrane:				
- Present	4 (4.5)	18 (10.1)	0.42	0.12
- Absent	84 (95.5)	160 (89.9)	(0.13-1.29)	1
Premature rupture of				
membrane:		4		
- Present	9 (10.2)	13(7.3)	1.44	0.41
- Absent	79 (89.8)	165(92.7)	(0.59-3.52)	1
Premature labor				
- Present	10 (11.4)	7 (3.9)	0.31	0.02*
- Absent	78(88.6)	171(96.1)	(1.14-8.53)	1
Prolong 1st stage				
-Present	15 (17.0)	25(14.0)	1.25	0.51
-Absent	73 (83.0)	153(86 .0)	(0.62-2.59)	1
Prolong 2nd Stage				
-Present	11 (12.5)	5(2.8)	4.94	0.00*
-Absent	77 (87.5)	173(97.2)	(1.66-14.71)	1
APH*				
-Present	1 (1.1)	1 (.6)	2.03	0.55
-Absent	87 (98.9)	177 (99.4)	(0.12-32.9)	
Meconium Stained				
Liquor				
-Present	23(26.1%)	65(73.9%)	5.37	0.00*
-Absent	11(6.2%)	167(93.8%)	(2.47-11.64)	1
Prolong labour				
-Present	15(17%)	7(3.9%)	5.02	0.00*
-Absent	73(83%)	171(96.1%)	(1.96-12.82)	1

Table 4. Distribution of maternal care received factor by case and control including OR

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and p-value

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	Categories	Case(n= 94) No (%)	Control (n= 188) No (%)	OR (95% CI)	p value
	Antenatal care visits				
	0-2 nd visits	18(19.1%)	36(19.1%)	1.15(0.59-2.25)	0.66
	3 rd visits	32(34%)	50(26.6%)	1.48(0.84-2.61) 1	0.17
	4 th visits	44(46.8%)	102(54.3%)		
	Cytromis induction				
	Yes	4(5.40%)	8(4.9%)	1	
	No	70(94.6%)	156(95.1%)	1.11(0.32-3.82)	0.86
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Table 5. Risk factor for Birth asphyxia in new-born

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Factors	OR adjusted	р	OR unadjusted	р
	95% CI	Value	95% CI	Value
Birth weight of	3.49		3.21	
newborn <2500gm	(1.79-6.78)	0.00	(1.84-5.58)	0.00*
Malpresentation	14.52		15.21	
-	(1.59-132.62)	0.01	(1.84-125.67)	0.00*
Meconium stained	6.80		5.37	0.00*
liquor	(2.82 - 16.40)	0.00	(2.47-11.64)	
Prolong 2 nd Stage	4.96		4.94	
	(1.45 - 16.92)	0.01	(1.66-14.71)	0.00*
Premature labour	4.55	0.00	0.31	
	(1.62 - 12.74)		(1.15-8.53)	0.02*
Height of the mother	2.10	0.01	2.22	
C	(1.13-3.89)		(1.14-4.31)	0.01*

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