INCIDENCE OF ACUTE RENAL FAILURE IN FULL TERM NEONATES WITH BIRTH ASPHYXIA

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ABSTRACT

Introduction
Birth asphyxia is an eventuality having far reaching consequences in the neonatal period. Hypoxia and ischemia can cause damage to almost every tissue and organ in the body and various target organs involved. Renal insult is a recognized complication of birth asphyxia and carries a poor prognosis. Timely detection of renal dysfunction and appropriate management may favorably alter the prognosis in many neonates with birth asphyxia.

Objective
The present study was done to find out the incidence of acute renal failure in the full term neonates with birth asphyxia.

Methodology
A cross sectional study was conducted at Birat Medical College Teaching Hospital, Morang, Nepal from 1st September 2017 to 28th February 2018. Fifty full term neonates born with Apgar score of <6 at 5 minutes and fulfilling inclusion criteria were enrolled in the study. Asphyxiated neonates having Serum creatinine >1.5gm/dl or urine output<1ml/kg/hr were labeled as cases of Acute Renal Failure. Blood sample for serum creatinine was collected at 24hrs, 48 hrs and 72 hrs of life.

Results
A total of 50 term asphyxiated neonates were enrolled in the present study. Among them 54% and 46% were males and females respectively with male to female ratio of 1.2:1. In the present study 62% of cases developed acute renal failure in either of the first three days of life with mean urine output 1.02±0.27ml/kg/hr and mean serum creatinine of 1.49±0.32 mg/dL. The incidence of oliguric renal failure was 52% and non oliguric renal failure was 48%. The association between serum creatinine and urine output was statistically significant.

Conclusion
In the present study birth asphyxia has been an important cause of neonatal acute renal injury, revealing 31 (62%) cases. Monitoring urine output and serum creatinine has helped in detecting the asphyxiated neonates with acute renal injury in the early stage.

KEY WORDS
Acute renal failure, birth asphyxia, neonates
Case Report

INTRODUCTION

Birth asphyxia is an insult to the fetus or newborn due to lack of oxygen (hypoxia) and/or lack of perfusion (ischemia) to various organs. It is one of the major cause of neonatal death in developing countries like Nepal, India and other South East Asian countries as health services are not much more advance.

The incidence of perinatal asphyxia is about 1-1.57% in the most centers in developed countries and is usually related to gestational age, birth weight and maternal conditions account for almost 20% of perinatal mortality. Incidence of birth asphyxia is higher in developing countries and it is mainly due to poor health facilities, illiteracy and ignorance. Of 130 million newborn infants born each year globally, about 4 million die in the first 4 weeks of life and around 23% of all newborn deaths are caused by birth asphyxia, with a large proportion of stillbirths. Following improvement in primary and obstetric care in industrialized countries, the incidence of birth asphyxia has reduced significantly and less than 0.1% of newborn infants die from birth asphyxia. In developing countries, rate of birth asphyxia are several folds higher, ranging from 4.6 per 1000 in Cape Town to 26 per 3000 in Nigeria. Incidence of birth asphyxia was 76 per thousand of live births in a study conducted in India.

The assessment of birth asphyxia is made by APGAR score (heart rate, color, respiration, muscle tone and reflex responses) and the diagnosis is supported by other clinical parameters like multi organ failure, and blood gas analysis. The target organs of perinatal asphyxia injury are brain, lungs, kidney, liver, bowels etc. In a study on asphyxiated newborns, 34% had no evidence of organ injury, 23% had abnormality confined to one organ, 34% had involvement of 2 organs and 3% had 3 organs affected. Most frequent abnormalities include involvement of kidney (50%) followed by central nervous system (25%), cardiovascular system (25%) and chest (22%).

As the acute renal insult following perinatal asphyxia is one of the common complications in neonates, the present study is to carry out the incidence of renal failure in full term neonates with birth asphyxia.

The aim and objectives of this study was to determine the incidence of acute renal failure in term asphyxiated neonates.

METHODOLOGY

A cross-sectional study was undertaken at Neonatal Intensive Care Unit, Departments of pediatrics, Birat Medical College Teaching Hospital for a period of six months from 1st September 2017 to 20th February 2018. Fifty term neonates 37 to less than 42 weeks of gestational age with 5 minutes APGAR score <6 were included in the study where as neonates with gross congenital malformations, prematurity, postmaturity and low birth weight were excluded. The details of baseline information with regards to maternal age, maternal factors, maternal history of per vaginal leaking and its duration, meconium stained amniotic fluid and parity was recorded. First minute, fifth minute and tenth minute APGAR score was recorded in all asphyxiated full term neonates. Birth weight and general physical examination was performed to rule out any congenital anomalies.

In this study, birth asphyxia was defined as fifth minute APGAR score <6 and acute renal failure was defined as plasma creatinine more than 1.5 mg/dL or urine output <1 ml/kg/hr.

Those newborns with 5 minute APGAR score <6 were evaluated and renal function test (sodium, potassium, serum urea and creatinine) were performed on 24 and 48 hours of life. Urine output was monitored till the newborns renal status was normalized. The urine sample in newborns was collected (in female neonates via urinary catheter and in male neonates via urinary bag) for at least 72 hours or till hospital stay. Along with renal function, serum calcium and other investigation were done when required. The newborns were followed up during hospital stay and details of management were recorded in all the neonates.

The statistical analysis was done by using SPSS.16 software, Chi-square and Fisher exact test was used for statistical significance between the parameters. Sample size: 50 newborns were included for the research purpose.

RESULTS

A total of 50 term asphyxiated neonates were enrolled in the present study. Among them 54% and 46% were males and females respectively with male to female ratio of 1.2:1 (Figure 1). At 24 hrs, 48 hrs and 72 hrs of life 40% (20), 52% (26) and 24% (12) had oliguria respectively. Overall 52% (26) of them had oliguric renal failure on either of first three days of life (Table 1).

![Sex Distribution of Newborns](image-url)

**TABLE 1: Distribution of Newborns According to Urine Output on 1st 3 Days of Life**

<table>
<thead>
<tr>
<th>DAY</th>
<th>Number of newborns (%) having urine output &lt;1 ml/kg/hr</th>
<th>Number of newborns (%) having urine output ≥1 ml/kg/hr</th>
</tr>
</thead>
<tbody>
<tr>
<td>Day 1</td>
<td>20 (40%)</td>
<td>30 (60%)</td>
</tr>
<tr>
<td>Day 2</td>
<td>26 (52%)</td>
<td>24 (48%)</td>
</tr>
<tr>
<td>Day 3</td>
<td>12 (24%)</td>
<td>38 (76%)</td>
</tr>
<tr>
<td>On either of 3 days</td>
<td>26 (52%)</td>
<td>24 (48%)</td>
</tr>
</tbody>
</table>
Similarly at 24 hrs, 48 hrs and 72 hrs of life 32% (16), 26% (13) and 22% (11) had raised creatinine level respectively (Table 2). Thirty percent (15) of the studied neonates had acute renal failure and oliguria while 12% (6) of them had non oliguric renal failure (Table 3). In the present study 62% of cases developed acute renal failure in either of the first three days of life with mean urine output 1.02±0.27ml/kg/hr and mean serum creatinine of 1.49±0.32 mg/dl. The incidence of oliguric renal failure was 52% and non oliguric renal failure was 48%. The association between serum creatinine and urine output was statistically significant.

**Table 2: Distribution of Newborns According to Serum Creatinine up to 3 Days of Life**

<table>
<thead>
<tr>
<th>DAY</th>
<th>Number of newborns having serum creatinine (%)</th>
<th>Number of newborns having serum creatinine (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>&gt;1.5mg/dL</td>
<td>&lt;1.5mg/dL</td>
</tr>
<tr>
<td>Day 1</td>
<td>16 (32%)</td>
<td>34 (68%)</td>
</tr>
<tr>
<td>Day 2</td>
<td>13 (26%)</td>
<td>37 (74%)</td>
</tr>
<tr>
<td>Day 3</td>
<td>11 (22%)</td>
<td>39 (78%)</td>
</tr>
<tr>
<td>On either of 3 days</td>
<td>21 (42%)</td>
<td>29 (58%)</td>
</tr>
</tbody>
</table>

**Table 3: Relation Between Serum Creatinine and Urine Output (n=50)**

<table>
<thead>
<tr>
<th>Number of cases with serum creatinine</th>
<th>Number of cases with urine output &lt;1ml/kg/hr</th>
<th>Number of cases with urine output ≥1ml/kg/hr</th>
<th>Total</th>
<th>p value</th>
</tr>
</thead>
<tbody>
<tr>
<td>&gt;1.5mg/dL</td>
<td>15 (30%)</td>
<td>6 (12%)</td>
<td>21</td>
<td>0.019</td>
</tr>
<tr>
<td>≤1.5mg/dL</td>
<td>11 (22%)</td>
<td>18 (36%)</td>
<td>29</td>
<td></td>
</tr>
<tr>
<td>Total</td>
<td>26 (52%)</td>
<td>24 (48%)</td>
<td>50</td>
<td></td>
</tr>
</tbody>
</table>

**DISCUSSION**

Birth asphyxia is an insult to the fetus or the newborn due to lack of oxygen (hypoxia) and/or lack of perfusion (ischemia) to various organs of sufficient magnitude and duration to produce functional and/or biochemical changes. As kidneys are very sensitive to oxygen deprivation, renal insufficiency may occur within 24 hours of a hypoxic ischemic episode which if prolonged may even lead to irreversible cortical necrosis. ARF is the commonest complication and carries a poor prognosis and may even result in permanent renal damage in up to 40% of survivors.

This prospective cross sectional study was carried out in 50 full term asphyxiated neonates to assess the renal functions, to find out the incidence of acute renal failure. Renal function was evaluated by monitoring urine output serum creatinine, Serum creatinine, urine output were estimated at 24 hrs, 48hrs and 72 hrs. Those having either raised serum creatinine (>1.5mg/dL) or oliguria (<1ml/kg/hr) on either of the three days were considered as cases with acute renal failure.

In the present study 27(54%) were males and 23(46%) were females with the ratio of 1.2:1, which is almost in conformity with the studies done by Roy B India(1) (1.28:1), Dalal EA India 1.1:1, Dongol S et al Nepal 1.26:1. The results in the present study is not in conformity to the study done by Gupta BD et al13 and Mangi et al19 who reported male to female ratio 1.96:1, 2.8:1 respectively. The birth weight of the newborns in this study ranged from 2.6-3.8 kg with mean 2.8±0.35 kg which is similar to the study conducted by Gupta et al14 (2.8±0.44 kg), Mangi et al19 (2.5 kg).

In this study the serum creatinine value ranged from 0.9-1.93mg/dl with mean 1.41±0.33.Serum creatinine value on the 1st, 2nd and 3rd day of life were 1.3±0.47,1.6±0.63, 1.31±0.67 respectively. Mean creatinine value was raised in the second day of life of studied neonates. According to the study done by Roy12 et al serum creatinine on day 1 was 1.14±0.57 and on 3rd day of life 1.24±0.5 which is nearly similar to the present study. The mean serum creatinine value in the present study is comparable to that done by Jayashree et al16 in which the mean serum creatinine value was 1.58±0.58, but is not comparable to the study done by Gupta et al14 and Aggarwal et al20 where the values were 1.08±0.49 and 1.0±0.5 respectively. The number of cases having raised serum creatinine(>1.5mg/dl) in the 1st, 2nd and 3rd day of life were 16 (32%), 13 (26%) and 11 (22%) respectively. 21 (42%) cases had serum creatinine values raised on either of 3 days of life. The present study was similar to the study done by Gupta et al (30%) Aggarwal et al(56%) and Hankins Gary D.V. et al (59%). The present study was dissimilar to the study done by Martin Ancel et al14 and Perlman JM et al19 who reported the elevated serum creatinine values 15% and 17% respectively.

In the present study the range of urine output varied from 0.6-1.5ml/kg/hr. The mean urine output on day 1, day 2, day 3 was 0.9±0.23, 1.0±0.38, 1.7±0.38 ml/kg/hr. Urine output improved day wise which could be attributed to prompt recognition of it and early administration of fluid boluses and furosemide. Urine output improved on 3rd day of life. This is similar to the observations of Perlman et al who reported transient oliguria on 1st day in 23% of newborns and urine output increased to normal values on 3rd day of life. In this study the mean urine output of three days was 1.02±0.27ml/kg/hr. The mean urine output was comparable to the study done by Gupta et al(14) which was 1.28±0.36ml/kg/hr.

The incidence of oliguric renal failure was 40%(20), 52%(26), 24%(12) on 1st, 2nd and 3rd day of life respectively in the present study. The incidence of oliguria was more on 2nd day of life and again decreased the next day owing to the early intervention and fluid management. This study was comparable to the study done by Shah GS et al15 in which the incidence of oliguric renal failure were 58.3%,30%,23.3%. on 1st three days of life respectively. The incidence of oliguric renal failure on either of the three days in this study is 52% which is comparable to the study done by Aggarwal et al(17) (42%) and Mohan PV17 et al (44%) but is dissimilar to the study done by Gharehbaghi et al(10%) and Mangi et al19. The incidence of non oliguric renal failure was 48% where serum creatinine was raised(>1.5mg/dL) despite normal urine output(>1ml/kg/hr).

Oliguria in this study could be attributed to birth asphyxia which leads vasoconstriction of the afferent arterioles and decreases the blood flow to the kidneys so that glomerular filtration rate decreases leading to oliguria and even anuria. Non oliguric renal failure is a recognized entity secondary to birth asphyxia. Renal parenchymal injury in non oliguric as...
well as oliguric renal failure is essentially similar but heterogenous response of individual nephron and variable damage to tubular epithelium results in anatomical damage of different severity. If damage to tubular epithelium is less severe there occurs decrease in fractional reabsorption which exceeds the decrease in single nephron GFR leading to polyuria in non oliguric renal failure.  

The incidence of ARF in the present study is 31(62%). The incidence of oliguric and non oliguric renal failure were 52 % and 48% respectively. The incidence of ARF(62%) in the present study is similar to the study results documented by Karlowicz et al27 (61%), Aggarwal et al27 (56%) and Mangi et al24(46%), but is not in similarity to the study by Gharhebaghi et al21 (36.5%), Gupta et al (47.14%).  

The variability in the results could be due to difference in place of study, different populations studied and different methods used for assessment of renal functions. 

The incidence of oliguric renal failure in this study is 52% which is comparable to the study done Aggarwal et al24 (42%) and Mohan PV et al22 (44%) but is dissimilar to the study done by Gharhebaghi et al21. 

The lesser incidence in Gharehbaghi study could be due to the fact that they excluded sepsis, respiratory distress syndrome and heart failure while evaluating renal failure which were not excluded in the present study. The incidence of non oliguric renal failure in this study is 48% which is comparable to Aggarwal et al24 (58%) and Mohan PV et al22 (56%), but was not comparable to the study done by Gharhebaghi et al21 (90%) and which was in much higher sides. 

CONCLUSION 

Out of the 50 term asphyxiated newborns in our study, 31(62%) of the cases developed acute renal failure in the first three days of life considering serum creatinine>1.5 mg/dL or oliguria(<1ml/kg/hr). The limitation of the study was the sample size which was small and the cases were included from only one center.

LIMITATION OF THE STUDY 

The limitation of the study was the sample size which was small and the cases were included from only one center.

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We are thankful to all the children and their parents who agreed to be the part of this assessment.

CONFLICT OF INTEREST 

None

FINANCIAL DISCLOSURE 

None

REFERENCES 


