A comparative study of serum electrolytes in newborns with birth asphyxia and non-asphyxiated newborns

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Abstract

Background: Birth asphyxia is a common neonatal problem and contributes significantly to neonatal morbidity and mortality. HIE is the foremost concern in asphyxiated neonate because contrary to other system derangements this has the potential to cause serious long term neuromotor sequel among survivors. Sodium, potassium and calcium are the major electrolytes in human body, and any deviation from their normal levels in blood might cause convulsions, shock and other metabolic abnormalities. Immediate aggressive treatment of these abnormalities could modify the entire outcome of the babies.

Methods: The study was a case control study conducted over a period of 2 years. Case (asphyxiated) group comprised of term babies weighing 2.5 kg or more with birth asphyxia. Birth asphyxia was said to be present in those babies who had a definite APGAR score of less than 7 at 1 minute of birth. Control (non-asphyxiated) group was made up of term babies weighing 2.5 kg or more with APGAR scores of 7 or more at 1 minute of birth. Blood samples collected were immediately sent to the biochemistry lab for evaluation of serum sodium, potassium and calcium.

Results: A total of 100 newborn babies were included in the study, 50 babies in case group and 50 babies in control group. Mean serum sodium and calcium values were significantly lower in the asphyxiated group when compared to the non-asphyxiated group. Mean serum potassium was significantly higher in the asphyxiated group when compared to the non-asphyxiated group. The mean serum sodium and calcium values in severely asphyxiated babies were significantly lower when compared to mild/moderately asphyxiated babies and the non-asphyxiated group. Conclusions: As serum sodium levels are low in birth asphyxia, fluids must be managed judiciously in asphyxiated newborns.

Key words: Asphyxia, Calcium, Potassium, Sodium

Introduction

Birth asphyxia is a common neonatal problem and contributes significantly to neonatal morbidity and mortality[1]. The signs of asphyxial injury are nonspecific and overlap with other illnesses. In the absence of perinatal records, it is difficult to retrospectively diagnose perinatal asphyxia. Perinatal asphyxia may result in adverse effects on all major body systems. In a term infant with perinatal asphyxia renal, neurologic, and cardiac and lung dysfunction occurs in 50%, 28%, 25% and 23% cases respectively [2]. HIE is the foremost concern in asphyxiated neonate because contrary to other system derangements this has the potential to cause serious long term neuromotor sequel among survivors. Despite the increasing understanding of the mechanisms leading to and resulting from neonatal asphyxia, early determination of brain damage following hypoxic-ischemic events still remains the hardest problems in neonatal care [3,4,5]. Brief hypoxia impairs cerebral oxidative metabolism leading to an anaerobic glycolysis to generate ATP.

Anaerobic metabolism ensues with production of large quantities of metabolic degradation products like lactic acid [6,7,8,9,10]. During prolonged hypoxia, cardiac output falls, cerebral blood flow is compromised and a combined hypoxic-ischemic insult produces further failure of oxidative phosphorylation and ATP production, sufficient to cause cellular damage. Due to cellular damage the levels of serum potassium increase,
consequently the levels of sodium reduce due to increase secretion of ADH and water retention and hypoxic injury to the renal tubules. Serum levels of calcium also tend to drop due to hypoxic ischemic damage to the parathyroid glands [6,7,8,9,10]. Sodium, potassium and calcium are the major electrolytes in human body, and any deviation from their normal levels in blood might cause convulsions, shock and other metabolic abnormalities.

Calcium is an important second messenger in our body and also helps carrying out muscle function and acts as cofactor for several enzymatic activities. Knowledge of these abnormalities to the clinician is very valuable as it is an important variable affecting perinatal mortality. Immediate aggressive treatment of these abnormalities could modify the entire outcome of the babies. Though there are more and more studies for understanding mechanisms leading to birth asphyxia, studies for early determination of tissue damages due to birth asphyxia are still lacking.

**Methodology**

The study is a case control study conducted at the Neonatal Intensive Care Unit Vydehi medical college hospital Bangalore over a period of 2years. Case (asphyxiated) group comprised of term babies weighing 2.5 kg or more with birth asphyxia. Birth asphyxia was said to be present in those babies who had a definite APGAR score of less than 7 at 1 minute of birth. Severity of Birth Asphyxia was further classified as mild/moderate or severe depending on the APGAR Score at 1 minute as follows: Mild/Moderate birth asphyxia: APGAR Score 4-6, Severe birth asphyxia: APGAR Score 0-3. Control (non-asphyxiated) group was made up of term babies weighing 2.5 kg or more with APGAR scores of 7 or more at 1 minute of birth.

The babies admitted to the NICU for observation and not having birth asphyxia or any of the exclusion criteria were included in the non-asphyxiated group. Babies in the non-asphyxiated group were matched to the asphyxiated group based on the mode of delivery. A written informed consent of parents was taken before inclusion of the newborns into the study. All the neonates included in the study were subjected to detailed maternal history, thorough clinical examination, recording of APGAR score at 1 minute & 5 minutes, assessment of staging of HIE by Sarnat and Sarnat Staging and collection of blood sample using aseptic precautions for serum electrolytes (Na, K & Ca) of newborn baby immediately after birth and the details were recorded in the pre-structured proforma.

Inclusion criteria: 1) Asphyxiated group - Term babies weighing 2.5 kg or more with APGAR scores of < 7 at 1min, 2) Non-asphyxiated group- Term babies weighing 2.5 kg or more with APGAR scores of ≥ 7 at 1 minute of birth. Exclusion criteria: Suspected metabolic disease, Babies treated with diuretics, Babies born to mothers receiving drugs likely to cause CNS depression or dyselectrolytemia in babies. The gestational age was determined by using new Ballard’s score. The birth weight was measured by electronic weighing machine.

Blood samples collected were immediately sent to the biochemistry lab for evaluation of serum sodium, potassium and calcium. Serum sodium, potassium were estimated by ion selective electrode method analyzed by a radiometer. Serum calcium was estimated by Arsenazo III method using a BS380 auto-analyzer. The serum electrolyte values in the asphyxiated and non-asphyxiated groups were compared in order to evaluate whether there was any difference in serum sodium, potassium and calcium levels. Serum sodium, potassium and calcium values were compared between mild/moderate and severely asphyxiated groups, to determine if there was any variation in their values based on the severity of asphyxia.

**Statistical Methods-** Descriptive and inferential statistical analysis has been carried out in the present study. Analysis of variance (ANOVA) has been used to find the significance of study parameters between three or more groups of patients. Chi-square/Fisher Exact test has been used to find the significance of study parameters on categorical scale between two or more groups.
asphyxiated groups had a gestational age of 38 weeks followed by 37 weeks. Among the 50 neonates in the asphyxiated group, 62% were born to primi para mothers and 38% were born to multi para mothers. Among the non-asphyxiated group of 50 neonates, 56% were born to primi para mothers and 44% were born to multi para mothers [Table.1].

Table-1: Characteristics of study newborns.

| Character                  | Asphyxiated group | Non-asphyxiated group | Total | “p” Value |
|----------------------------|-------------------|-----------------------|-------|-----------|
|                            | n (%)             | n (%)                 |       |           |
| Mode of Delivery           |                   |                       |       |           |
| LSCS                       | 33 (66%)          | 33 (66%)              | 66 (66%) | 0.98  |
| Vaginal Delivery           | 17 (34%)          | 17 (34%)              | 34 (34%) |       |
| Sex                        |                   |                       |       |           |
| Male                       | 29 (58%)          | 30 (60%)              | 59 (59%) | 0.84  |
| Female                     | 21 (42%)          | 20 (40%)              | 41 (41%) |       |
| Gestational Age            |                   |                       |       |           |
| 37 Weeks                   | 16 (32%)          | 13 (26%)              | 29 (29%) | 0.42  |
| 38 Weeks                   | 22 (44%)          | 20 (40%)              | 42 (42%) |       |
| 39 Weeks                   | 7 (14%)           | 11 (22%)              | 18 (18%) |       |
| 40 Weeks                   | 4 (8%)            | 5 (10%)               | 9 (9%)   |       |
| 41 Weeks                   | 1 (2%)            | 1 (2%)                | 2 (2%)   |       |
| Parity of the Mother       |                   |                       |       |           |
| Primipara                  | 31 (62%)          | 28 (56%)              | 59 (59%) | 0.54  |
| Multipara                  | 19 (38%)          | 22 (44%)              | 41 (41%) |       |

In our study 94% of the asphyxiated group had hyponatremia (serum sodium< 135mmol/l) whereas only 8% of the non-asphyxiated group had hyponatremia and this difference was observed to be statistically significant. In our study 32% of the asphyxiated group had hyperkalemia (serum potassium>5.5mmol/l) whereas 14% of the non-asphyxiated group had hyperkalemia and this difference was observed to be statistically significant. In our study, the asphyxiated group had 96% of babies had Serum Calcium≥ 7mg/dl and only 4% had hypocalcaemia whereas all the babies in the non-asphyxiated group had Serum Calcium≥ 7mg/dl [Fig.1]. It was observed that the difference was not statistically significant (p-value>0.05). Mean serum sodium and calcium values were significantly lower in the asphyxiated group when compared to the non-asphyxiated group. Mean serum potassium was significantly higher in the asphyxiated group when compared to the non-asphyxiated group [Table2].

Table-2: Serum electrolytes status in the asphyxiated and non-asphyxiated groups.

| Serum Electrolytes | Asphyxiated group | Non-asphyxiated group | p-value |
|--------------------|-------------------|-----------------------|---------|
|                    | n=50 (MEAN±SD)    | n=50 (MEAN±SD)        |         |
| Sodium (mmol/L)    | 128.96±3.9        | 139.58±3.78           | <0.001  |
| Potassium (mmol/L) | 5.32±0.57         | 4.92±0.49             | <0.001  |
| Calcium (mg/dl)    | 7.95±0.56         | 8.80±0.63             | <0.001  |

The mean serum sodium and calcium values in severely asphyxiated babies were significantly lower when compared to mild/moderately asphyxiated babies and the non-asphyxiated group. It was also observed that the mean serum sodium and calcium levels were significantly lower in mild/ moderately asphyxiated babies when compared to the non-asphyxiated group.

The mean serum potassium value in severely asphyxiated babies when compared to mild/moderately asphyxiated babies was only marginally raised, which was not statistically significant. It was also observed that the mean serum potassium level was significantly high in mild/ moderately and severely asphyxiated babies, when compared to the non-asphyxiated group, which was statistically significant [Table.3].
Table-3: Serum electrolytes status in asphyxiated newborns of varying severity and non-asphyxiated groups

| Serum Electrolytes | Severe Birth Asphyxia n=17 (MEAN±SD) | Mild/Moderate Birth Asphyxia n=33 (MEAN±SD) | Non-asphyxiated groups n=50 (MEAN±SD) | P- values |
|--------------------|--------------------------------------|---------------------------------------------|--------------------------------------|-----------|
| Sodium (mmol/L)    | 127.18±4.864                         | 129.88±3.140                                | 139.58±3.786                        | 0.019     |
|                    |                                     |                                             |                                      | <0.001    |
|                    |                                     |                                             |                                      | <0.001    |
| Potassium (mmol/L) | 5.28±0.722                           | 5.34±0.494                                  | 4.92±0.499                          | 0.711     |
|                    |                                     |                                             |                                      | 0.020     |
|                    |                                     |                                             |                                      | 0.001     |
| Calcium (mg/dl)    | 7.69±0.488                           | 8.08±0.565                                  | 8.80±0.635                          | 0.029     |
|                    |                                     |                                             |                                      | <0.001    |
|                    |                                     |                                             |                                      | <0.001    |

Fig-1

Distribution of cases with electrolytes levels

- Serum sodium ≥ 135 mmol/L
- Serum sodium < 135 mmol/L
- Serum Potassium ≤ 5.5mmol/L
- Serum Potassium > 5.5mmol/L
- Serum Calcium ≥ 7mg/dl
- Serum Calcium < 7mg/dl

Odds Ratio: 180.17 (95%C.I 38.19-849.86). p: <0.001

Odds Ratio: 2.89 (95%C.I 1.07-7.82). p :0.032

p-value: 0.495

Discussion

This study conducted on asphyxiated and non-asphyxiated newborns, compared the pattern of electrolyte disturbances in them. It also correlated the serum electrolyte pattern with severity of birth asphyxia based on APGAR score.

In our study the incidence of asphyxia was higher in the male babies and constituted 58% of total asphyxiated cases which is comparable to the study done by Pallab Basu et al where the male babies constituted 64% of the total asphyxiated cases [11]. It was observed that the mean birth weight in the asphyxiated group was 2.74±0.36 and non-asphyxiated group was 2.78±0.33, which is almost similar to the study done by Pallab Basu et al [11]. The serum electrolyte values in the asphyxiated and non-asphyxiated groups were compared in order to evaluate whether there was any difference in their levels. In the present study, mean serum sodium level in the asphyxiated neonates was 128.96±3.9 mmol/L, which was significantly lower in comparison to the non-asphyxiated neonates, who had a value of 139.58±3.7 mmol/L (p < 0.001). The results of the present study
were in concordance with those of Pallab Basu et al [11]. They reported that the mean serum sodium level was 122.1±6.0 mmol/L, which was significantly lower compared to the non-asphyxiated neonates, who had a value of 138.8±2.7mmol/L (p < 0.001). We noticed that serum sodium levels were decreasing with the increasing severity of asphyxia showing a significant positive correlation with a r value of 0.808 and a p-value< 0.001. Pallab Basu et al also found a significant positive correlation between serum levels of sodium and APGAR score [11].

Low serum sodium levels in birth asphyxia may be due to dilutional hyponatremia caused by inappropriate secretion of anti diuretic hormone. The mean serum potassium level in the asphyxiated group was 5.32±0.57 mmol/L, which was significantly higher when compared to the non-asphyxiated group which had a value of 4.92±0.49 mmol/L ( p< 0.001).

The mean serum potassium level in the asphyxiated group was within the upper limit of the normal range. Pallab Basu et al found out that mean serum potassium level in the asphyxiated group was 5.05±0.63 mmol/L, which was slightly higher as compared to the non-asphyxiated group, which had a value of 4.19±0.40 mmol/L.(p < 0.001), but the levels were found to be within the normal range[11].

In this study serum potassium levels were increasing with the increasing severity of asphyxia showing a negative correlation with a r value of -0.325 and a p-value< 0.001. Pallab Basu et al also found a significant negative correlation between serum levels of potassium and APGAR score [11].

Higher serum potassium levels in birth asphyxia may be due to acute renal failure or hypoxic cell injury leading to release of potassium into the extracellular fluid. The mean serum calcium level in the asphyxiated group was 7.95±0.56 mg/dl, which was significantly lower in comparison to the non-asphyxiated group which had a value of 8.80±6.3 mg/dl, ( p < 0.001) but the levels were found to be within the normal range.

In the PallabBasu et al study, the mean serum calcium level in the asphyxiated group was 6.85±0.95 mg/dl and was significantly lower when compared to the non-asphyxiated group, which had a value of 9.50 ± 0.51 mg/dl; (p< 0.001)[11]. In the present study serum calcium levels were decreasing with the increasing severity of asphyxia showing a positive correlation with a r value of 0.549 and a p-value< 0.001. PallabBasu et al also found a significant positive correlation between serum levels of calcium and APGAR score [11]. Another study was done by Alphonsus N. Onyiriuka on serum calcium levels in asphyxiated term newborns, however it is not comparable with our study as they have studied serum calcium levels at 12,24 and 48 hours after birth [12]. Study done by Deepak Jajoo et al observed that serum calcium levels were significantly lower in term appropriate for gestational age asphyxiated newborns [13]. They were of the opinion that hypoxia impairs the functions of parathyroid gland resulting in lower calcium levels. However, their sample size was smaller in comparison to our study & PallabBasu et al’s study and they have also included pre-term neonates in their study.

**Conclusion**

Serum sodium levels in the asphyxiated newborns were in the hyponatremic range and in proportion to the severity of asphyxia. As serum sodium levels are low in birth asphyxia, fluids must be managed judiciously in asphyxiated newborns.

**Contribution details**

- Dr. Manjunatha Babu R- Study designing and Tabulation
- Dr. Pavan Kumar Jerry- Data collection & Data entry
- Dr. Harish G – Statistical analysis
- Dr. Susheela C- Guidance and over all supervision.

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