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Original Research Article

Clinico-Etiologic and Outcome Assessment of Modality of Respiratory Support in Preterm Neonates with Respiratory Distress

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Abstract

Aim: The aim of the present study was to determine causes of respiratory distress, pattern of respiratory modality used and its outcome among admitted preterm newborns.

Methods: A retrospective descriptive study was conducted in the Department of Pediatrics, DMCH, Darbhanga, Bihar, India over a period of two years targeting all neonates admitted to neonatology department. A Total of 200 patients were admitted in NICU during the study.

Results: Mean gestational age was 32.46±2.58 weeks. Most of the newborns (64%) belonged to the gestational age within 28 - <34 weeks category. Mean birth weight was 1665.43±585.58 g among them very low birth weight infants were 42%. Male out numbered female newborns (55% vs 45%). Total 12 (6%) infants had Apgar scores <7 at 5 minutes who required some degree of resuscitation just after birth. After admission 8 (4%) patients got single dose of surfactant. Most of the mother (66%) was multiparous and 51% of them did not receive even a single dose of antenatal corticosteroid. All of them were inborn and cesarean section was the mode of delivery for 80% of the enrolled neonates. Maternal hypertension and diabetes mellitus were present in 140 (70%) and 64 (32%) of mother respectively. Maternal risk factors for sepsis were present in 48 (24%) of infants admitted to the NICU.

Conclusion: Respiratory distress syndrome is the commonest cause of respiratory distress. Two third of preterm newborns required respiratory support. Most common mode of respiratory support was non invasive mode in the form of supplemental oxygen, Heated humidified high flow nasal cannula and continuous positive airway pressure. Short term morbidities like nasal trauma, sepsis, septic shock, disseminated intravascular coagulation, necrotising enterocolitis and intraventricular haemorrhage were more common in newborns who required invasive respiratory support ($p < 0.05$). Retinopathy of prematurity and mortality was significantly higher in invasive respiratory support group.

Keywords: Neonates, Preterm, Mechanical ventilation, Non-invasive ventilation

Introduction

Preterm neonates often develop respiratory distress syndrome (RDS) due to a surfactant deficiency. [1] Incidence of RDS is inversely proportional to gestational age. It affects approximately 98% of neonates born at or before 24 weeks, while only a quarter of Very Low Birth Weight (VLBW) neonates develop RDS. [2] With advancement in the field of neonatology and usage of maternal antenatal steroids for fetal lung maturation, the incidence and severity of RDS have decreased by 34%. [3] Despite this, RDS remains a leading complication among preterm neonates. RDS presents early with difficulty in breathing, grunting and desaturation, which may lead to type 1 and type 2 respiratory failure and ultimately developing multiorgan dysfunction if not treated optimally and timely. [1]

Treatment strategies for RDS include surfactant administration and respiratory support, including invasive and non-invasive ventilation. These respiratory support strategies have led to improved neonatal survival and outcomes. [4] Invasive ventilation may cause significant complications like atelectasis, pneumothorax and ventilator-associated pneumonia compared with non-invasive ventilation. There is an increasing trend to initiate non-invasive ventilation. [5,6] This is particularly true for low- and middle-income countries (LMIC) due to the limited resources and expertise for invasive ventilation. [7]

Respiratory distress syndrome (RDS) in newborns is the most common cause of morbidity and mortality and is an indication for ventilation in preterm

infants. [8] In recent years, the widespread implementation of nasal continuous positive airway pressure (CPAP) as the initial means of respiratory support for preterm infants has fundamentally changed respiratory management in the first hours of life. The universal use of CPAP has reduced the need for endotracheal intubation and mechanical ventilation (MV) [9] and their associated lung injuries. [10,11] The American Academy of Pediatrics [12] and the European Consensus Guidelines for the Management of RDS [13] recommended that the initial application of CPAP be considered as the optimal mode of respiratory support.

The aim of the present study was to determine causes of respiratory distress, pattern of respiratory modality used and its outcome among admitted preterm newborns.

Materials and Methods

A retrospective descriptive study was conducted in the Department of Pediatrics, DMCH, Darbhanga, Bihar, India over a period of two years targeting all neonates admitted to neonatology department. A Total of 200 patients were admitted in NICU during the study.

All inborn preterm neonates having gestational age of <37 weeks and admitted with respiratory distress were included in the study. Neonates with incomplete data and or lethal congenital anomalies were excluded from the study. Data were collected from the neonatal admission, discharge and death registers. The register contained each neonate's date of admission and discharge or death, sex, weight at admission or at birth, gestation age at birth, mode of delivery, duration of stay at the hospital, diagnosis and outcomes. The primary causes of admissions and deaths were defined as the underlying obstetric and neonatal factors or conditions, which resulted in the admission or death of the neonate. Standard definitions of the medical conditions were used for diagnosis. [14] All medical and nursing staff working at the neonatal unit was oriented on recording of the neonatal admission, discharge and death registers, clinical guidelines of diagnosis and compilation of monthly summaries for presentation at monthly perinatal mortality meetings. Extracted data included: mode of delivery, multiple gestations, use of antenatal corticosteroid, gestational diabetic mellitus (GDM), Pregnancy induced hypertension (PIH), risk factor for sepsis, sex of the baby, gestational age (weeks), birth weight (g), APGAR score at 5th minutes, Silverman- Anderson score (SAS), neonatal resuscitation, surfactant administration, fetal growth at birth (SGA, AGA, LGA), respiratory distress, cause of respiratory distress, mode of respiratory support (NIV, MV) and days of hospitalization.

Neonates born at less than 37 completed weeks (less than 259 days) of gestation were termed as preterm and those having birth weight of < 2500 g were defined Low birth weight (LBW).[14] Newborns was defined small for gestational age (SGA) if the birth weight less than the 10th percentile. Resuscitation was defined as need for intermittent positive pressure ventilation and/or cardiac compression and/or drug administration in the neonatal stabilization period. Respiratory distress in newborn was labelled when a baby had one or more signs of increased work of breathing, such as tachypnea, nasal flaring, chest retractions, or grunting.8 Severity of respiratory distress was categorized using Silverman-Anderson scoring system where Score ≥ 4 indicate clinical respiratory distress and score ≥ 7 indicate respiratory failure. [15]

Respiratory distress syndrome (RDS) was defined in neonates with increasing oxygen dependence during the first 24 h, typical radiological findings: like reduced air content, reticulo-granular pattern of the lungs, air bronchogram and/or white out lung. When newborns developed respiratory distress soon after birth and resolves within 18-24 hours of life with normal chest X-ray finding or show reduced translucency, infiltrates and hyperinflation of the lungs were labelled as Transient Tachypnea of newborn (TTN). Breathing pauses that last for > 20 seconds or for > 10 seconds if associated with bradycardia or oxygen desaturation was termed as apnea. Bronchopulmonary Dysplasia (BPD) was defined when a neonate was requiring oxygen at 36 weeks of post gestational age for babies born < 32 weeks of gestation or 28 days of age for neonates born ≥ 32 weeks of gestation or later. [15]

Maternal characteristics included as maternal age, maternal diseases like gestational and non-gestational diabetes mellitus, Pregnancy induced hypertension, infections and use of antenatal corticosteroids. Deliveries were categorized as vaginal or Cesarean Section (CS). The Data on respiratory support such as oxygen therapy administered through nasal-cannula, nasal continuous positive airways pressure (CPAP), heated humidified high flow nasal cannula (HHHFNC) and mechanical ventilation (MV) was collected. When analyzing the data, for each neonate only the highest level of respiratory support was considered. The need for specific adjunctive therapy (surfactant Administration) and the short-term outcome including morbidities and in hospital mortality were recorded. Respiratory support was divided into two groups as Invasive (Mechanical ventilation) and noninvasive which included supplemental oxygen through nasal cannula, head box, High flow nasal cannula and continuous positive airway pressure (CPAP). All the variables and mortality were compared between the two groups

Statistical analysis: Data entry and analysis was carried out by using the Statistical Package of Social Science Software program (SPSS), version 22. Categorical variables were expressed in frequency and statistical analysis was done by Chi-Square test or Fisher exact test. Continuous variable was seen in mean ± SD and statistical analysis was done by student t-test. P value <0.05 was considered statistically significant.

Results

Table 1: Baseline neonatal characteristics in studied neonates

Parameter	Value
Gestational age (weeks), Mean ±SD	32.46±2.58
Gestational age category, n (%)	
<28 weeks	12 (6)
28-<34 weeks	128 (64)
34-<37 weeks	60 (30)
Birth weight (g), Mean ± SD	1665.43±585.58
Birth weight category, n (%)	
<1000 g	20 (10)
1000-1499 g	84 (42)
1500-2499 g	80 (40)
≥2500 g	16 (8)
Sex of the baby, n (%)	
Male	110 (55)
Female	90 (45)
Multiple birth, n (%)	44 (22)
Neonatal resuscitation, n (%)	12 (6)
APGAR score at 5th minute	
≥7	188 (94)
<7	12 (6)
Silverman Anderson Score at randomization	
<4	28 (14)
4-7	172 (86)
>7	0
Surfactant administration, n (%)	8 (4)

Mean gestational age was 32.46±2.58 weeks. Most of the newborns (64%) belonged to the gestational age within 28 - <34 weeks category. Mean birth weight was 1665.43±585.58 g among them very low birth weight infants were 42%. Male out numbered female newborns (55% vs 45%). Total 12 (6%) infants had Apgar scores <7 at 5 minutes who required some degree of resuscitation just after birth. After admission 8 (4%) patients got single dose of surfactant.

Table 2: Baseline maternal characteristics in studied group

Parameter	Value
Consanguinity present, n (%)	4 (2)
Parity, n (%)	
Primipara	68 (34)
Multipara	132 (66)
Exposure to ACS, n (%)	
Complete	34 (17)
Incomplete	64 (32)
None	102 (51)
Mode of delivery, n (%)	
NVD	40 (20)
LUCS	160 (80)
GDM, n (%)	64 (32)
PIH, n (%)	140 (70)
Risk factors for sepsis, n (%)	48 (24)

Most of the mother (66%) was multiparous and 51% of them did not receive even a single dose of antenatal corticosteroid. All of them were inborn and cesarean section was the mode of delivery for 80%

of the enrolled neonates. Maternal hypertension and diabetes mellitus were present in 140 (70%) and 64 (32%) of mother respectively. Maternal risk factors for sepsis were present in 48 (24%) of infants admitted to the NICU.

Table 3: Primary disease requiring respiratory support

Parameter	Value
RDS, n (%)	102 (51)
Transient tachypnoea of newborn, n (%)	46 (23)
Congenital Pneumonia, n (%)	40 (20)
Perinatal asphyxia, n (%)	12 (6)
Meconium aspiration syndrome, n (%)	0

The respiratory distress syndrome (RDS) was the most common cause 102 (51%) following TTN 46 (23%), congenital pneumonia 40 (20%) and PNA 12 (6%).

Table 4: Level of respiratory support

Parameter	Value
Noninvasive support, n (%)	130 (65)
Invasive support, n (%)	70 (35)

Among total of 200 patients, 130 (65%) patients required NIV support including oxygen, CPAP or HFNC and 70 (35%) need Invasive support during the hospital course.

Table 5: Associated mortality and morbidity of neonates who required respiratory support

Parameters	NIV group (n=130)	IV group (n=70)	P value
PDA, n `	40	32	0.48
Sepsis, n	56	36	<0.001
Septic Shock, n	24	28	<0.001
DIC, n	8	24	<0.001
AKI, n	22	16	0.36
NEC, n	12	20	0.001
In-hospital mortality, n	12	52	<0.001

Among the associated mortality and morbidity, sepsis, septic shock, DIC, NEC and in-hospital mortality occurred significantly higher in the invasive support group (IV) in comparison to the NIV support group and the p-value were <0.05.

Table 6: Complication of Respiratory support modality in preterm neonates

Parameters	NIV group (n=130)	IV group (n=70)	P value
Nasal trauma, n	14	24	0.001
Pneumothorax, n	0	2	0.16
ROP, n	8	16	0.018
BPD, n	2	6	0.090
IVH, n	0	10	0.001

Among the complication of respiratory support only nasal trauma, sepsis, ROP and IVH occurred significantly higher in the MV support group in comparison to the NIV support group and the p-value were <0.05.

Discussion

The first 28 days of life, defined as neonatal period is the most vulnerable time for a child's survival. Every year an estimated 4 million babies die in the first 4 weeks of life [16] accounting for more than half of the under-five child deaths in most regions of the world. [17] Almost all (99%) neonatal deaths are happening in developing countries. [18] Neonatal mortality rate is one of the indicators for measuring the health status of a nation. The mortality and morbidity of preterm neonates are significantly higher than those of full-term neonates because preterm neonates are more prone to develop respiratory failure. [19] The functional immaturity of their lung structure, can lead to impaired gas exchange and requires respiratory support. [20]

Mean gestational age was 32.46±2.58 weeks. Most of the newborns (64%) belonged to the gestational age within 28 - <34 weeks category. Mean birth weight was 1665.43±585.58 g among them very low birth weight infants were 42%. Male out numbered female newborns (55% vs 45%). Total 12

(6%) infants had Apgar scores <7 at 5 minutes who required some degree of resuscitation just after birth. After admission 8 (4%) patients got single dose of surfactant. Nemr CN, et al. showed 100 cases with gestational age ranging from 27 to 40 weeks having mean of 33.98 ± 3.44 weeks. Mean birth weight in our study was 1631.44 ± 578.57g which was comparable with the same study where mean birth weight was 1580 gm. [21] Male outnumbered female newborns (54.2% vs 45.8%). Similar finding was found by Nemr CH where sixty- three (63%) were boys and thirty-seven (37%) were girls and by Iqbal Q (60% males). [22]

Most of the mother (66%) was multiparous and 51% of them did not receive even a single dose of antenatal corticosteroid. All of them were inborn and cesarean section was the mode of delivery for 80% of the enrolled neonates. Maternal hypertension and diabetes mellitus were present in 140 (70%) and 64 (32%) of mother respectively. Maternal risk factors for sepsis were present in 48 (24%) of infants admitted to the NICU. In Lategan I, et al [23] 17.3% infants of < 34 weeks had optimal ANS, birth by Cesarean Section done in 60.3% of cases, maternal hypertension was present in 43.6% of mothers. The respiratory distress syndrome (RDS) was the most common cause 51 (51%) following TTN 23 (23%), congenital pneumonia 20 (20%) and PNA 6 (6%). Among total of 100 patients, 65 (65%) patients required NIV support including oxygen, CPAP or HFNC and 35 (35%) need Invasive support during the hospital course. Among the associated mortality and morbidity, sepsis, septic shock, DIC, NEC and in-hospital mortality occurred significantly higher in the invasive support group (IV) in comparison to the NIV support group and the p-value were <0.05. Among the complication of respiratory support only nasal trauma, sepsis, ROP and IVH occurred significantly higher in the MV support group in comparison to the NIV support group and the p- value were <0.05. Nemr CN, et al. showed 72% of the studied patients underwent CPAP, 14% underwent oxygen support by nasal cannula and 14% underwent mechanical ventilation. [21] The need of invasive support was less than our study as they utilize non-invasive respiratory support as most widely used modality and practiced different ways of delivering CPAP.

In the current study, non- invasive respiratory support was given by nasal cannula, head box, CPAP and invasive support by mechanical ventilation. We used nasal CPAP with preterm neonates with recurrent apneas or early features of RDS. Those who had a failure of nasal CPAP therapy or had respiratory acidosis were ventilated. This is similar to the results of Iqbal Q, et al. in which all the preterm neonates with gestational age < 32 weeks or recurrent apnea or early features of RDS were given nasal CPAP therapy, and those who had a failure of nasal CPAP therapy were ventilated.[22] Judicious ventilator strategies, proper sedation, timely extubation and readily available X-ray and surgical facilities will help to control these problems. Mortality among sick neonates in NICU is high, but mortality among mechanically ventilated neonates is even higher. In this study, mortality in ventilated neonates was 60%, which is comparable to mortality by Hossain et al. 70.6 %. [24]

Conclusion

Respiratory distress syndrome is the commonest cause of respiratory distress. Two third of preterm newborns required respiratory support. Most common mode of respiratory support was non invasive mode in the form of supplemental oxygen, Heated humidified high flow nasal cannula and continuous positive airway pressure. Short term morbidities like nasal trauma, sepsis, septic shock, disseminated intravascular coagulation, necrotising enterocolitis and intraventricular haemorrhage were more common in newborns who required invasive respiratory support (p <0.05). Retinopathy of prematurity and mortality was significantly higher in invasive respiratory support group.

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