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# Respiratory Distress and Its Outcome among Neonates Admitted to Neonatal Intensive Care Unit of Mukalla Maternity and Child Hospital, Yemen

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## Abstract

**Background:** Respiratory distress is one of the most common causes of admission in neonatal intensive care unit .

**Objective:** To determine the causes and to study the various risk factors associated with development of respiratory distress and outcomes of respiratory distress in neonates admitted to neonatal care unit of Mukalla Maternity and Child Hospital in Al- Mukalla city, Hadhramaut Governorate, Yemen.

**Patients and Methods:** This is a prospective study covering the 12-month period between April 2018 to March 2019 to neonates admitted to neonatal care unit of Mukalla MaternityandChild hospital.

**Results:** A total of 430 patients were admitted to the neonatal care unit. Number of cases presented with respiratory distress was 250, representing 58.1% of all cases admitted. The most frequent underlying cause for respiratory distress in children was respiratory distress syndrome (44%), followed by transient tachypnea of the newborn (18.8%), birth asphyxia (14%), meconium aspiration syndrome (12%), and other causes (11.2) (CHD 10(4%), sepsis (3.2%), congenital pneumonia (2.4%), and congenital anomalies (1. 6%). The majority of cases of respiratory distress syndrome and transient tachypnea of newborn were males with statistical significant difference. The outcome of neonatal respiratory disorders was: cure in 40.4% of cases, patients discharged with complications in 10.4 % of cases, and death in 49.2% of cases. The highest case fatality rate of neonatal respiratory distress diseases were respiratory distress syndrome (81.8%) followed by meconium aspiration syndrome (56.7%) and birth asphyxia (34.3%).

**Conclusion:** Respiratory distress syndrome was the main cause of respiratory distress followed by transient tachypnea of newborn.

Keywords: Respiratory distress, neonate, causes, outcome, risk factor, NICU.

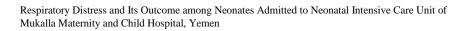
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### Introduction

Respiratory distress (RD) is a challenging problem and is one of the most common causes of admission in neonatal intensive care unit (NICU) [1]. Respiratory distress is reported in 13.3-88% of ill neonates [2-7]. A wide variety of pathologic lesions may be responsible for respiratory disturbances, including pulmonary, airway, cardiovascular, central nervous, infection, and other disorders [8]. Some risk factors increase the likelihood of neonatal respiratory disease; prematurity, meconium stained amniotic fluid, cesarean delivery, gestational diabetes, maternal chorioamnionitis and presence of prenatal ultrasonographic findings such as oligohydramnios or structural lung abnormalities [9]. Irrespective of the underlying etiology, neonatal respiratory diseases NRD has been reported to contribute to poor neonatal outcomes particularly in the developing countries [6,10]. Therefore, any healthcare professional who cares for newborn babies should easily recognize the symptoms and signs of respiratory distress, distinguish between various causes and initiate management strategies to prevent significant complications or death [11]. The aim of this study was to determine the causes and to study the various risk factors associated with development of RD and outcomes of respiratory distress in neonates admitted to NICU of Mukalla Maternity and Child Hospital (MMCH).

## **Patients and Methods**

This is a prospective study covering the 12month period between April 2018 to March 2019 to neonates admitted to NICU of MMCH in Al- Mukalla city, Hadhramaut Governorate, Yemen. Out of 430 neonates admitted to NICU of MMCH during study period 250 of them had RD which constituted 58.1% of all cases admitted.

Inclusion criteria: Were all neonate admitted to NICU in MMCH with a diagnosis of NRD. Exclusion criteria: Babies weight less than 1000 gms and babies less than 28 wks of age were excluded from the study. The following information's were recorded: The maternal data included: maternal age at pregnancy, parity, maternal hypertension, maternal diabetic, multiple gestation, antepartum hemorrhage, maternal fetal infection, Pre-eclampsia, mode of delivery, prolonged labor. Neonatal data included : Gender, gestational age, birth weight.

#### **Definition of terms**

Neonatal respiratory distress: The presence of one or more of the following signs: tachypnea > 60 breaths/min, or signs of labored breathing (expiratory grunting, nasal flaring, intercostal recessions, xyphoid recessions, thoraco-abdominal or asynchrony), with or without cyanosis criteria RDS [4].Diagnostic for were considered by the presence of clinical sign, grunting, flaring, such as tachypnea, retractions, requiring a respiratory support (supplemental oxygen requirement and/or non-invasive or invasive ventilation). Typical radiological findings were reticulogranular patterns, air bronchograms and ground glass appearance [12]. Transient tachypnea of newborn (TTN) is characterized by the early



onset of tachypnea (>60 breaths/min), sometimes with retractions or expiratory grunting and occasionally with cyanosis that is relieved by minimal O2 supplementation (<40%). The chest generally sounds clear without crackles or wheeze, and the chest radiograph shows prominent perihilar pulmonary vascular markings, fluid in the intralobar fissures, and rarely small pleural effusions [13].

Birth asphyxia was presumed in outborns when there was a history that the baby had failed to cry or breathe at birth [14]. Apgar score was used in all inborn; a score <7 in the first minute and fifth minute was regarded as asphyxia [14].

Meconium aspiration syndrome was defined as the onset of respiratory distress after birth with meconium-stained body and liquor with or without features of air leaks[15] Chest radiograph may show widespread patchy or homogenous opacities infiltrates and evidence of air trapping[15]. Diagnosis of sepsis was made when the baby had a bacterial growth on blood culture or cerebrospinal fluid culture, respectively [16]. Case Fatality Rate (CFR) = (No. of deathsdue to certain disease/ Total cases from that disease) x100 [17].

# Statistical analysis

The collected data were coded, tabulated, and statistically analyzed using SPSS program and version 17. Data were described using frequencies and percentages. The differences between proportions were tested using chi-square test. A p-value less than 0.05 was considered statistically significant.

# Results

From April 2018 to March 2019, a total of 430 patients were admitted to the NICU. Number of cases presented with respiratory distress was 250, representing 58.1% of all cases admitted. Due to the small percentage of congenital heart disease, congenital anomalies, congenital pneumonia, sepsis, they were combined as the term of others for subsequent analysis.

Table (1) illustrates that babies delivered by CS showed significantly high rates of both RDS and TTN when compared to delivered by vaginal delivery. Prolonged labor were significantly associated with TTN, BA and MAS. None of the other maternal characteristics were associated with causes of NRD.

Table (2) illustrates no significant association between maternal hypertension, maternal diabetes, multiple gestation and causes of NRD.

Table (3) shows no significant association between antepartum hemorrhage, maternal fetal infection, pre-eclampsia and causes of NRD.

Table (4) illustrates that the majority of cases of RDS and TTN were male with statistical significant difference. The only neonatal RD disorders that were associated with low birth weight was RDS. It was found that RDS was significantly associated with prematurity, while TTN and MAS were significantly associated with full term.

Table(5) shows the highest case fatality rate of neonatal respiratory distress diseases were RDS (81.8%) followed by MAS



(56.7%), BA (34.3%), congenital anomalies(25%), CHD (20%) and neonatal sepsis(12.5).

Table (6) summarizes the underlying illnesses of NRD in this study, compared with results from previous studies.

Figure(1) summarizes the most frequent underlying causes for NRD were respiratory distress syndrome(RDS)110(44%), followed by transient tachypnea of the newborn (TTN) 47(18.8%), birth asphyxia (BA) 35(14%), meconium aspiration syndrome (MAS)12(12%), others 28(11.2).

Figure(2) shows that the outcome of NRD was: cure in 40.4% of cases, patients discharged with complications in 10.4 of cases, and death in 49.2% of cases.

	Total	Ma	P-value			
Causes of RD	(N=250)	<35 (200)		>35 (50)		1
		N (%)		N (%)		
RDS	110	90(81.8)			20(18.2)	0.52
TTN	47	36(76.6)			11(23.4)	0.51
BA	35	30(85.7)			5(14.3)	0.36
MAS	30	23(76.7)			7(23.3)	0.62
Others	28	21(75)			7(25)	0.48
	Ì		Pa	rity		P-value
Causes of RD	Total	Primiparous(60)N	1-2	(100)	>3(90)	1
	(N=250)	(%)	N	(%)	N (%)	
RDS	110	30(27.3)	45(	(40.9)	35(31.8)	0.28
TTN	47	14(29.8)	17(	(36.2)	16(34)	0.30
BA	35	8(22.8)	12(34.3)		15(42.9)	0.86
MAS	30	4(13.3)		(56.7)	9(30)	0.15
Others	28	4(14.3)	9(3	32.1)	15(53.6)	0.20
	Total	N	lode of	f delivery	/	P-value
Causes of RD	(N=250)	Vaginal delivery (	73)	Cesarean delivery (177)		
		N (%)			N (%)	
RDS	110	25(22.7)			85(77.3)	0.04
TTN	47	20(42.6)			27(57.4)	0.02
BA	35	15(42.9)			20(57.1)	0.05
MAS	30	5(16.7)		25(83.3)		0.11
Others	28	8(28.6)		20(71.4)		0.93
	Total	F	rolong	ged labor		P-value
Causes of RD	(N=250)	Yes (175) N (%)		No (75) N (%)		1
RDS	110	74(67.3)		36(32.7)		0.40
TTN	47	25(53.2)		22(46.8)		0.006
BA	35	30(85.7)			5(14.3)	
MAS	30	26(86.7)				0.04
Others*	28	20(71.4)			8(28.6)	0.86

Table (1) Polationshi	n of motornal aga	nority	mode of delivery	prolonged labor	and courses of NPD
Table (1): Relationshi	p of maternal age.	, painty,	, mode of derivery.	, prototigeu labor	and causes of MKD



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 \*\*RDS : Respiratory distress syndrome
 TTN : Transient tachypnea of newborn

 \*\*BA: Birth asphyxia
 MAS : Meconium aspiration syndrome

 Others (congenital heart disease, congenital pneumonia, sepsis and congenital anomalies)

Congenital anomalies(diaphragmatic hernias, tracheoesophageal fistula, congenital lobar emphysema)

Table (2): Relationship of maternal hypertension, maternal diabetes, multiple gestation and causes of

NRD

		INKD		
Causes of RD	Total	Maternal h	P-value	
	(N=250)	Yes(15)	No(235)	1
		N (%)	N (%)	
RDS	110	5(4.5)	105(95.5)	0.39
MAS	47	3(6.4)	44(93.6)	0.90
BA	35	2(5.7)	33 (94.3)	0.93
MAS	30	4(13.3)	26(86.7)	0.08
Others	28	1(3.6)	27(96.4)	0.57
Causes of RD	Total	Materna	diabetes	P-value
	(N=250)	Yes(10)	No(240)	
		N (%)	N (%)	
RDS	110	5(4.5)	105(95.5)	0.21
TTN	47	1(2.1)	46(97.9)	0.58
BA	35	0.(0.0)	35 (100)	0.42
MAS	30	2(6.7)	28(93.3)	0.37
Others	28	2(7.1)	26(92.9)	0.32
Causes of RD	Total	Multiple	gestation	P-value
	(N=250)	Yes (6)	No(244)	
		N (%)	N (%)	
RDS	110	3(2.7)	107(97.3)	0.76
TTN	47	2(4.3)	45(95.7)	0.36
BA	35	1(2.9)	34(97.1)	0.84
MAS	30	0(0.0)	30(100)	0.67
Others	28	0(00)	28(100)	0.71

 Table (3): Relationship of antepartum hemorrhage, maternal fetal infection, pre-eclampsia and causes of NRD

		TURE		
Causes of RD	Total	Antepartum	P-value	
	(N=250)	Yes (18)	No(232)	
		N (%)	N (%)	
RDS	110	8(7.3)	102(92.7)	0.96
TTN	47	2(4.2)	45(95.8)	0.39
BA	35	5(14.3)	30(85.7)	0.09
MAS	30	3(10)	27(90)	0.52
Others	28	0(00)	28(100)	0.25
Causes of RD	Total	Maternal fetal infection		P-value
	(N=250)	Yes (15)	N(235)	1
		N (%)	N(%)	
RDS	110	5(4.5)	105(95.5)	0.90
TTN	47	3(6.4)	44(93.6)	0.72



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Birth asphyxia	35	1(2.9)	34(97.1)	0.46
MAS	30	3(10)	27(90)	0.28
Others	28	3(10.7)	25(89.3)	0.23
Causes of RD	Total	Pre-ecl	ampsia	P-value
	(N=250)	Yes (6)	N (244)	
		N (%)	N (%)	
RDS	110	2(1.8)	108(98.2)	0.44
TTN	47	0(0.0)	47(100)	0.73
BA	35	1(2.9)	34(97.1)	0.35
MAS	30	0(0.0)	30(100)	0.99
Others	28	0(00)	28(100)	0.95

 Table (4): Relationship of between sex of neonates, gestational age, birth weight and causes of NRD

	Total		Sex				
Causes of RD	(N=250)	Male (180)		D) Female (70)			
		N (%)		N (%)			
RDS	110	90(81	.8)		20(18.2)	0.002	
TTN	47	28(59	.6)		19(40.4)	0.03	
BA	35	21(6	0)		14(40)	0.09	
MAS	30	18(6	0)		12(40)	0.12	
Others	28	23(82	.1)		5(17.9)	0.211	
			Birth we	eight		P-value	
Causes of RD	Total	<2.5(165)	2.5-3.9(	74)	≥4(11)		
	(N=250)	N (%)	N (%)		N (%)		
RDS	110	80(72.7)	30(27.3	3)	0(0)	0.04	
TTN	47	30(63.8)	15(32)	)	(2)(4.2)	0.72	
BA	35	20(57.2)	11(31.4)		4(11.4)	0.23	
MAS	30	17(56.7)	10(33.3)		3(10)	0.25	
Others	28	18(64.3)	8(28.6)		2(7.1)	0.83	
			Gestation	al ag	e	P-value	
Causes of RD	Total	Preterm<3	Fullterm	37-	Postterm		
	(N=250)	7 w(142)	42(92)	)	>16 (16)		
		N (%)	N (%)		N (%)		
RDS	110	100(90.9)	10(9.1)		0(0.0)	P < 0.001	
TTN	47	2(4.3)	40(85.1)		5(10.6)	P < 0.001	
BA	35	15(42.9)	16(45.7)		4(11.4)	0.24	
MAS	30	10(33.3)	18(60)	)	2(6.7)	0.006	
Others	28	15(53.6)	8(28.6	)	5(17.8)	0.340	



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Table (5): Frequency of causes of NRD and their outcome							
Causes		Immediate outcome	Case	Total			
	Cured N=101	Discharged with complication N=26	Died N=123	fatality rate (%)	N=250		
RDS	15	5	90	81.8	110		
TTN	47	0	0	0	47		
Birth asphyxia	15	8	12	34.3	35		
MAS	11	2	17	56.7	30		
CHD	0	8	2	20	10		
Neonatal sepsis	6	1	1	12.5	8		
Congenital pneumonia	6	0	0	0	6		
Congenital anomalies	1	2	1	25	4		

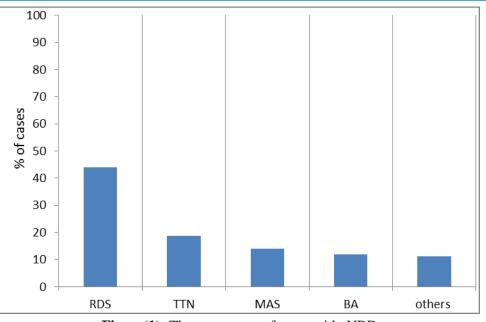
#### Table (5): Frequency of causes of NRD and their outcome

Table (6): Underlying illnesses	of NRD in this study, co	ompared with results from	previous studies

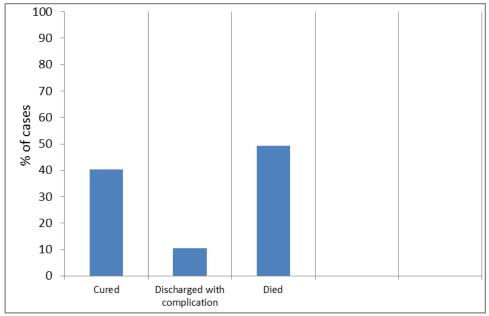
Study	Current study	Abdelrah man et al., (3)	Tochie et al., (4)	Rao and Rao (5)	Abou- Faddan and abdelaziz. (6)	Rijal and Shrestha (7)
Published year	2019	2014	2016	2017	2018	2018
Country	Yemen	Sudan	Cameroon	India	Egypt	Nepal
Number of cases	250 (52.9%)	100 (56.5%)	334 (47.5%)	200 (13.3%)	487 (52.9%)	109 (34.3%)
RDS	110 (44%)	15 (15%)	47 (14%)	64 (32%)	223 (45.8%)	13 (11.9%)
TTN	47 (18.8%)	28 (28%)	83 (25%)	18 (9%)	83 (17%)	17 (15.6%)
Birth asphyxia	35 (14%)	-	27 (8%)	37 (18.5%)	-	13 (11.9%)
MAS	30 (12%)	6(6%)	37 (11%)	70 (35%)	9 (1.8%)	23 (21.1%)
Others						
CHD	10 (4%)	9 (9%)	-	7 (3.5%)	-	7 (6.4%)
Neonatal sepsis	8 (3.2%)	24 (24%)	103 (31%)	-	-	18 6.5%)
Congenital pneumonia	6 (2.4%)	18 (18%)	37 (11%)	2 (1%)	86 (17.7%)	2 (1.8%)
Congenital anomalies	4 (1.6%)	18 (18%)	37 (11%)	2 (1%)	86 (17.7%)	2 (1.8%)



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Figure(1): The percentage of cases with NRD



Figure(2): Frequency of immediate outcome of NRD

### Discussion

We had observed that, the most frequent underlying cause for RD in children was RDS 44% of cases, followed by TTN 18.8%, birth asphyxia 14%, MAS (12%), others 11.2% (CHD 4%, sepsis 3.2%, congenital pneumonia 2.4%, and congenital anomalies 1.6%. Compared to previous studies [3-7], our results showed some similarities and



differences. This wide variation in proportion of causes of respiratory distress among various studies may be due to antenatal care in that community, availability of obstetricians including trained birth attendants, use of preventive measures during antenatal period, proportion of term and preterm deliveries, non-uniform inclusion criteria, various other factors such as status of community and type of facilities in institution [18]. The first important cause of respiratory distress in this study was RDS, which was found to be 110(44%). Similarly, RDS was found to be the first cause of respiratory distress with an proportion of 45.8% by Abou-Faddan and Abdelaziz [6] and Sabzehei et al., (36.6%) [19]. low proportion were found by Rijal and Shrestha (11.9%) [7], abdulrahaman et al., (15%) [3] and Tochie et al., (14%) [4] .This can be explained by a higher proportion of neonates in our study were premature. In the current study, it was observed that cesarean delivery was significantly higher in RDS (77.3% versus 22.7% for babies with normal vaginal delivery ,p=0.04). This is in agreement with the results of other studies [7,20,21]. Neonates born by cesarean section have a larger residual volume of lung fluid, secrete less surfactant to the alveolar surface and have a delayed clearance of lung fluid[22]; thus, they are at higher risk of developing RDS. Male babies had higher rates of RD in RDS (81.8% versus 18.2% for females, p=0.002). This is consistent with several studies [20,21]. Differences in hormonal regulation of lung development provide

candidate mechanisms to account for an increased risk of RDS associated with male sex [23]. Androgens delay lung fibroblast secretion of fibroblast pneumocyte factor, which can delay the development of alveolar type II cells; furthermore, they reduce the release of surfactant. On the contrary, estrogens promote the synthesis of surfactant, including phospholipids, lecithin and surfactant proteins A and B, and improve fetal lung development by increasing the number of alveolar type II cells and the synthesis of lamellar bodies[24].

In the current study, it was observed that prematurity was significantly higher in RDS (90.9% versus 9.1% for babies with term, p < 0.001). This is in agreement with the of other studies results [6.7.20.21].Respiratory distress syndrome (RDS) occurs primarily in premature infants; its incidence is inversely related to gestational age and birth weight. Surfactant deficiency (decreased production and secretion) is the primary cause of RDS [12]. In the current study, it was also observed that low birth was significantly higher in RDS weight (72.7% versus 27.3% for babies with normal weight p=0.04) [6,7,20]. The premature and low-birth-weight neonates usually have pulmonary immaturity and limited respiratory muscle strength. It is important to make a diagnosis and find the etiology to provide appropriate management to prevent preterm delivery [24]. Transient tachypnea of newborn was the second most common cause of respiratory distress found in 47(18.8%) with similar proportion as reported by other



studies [6,7]. But, kommawar et al., found TTN was the most common cause of respiratory distress in newborns with proportion of 40%.[25]. In the study done by Sodawat et al., a transient tachypnea of newborn was found to be the commonest of respiratory distress in newborns with incidence of (32.6%) [18].

In the current study, it was observed that cesarean delivery was significantly higher in TTN (57.4% versus 42.6% for babies with normal vaginal delivery, p=0.02). This is consistent with previously reported findings [7,26]. These have been attributed to delayed or abnormal fetal lung fluid clearance due to the absence of the hormonal changes that accompany spontaneous labor [27]. Transient tachypnea of newborn is believed to be secondary to slow absorption of fetal lung fluid, resulting in decreased pulmonary compliance and tidal volume and increased dead space [13]. It was also observed that prolonged labor was significantly higher in TTN (53.2% versus 46.8% for babies with normal labor, p=0.006), which is consistent with other results [26,27,28]. Prolonged labor may cause dysfunctional catecholamine regulation, mild pulmonary capillary leak and myocardial dysfunction [28]. In the current study, it was observed that male gender was significantly higher in TTN (59.6% versus 40.4% for female, p=0.03). This is in agreement with the results of other studies [26,27]. It was also observed that full term was significantly higher in TTN, which is consistent with other results [6,26]. Birth asphyxia constituted 35(14%) of the total

respiratory distress cases in our study. A proportion of birth asphyxia of 12.2% and 11.9% were found by konmnawar et al., [25] and Rijal and Shrestha [7] respectively. Very low proportion were found by Sabzehei et al., (1.1%)[19]. It was observed that prolonged labor was significantly higher in BA (85.7%) versus 14.3% for babies with normal labor, P=0.03). This was agreement with other studies [29,30]. It is clear that when labor is prolonged, there is a high probability for the fetus to become distressed[31] Meconium aspiration syndrome constituted 30(12%) of the total respiratory distress cases in our study. Proportion of MAS of 9.3% and 11% were found by Sodawat et al., [18] and Tochie et al. [4] respectively. Very high proportions were found by Rao and Rao (35 %) [5]. It was observed that prolonged labor was significantly higher in MAS (86.7% versus 13.3% for babies with normal, p=0.04). This is consistent with previously reported findings [32,33]. Probably because this is the referral hospital catering more than 250 km radial distance hence the referred patient took longer time to reach the hospital. It was also observed that full term was significantly higher in MAS, which is consistent with other results [33,34]. This was related to maturity and increased levels of intestinal hormone, motilin with increasing gestational age [34]. The immediate outcome of NRD is cure in 40.4%, patients discharged with complications are 10.4% and death occurred in 49.2% of cases. In a study done by Abdelrahman et al.,[3] the immediate outcome of NRD is cure in 56%, patients



discharged with complications are 8% and death occurred in 36% of cases. Sabzehei et al., [19] reported that 19.3 of infants with NRD died of the disease. Sodawat et al., [18] found that overall death among cases of respiratory distress was 28%. But, Rao and Rao [5] found a very low death rate among NRD (2.5%). Comparison between the causes and their fatalities in this study revealed that hyaline membrane disease has case fatality rate of (81.8) followed by MAS (56.7%) and BA (34.3%). This result is consistence with other studies [5,6,10]. In a study done by Sodawat et al., the fatality rate were 47.58% for RDS, 42.24% for sepsis, 35.71% for MAS, 35.55% for HIE and 33.33% for cases of CHD[18]. Kommawar et al.,[25] reported the highest case fatality rate were 61.6% for HMD, 17.4% for BA and 8.13% for MAS. The high fatality rate can be explained by the higher percentages of preterm and LBW, inadequate antenatal care, less use of steroid in prematurity antenatal, less meticulous management of high risk pregnancies. In addition to our study was conducted in children hospital, which represents the reference hospital for neonates of high risk pregnancies and complicated deliveries and lack of surfactant replacement therapy in our hospital and mechanical ventilation. In conclusions respiratory distress was the major cause of admission in our NICU. The most common causes of respiratory distress were RDS, perinatal asphyxia, and MAS. The common cause of death was RSD. RDS is significantly correlated with cesarean section, prematurity,

male sex and low birth weight, while prolonged labor were significantly associated with TTN, BA and MAS. From this study we may recommend the followings; early detection and appropriate management of risk factor is essential to ensure better outcome in all infants presenting with respiratory distress. Good antenatal care with prompt management of high risk pregnancies. Early recognition of potential risk factors for respiratory distress will be helpful in decreasing the burden associated with neonatal respiratory distress.

Proper training of physicians in the management of neonatal respiratory distress, including availability of equipped NICU with ventilator and surfactant facilities.

### **Conflict of interest**

The research protocol and the questionnaires were conducted according to principles of the Declaration of Helsinki, as well as reviewed and approved by the college ethical research committee. Verbal consents were also taken from the parents and caregivers of children involved in the study.

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