## Audit on the Application of Surfactant in Premature Babies in Assiut University Children Hospital

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#### Abstract

*Background:* To evaluate the use of exogenous natural surfactant for Respiratory Distress Syndrome (RDS) in premature babies, identify areas where European Consensus Guidelines on the Management 2013 Update, modified by The Egyptian Consensus on the Management of Neonatal Respiratory Distress Syndrome in Preterm Infants 2014 are applied to as a reference standard in Assiut University Children Hospital and identify those points to be targeted for improvement.

*Patients and Methods:* A retrospective audit was undertaken over a 12-month period between 1 st of January, 2016 to the 31 st of December 2016. It included all babies less than 37 weeks gestation with respiratory distress syndrome who received surfactant as a rescue treatment, guided by European Consensus Guidelines on the Management 2013 Update, modified by The Egyptian Consensus on the Management of Neonatal Respiratory Distress Syndrome in Preterm Infants 2014 and admitted to Neonatal Intensive Care Unit of Assiut University Children Hospital during this period.

*Results:* All cases with severe respiratory distress that needed surfactant injection and admitted in Assiut university children hospital from 1 st of January to 31<sup>th</sup> of December 2016 were included in the study. The study included 50 patients with severe respiratory distress syndrome.

*Conclusion:* Surfactant replacement therapy has been available for about 37 years, revolutionizing neonatal respiratory care after its introduction in the 1980s. Along with antenatal steroids, surfactants improve survival for preterm babies and they are now recommended routinely as early in the course of Respiratory Distress Syndrome (RDS) as possible. The present study aimed to evaluate the use of surfactant for Respiratory Distress Syndrome (RDS) in premature babies admitted at NICU of Assiut university children hospital during the period from 1 st of January to 31 st of December 2016 using European Consensus Guidelines on the Management 2013 Update, modified by The Egyptian Consensus on the Management of Neonatal Respiratory Distress Syndrome in Preterm Infants 2014.

The study included 50 cases with respiratory distress syndrome who received surfactant therapy.

- Data of the study showed that treatment regimens for cases of RDS in Assiut University Children Hospital partially followed the reference standard of the study concerning application of surfactant in premature babies.
- Data of the gestational age and weight and sex were recorded in 100% of cases.
- Data concerning clinical evaluation of the studied group were recorded in 100% cases.
- Data concerning obstetric history and prenatal care were fulfilled well but there were no records concerning receiving a course of antibiotics if there is preterm, pre-labor rupture of membranes nor short term tocolytic drugs. Also, the data showed that only 28% of mothers received a course of prenatal steroids if preterm labor was expected.
- Data concerning delivery room stabilization were fulfilled well but there were no records concerning delayed cord clamping and there was no available CPAP at the delivery room nor plastic bag for stabilization of preterm baby with respiratory distress syndrome.
- Data concerning surfactant therapy were fulfilled well but there was no data that any case received prophylactic surfactant therapy neither with nor without INSURE technique.
- Data concerning respiratory support were fulfilled well but no cases received CPAP as a 1 st line respiratory support for any spontaneously breathing preterm with RDS at delivery room and no cases received CPAP with early rescue surfactant which is now considered the optimal management for babies with RDS.
- Data concerning supportive care were fulfilled in 100% of cases but no cases received enteral feeding from the 1 st day which is strongly recommended with careful fluid balance for early aggressive nutritional support.
- Data concerning prognosis showed that the vast majority of dead cases were below 32 weeks and below 1KG.
- Key Words: Antenatal corticosteroids Continuous positive airway pressure – Meconium aspiration syndrome – Nasal intermittent positive pressure ventilation – Necrotizing enterocolitis – Patent ductus arteriosus – Respiratory distress syndrome.

Abbreviations:

ACS : Antenatal Corticosteroid. HMD: Hyaline Membrane Disease.

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### Introduction

**IN 1959**, Avery and Mead reported on the deficiency of surface-active material in the lungs of preterm babies with respiratory distress syndrome (RDS). This led to clinical trials of artificial surface-active materials in babies with RDS.

The term "hyaline membrane disease" refers to the histological aspect of the most frequent pulmonary pathology in preterm newborn patients [7].

Surfactant deficiency in the immature lungs causes alveolar instability and collapse, capillary edema and the formation of hyaline membrane. Thus, the hyaline membranes are epiphenomena and are not the cause of respiratory failure in infants with immature lungs. This definition is presently used to indicate surfactant deficit alone and should not be used for other causes of respiratory distress, Clinicians prefer to talk of "respiratory distress syndrome" (RDS) [1].

Improvement in neonatal treatment has changed the natural course of the illness, its clinical and radiological features and has enabled extremely low birth weight newborns (ELBW) to survive, Alveoli paucity and pulmonary interstitial thickness in ELBW impair gas exchange and may necessitate prolonged ventilation treatment, increasing the risk of ventilator-induced lung injury (VILI) and bronchopulmonary dysplasia [13].

RDS, therefore, is a complex illness where pulmonary immaturity and surfactant deficit play a role together with other pathological conditions that determine the course of the illness and both short and long-term results.

RDS is usually defined by the presence of acute respiratory distress with disturbed gas exchange in a preterm infant with a typical clinical course or X-ray (ground glass appearance, air bronchograms and reduced lung volume [12].

The lungs of preterm babies with RDS are both anatomically and biochemically immature; they neither synthesize nor secrete surfactant well.

Surfactant normally lines the alveolar surfaces in the lung, thereby reducing surface tension and preventing atelectasis [8]

Surfactant replacement therapy, either as a rescue treatment or a prophylactic natural surfactant therapy, reduces mortality and several aspects of morbidity in babies with RDS. These morbidities include deficits in oxygenation, the incidence of pulmonary air leaks (pneumothorax and pulmonary interstitial emphysema) and the duration of ventilatory support.

Surfactant replacement increases the likelihood of surviving without bronchopulmonary dysplasia (BPD, also known as chronic lung disease of the preterm) [2].

Babies treated with surfactants have shorter hospital stays and lower costs of intensive car treatment compared with randomized control infants receiving no surfactants.

Surfactant replacement therapy has been available for about 25 years, revolutionizing neonatal respiratory care after its introduction in the 1980s. Along with antenatal steroids, surfactants improve survival for preterm babies and they are now recommended routinely as early in the course of Respiratory Distress Syndrome (RDS) as possible [1]. Prophylactic treatment, although appearing ideal, exposes some babies who might manage perfectly well on Continuous Positive Airway Pressure (CPAP) to intubation and ventilation, which may increase the risk of bronchopulmonary dysplasia. Recent studies attempt to determine the optimal balance between avoiding ventilation by using CPAP and giving surfactant in a timely fashion to babies with RDS. The key feature of Respiratory Distress Syndrome (RDS) is the insufficient production of surfactant in the lungs of preterm infants [11] As a result, researchers have looked into the possibility of surfactant replacement therapy as a means of preventing and treating RDS. We sought to identify the role of surfactant in the prevention and management of RDS, comparing the various types, doses, and modes of administration, and the recent development [9].

Natural, or animal-derived, surfactant is currently the surfactant of choice in comparison to protein-free synthetic surfactant. However, it is hoped that the development of protein-containing synthetic surfactant, such as lucinactant, will rival the efficacy of natural surfactants, but without the risks of their possible side effects. Administration techniques have also been developed with nasal Continuous Positive Airway Pressure (nCPAP) and selective surfactant administration being now recommended, multiple surfactant doses have also reported better outcomes [10]. An aerosolized form of surfactant is being trialed in the hope that surfactant can be administered in a non-invasive way [3]. Overall, the advancement, concerning the structure of surfactant and its mode of administration,

offers an encouraging future in the management of RD.

## **Patients and Methods**

All cases with severe respiratory distress that needed surfactant injection and admitted in Assiut university children hospital from 1 st of January to 31 th of December 2016 were included in the study. The study included 50 patients with severe respiratory distress syndrome.

*Inclusion criteria:* All the recorded premature cases with Respiratory Distress Syndrome (RDS) admitted to neonatal intensive care unit during aforementioned period that received surfactant therapy and referred from Hospital of Women's Health in Assiut were included.

*Exclusion criteria:* All the recorded cases during aforementioned period who were admitted to neonatal unit with any other type of respiratory distress rather than respiratory distress syndrome were excluded.

*Tools of study:* Reviewing sheets of patient with severe respiratory distress admitted to Assiut University Children Hospital during the study duration.

The following data were collected and recorded for each patient in a master sheet for management of RDS by Surfactant:

- Socio-demographic characteristics such as name, gestational age and sex.
- Data about clinical evaluation of a neonate with respiratory distress using Downes' score as air entry, retractions, grunting and cyanosis.
- Data about former, present obstetric history (maternal age, fetal distress and perinatal asphyxia multiple gestation, prior preterm delivery, maternal diabetes and mode of delivery and prenatal care received by women at high risk of preterm birth as being transferred out to perinatal centers with experience in management of RDS and receiving antenatal steroids or tocolytics and antibiotics if there is history of PROM or not.
- Data about delivery room stabilization as cord clamping, stabilization by 21 %-30% oxygen, using CPAP, and stabilization under radiant warmer.
- Data about Surfactant therapy as it's nature either natural or synthesized, if we use prophylactic surfactant therapy, type of surfactant used, if we use INSURE technique and if repeated doses of surfactant needed.

- Data about further respiratory support as use of oxygen, CPAP, MV and caffeine.
- Data about prognosis: To identify areas where European Consensus Guidelines on the Management 2013 Update, modified by The Egyptian Consensus on the Management of Neonatal Respiratory Distress Syndrome in Preterm Infants 2014 are applied to as a reference standard in Assiut University Children Hospital and identify those points to be targeted for improvement.

## Results

Table (1) showed that the majority of cases that needed surfactant injection were males (72%), very preterm babies (28 to <32 weeks gestational age) (72%) and VLBW <1.5 KG (48%).

Table (2) showed evaluation of respiratory distress using downes' score, it showed that the majority of cases that needed surfactant injection were distressed with Downe score 4-7 (80%).

Table (3) showed that the majority of cases that needed surfactant injection had a previous history of prior preterm delivery (44%) and most of them delivered by elective CS (76%).

The table also showed that only 28% of the mothers of theses neonates received a prophylactic course of steroids prior to delivery.

Despite the importance of prenatal care as a cornerstone of a healthy pregnancy, labor, and baby, no sufficient information could be received especially about administration of antenatal corticosteroids.

Being with or without tocolytics and antibiotics that will affect the decision making for use of surfactant in premature babies <30 wks. Accurate and complete documentation in the medical record is lacking. Most of information was inaccurate, insufficient and derived from the parents.

Table (4) showed that there was lack of facilities at the delivery room as there was no available CPAP nor surfactant that could be used which delayed the management of cases of RDS which worsened the prognosis.

Table (5) showed that no cases received prophylactic surfactant therapy which is of great benefit to extremely preterm infants in whom the mother has not had antenatal steroids or those who require intubation for stabilization.

The table also showed that only 4% of cases received repeated doses of surfactant inspite of

presence of signs of respiratory distress even after injection in some cases.

INSURE technique was not used for rescue surfactant administration.

Table (6) showed that only 16% of cases that needed surfactant injection were connected to CPAP which should be started from birth in all babies at risk of RDS, such as those <30 weeks' gestation who do not need MV, until their clinical status can be assessed.

The table also showed that there were no cases received CPAP with early rescue surfactant which considered the optimal management for babies with RDS.

Table (7) showed that no cases received enteral feeding on day 1 of life but Careful fluid balance was required with early aggressive nutritional support using parenteral nutrition whilst enteral feeding was being established.

Table (8) Showed that the vast majority of dead cases were below 32 weeks and below 1 KG.

Table (1): Demographic data of the studied group.

Item	Ν	%	<i>p</i> -value
Gestation at Birth:			
28 to <32 weeks	36	72	_
32 to <34 weeks	6	12	_
34 to $<$ 37 weeks	8	16	-
Birth Weight (kilograms):			
<1.0 kg	14	28	_
<1.5 kg	24	48	_
1.5: 2.499 kg	12	24	_
Sex:			
Male	36	72	< 0.001**
Female	14	28	< 0.001*

\*: Statistically significant difference (*p*<0.05). \*\*: Statistically significant difference (*p*<0.01).

Item	Ν	%	<i>p</i> -value
Air entry:			
Good bilateral	12	24	< 0.001**
Mild Decrease in Air entry	8	16	< 0.001**
No Air Entry	30	60	0.046*
Respiratory rate:			
<60	0	_	_
60-80	12	24	< 0.001**
>80	38	76	< 0.001**
Retractions:			
No Retractions	44	88	< 0.001**
Mild Retractions	2	4	< 0.001**
Severe Retractions	4	8	< 0.001**
Grunting:			
No Grunting	10	20	< 0.001**
Audible by stethoscope	7	14	< 0.001**
Audible with ear	33	66	0.002**
Cyanosis:			
No Cyanosis	42	84	< 0.001**
Cyanosis relieved by $O_2$	0	_	_
Cyanosis on $O_2$	8	16	< 0.001**
Downe' score:			
4-7	40	80	< 0.001**
>7	10	20	< 0.001**
<4	0	_	_

Table (2): Evaluat	tion of respirator	y distress using	downes' score	
1 able (2). Evaluat	tion of respirator	y distress using	downes score.	

Item	Ν	%	<i>p</i> -value
Maternal age:			
<16 years	8	16	< 0.001**
16 - <35 years	28	56	0.005**
>35 years	14	28	< 0.001**
Fetal distress and perinatal asphyxia	14	28	<0.001**
Mutiple gestation	4	8	< 0.001**
Prior preterm delivery	22	44	< 0.001**
Maternal diabetes	10	20	< 0.001**
Delivery mode:			
- Elective and emergency CS	38	76	< 0.001**
- Vaginal delivery	10	20	< 0.001**
- Precipitous labor	2	4	< 0.001**
Prenatal care:			
Transferred out to perinatal centers with experience in management of RDS	50	_	_
Received a course of prenatal steroids if gestation between 23 and 34 weeks.	14	28	<0.001**
Received antibiotics if preterm, pre-labour rupture of membranes.	Ν	o available rec	cords
Received short-term tocolytic drugs to allow completion of a course of prenatal corticosteroids.	Ν	o available rec	cords

Table (4): Delivery room stabilization among the studied cases.

tem			<i>p</i> -value
Cord clamping delayed for at least 30 seconds (if possible)	No available records		
Stabilisation initiated in 21-30% oxygen using a blender	50	100	-
Applied pulse oximetry to the right wrist from birth	50	100	-
Spontaneously breathing baby stabilised with CPAP	No available CPAP facility at delivery room		
If required intubation, received surfactant	0	_	-
If <28 weeks gestation, delivered into a plastic bag	No available facility		
Servo controlled babies stabilized under a radiant warmer within 10min	50	100	-

Table (5): Surfactant Therapy among the studied cases.

Item	Ν	%	<i>p</i> -value
Received a natural surfactant preparation (if needed)	50	100	_
Prophylactic surfactant therapy:			
- In the delivery suite for extremely preterm infants in whom the mother has not		o facilities to injec	
had antenatal steroids or those $\leq 31$ who require intubation for stabilization	Surfactant a	t delivery room a	is
Early rescue surfactant administered:	a proph	ylactic therapy	
Babies with RDS if:			
<26 weeks FiO <sub>2</sub> $>30%$	0	_	-
>26 weeks and FiO <sub>2</sub> $>40%$	50	100	-
Type of surfactant administered:			
Poractant alfa (Curosurf) in an initial dose of 100 mg/kg 200mg/kg	We only us	e natural Survant	a
Beractant Beractant (Survanta) in an initial dose of 100mg/kg	(Beractant)a	at a dose of 4ml/k	g
Bovactant (Alveofact) in an initial dose of 50mg/kg			
Received a rescue dose of 200mg/kg surfactant	0	_	_
INSURE technique used for rescue surfactant administration.	0		
Repeat doses of surfactant given in ongoing evidence of RDS, e.g. O <sub>2</sub> requirement/need for MV	2	4	<0.001**
After surfactant given, rapid reduction in administered FiO <sub>2</sub> documented	50	100	_

Table (6): Further Respiratory Support among the studied cases.

Item	Ν	%	<i>p</i> -value
Oxygen Indications:			
Severe respiratory distress	40	80	< 0.001**
Low O2 saturation in air	50	100	_
Target saturations [SaO 2] 90-95% in preterm infants	50	100	
Non-Invasive Respiratory Support:			
CPAP first-line respiratory support used for RDS, if not	No available e	arly CPAP facility	
intubated	at deli	very room	
CPAP delivered through mask or bi-nasal prongs	8	16	< 0.001**
CPAP pressure of at least 6 cm of water applied	8	16	< 0.001**
CPAP with early rescue surfactant	0	-	_
Mechanical Ventilation Strategies:			
Targeted tidal volume ventilation If not used, what is documented reason why? (E.g. high leak, used. another lung pathology etc.)	Our machines deliver only Pressure controlled ventilation		
Caffeine used in baby if; apnoea, or, facilitate weaning from MV	50	100	_
Tapering course of steroids (dexamethasone) used if remains on MV after 1-2 weeks	8	16	<0.001**

Table (7): Supportive care among the studied cases.

Item	Ν	%	<i>p</i> -value
Body temperature Maintained in the normal range	50	100	_
Antibiotics started until sepsis has been ruled out (unless the risk of infection is low, for example after an elective caesarean section	50	100	_
Parenteral nutrition started on day 1 of life	50	100	_
Minimal enteral feeding/trophic feeds started on day 1 of life	0	_	_
Blood pressure monitored regularly, aiming to maintain normal tissue perfusion, if necessary usinginotropes	50	100	_
Hb conc. maintained within normal range	50	100	_
PDA, if needing treatment, is medically managed (ibuprofen/indomethacin)	13	100	_

Table (8): Prognosis of the studied Group of RDS.

Item	Ν	%	<i>p</i> -value
Living cases according to Gestational age:			
28 to $< 32$ weeks	2	20	0.007**
32 to $<$ 34 weeks	4	40	0.371
34 to < 37 weeks	4	40	0.371
Living cases according to Birth Weight:			
< 1.0 kg	-	_	-
<1.5 kg	4	40	0.371
1.5: 2.499 kg	6	60	0.371
Dead cases according to Gestational age:			
28 to $<$ 32 weeks	34	85	< 0.001**
32 to $<$ 34 weeks	2	5	< 0.001**
34 to < 37 weeks	4	10	< 0.001**
Dead cases according to Birth Weight:			
< 1.0 kg	14	35	0.007**
<1.5 kg	20	50	1
1.5: 2.499 kg	6	15	< 0.001**

#### Discussion

Surfactant replacement therapy has been available for about 25 years, revolutionizing neonatal respiratory care after its introduction in the 1980s. Along with antenatal steroids, surfactants improve survival for preterm babies and they are now recommended routinely as early in the course of Respiratory Distress Syndrome (RDS) as possible. Prophylactic treatment, although appearing ideal, exposes some babies who might manage perfectly well on Continuous Positive Airway Pressure (CPAP) to intubation and ventilation, which may increase the risk of bronchopulmonary dysplasia. Recent studies attempt to determine the optimal balance between avoiding ventilation by using CPAP and giving surfactant in a timely fashion to babies with RDS. The key feature of Respiratory Distress Syndrome (RDS) is the insufficient production of surfactant in the lungs of preterm infants. As a result, researchers have looked into the possibility of surfactant replacement therapy as a means of preventing and treating RDS [4].

We sought to identify the role of surfactant in the prevention and management of RDS, comparing the various types, doses, and modes of administration, and the recent development.

Natural, or animal-derived, surfactant is currently the surfactant of choice in comparison to protein-free synthetic surfactant [5]. However, it is hoped that the development of protein-containing synthetic surfactant, such as lucinactant, will rival the efficacy of natural surfactants, but without the risks of their possible side effects. Administration techniques have also been developed with nasal Continuous Positive Airway Pressure (nCPAP) and selective surfactant administration now recommended, multiple surfactant doses have also reported better outcomes. An aerosolized form of surfactant is being trialed in the hope that surfactant can be administered in a non-invasive way. Overall, the advancement, concerning the structure of surfactant and its mode of administration, offers an encouraging future in the management of RD.

It is very important to receive proper prenatal care as prenatal care is a cornerstone of a healthy pregnancy, labor, and baby [6].

Interventions to prevent RDS should begin before birth and involve both pediatricians and obstetricians as part of the perinatal team. The care team must be well attuned as prevention is better than a cure [14].

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Enhance communication among healthcare team members that influences the quality of working relationships, job satisfaction and has profound impacts on patient safety through accurate and complete documentation in the medical record especially the administration of antenatal corticosteroids (with or without tocolytics) that will affect the decision making for use of surfactant in premature babies <37 weeks [15]

Delivery room should be equipped with Facilities as Plastic bags to prevent hypothermia, CPAP machine or T-piece and Surfactant for use once indicated [16].

If possible, delay clamping of the umbilical cord for at least 30 s with the baby held below the mother to promote placento-fetal transfusion.

Try to initiate Early CPAP and selective surfactant administration to avoid intubation and reduce rates of death.

An experienced neonatal resuscitation/stabilization team is essential for surfactant administration [17].

Select the type of surfactant in terms of efficacy, safety, and total number of doses per patient and cost.

MV can be avoided by using the 'INSURE' (Intubate-SURfactant-Extubate to CPAP) technique

CPAP should be started from birth in all babies at risk of RDS, such as those <30 weeks' gestation who do not need MV, until their clinical status can be assessed [18].

CPAP with early rescue surfactant should be considered the optimal management for babies with RDS.

MV should be used to support babies when other methods of respiratory support have failed [19]

Minimal enteral feeding should also be started from the first day.

Careful fluid balance is required with early aggressive nutritional support using parenteral nutrition whilst enteral feeding is being established

Adherence to guidelines as a reference standard if possible to improve the prognosis [20].

Consideration should be given to pharmacological closure of the ductus arteriosus.

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# دراسة تدقيقية (أستخدام السيرفاكتانت في الأطفال الخدج) في مستشفى الأطفال الجامعي بأسيوط

أستخدام المادة الفعالة للسطح في علاج متلازمة الغشاء الزجاجي أتيح منذ ٣٧ عام مما أحدث ثورة عام ١٩٨٠.

أدت المادة الفعالة للسطح مع الكورتيزون فيما قبل الولادة إلى احسين نجاة الأطفال الخدع التى تعانى من متلازمة الضائقة التنفسية. نتيجة متلازمة الغشاء الزجاجي.

أهداف البحث: تقييم مدى إلتزام الأطباء بالبروتو كول العلاجى لحالات متلازمة الضائقة التنفسية نتيجة متلازمة الغشاء الزجاجى بإستخدام المادة الفعالة للسطح والمحجوزين بوحدة الأطفال حديثى الولادة بمستشفى الأطفال الجامعى بأسيوط خلال الفترة من اول يناير ٢٠١٦ الى ٣١ ديسمبر ٢٠١٦ بإستخدام المعايير الموضوعة من قبل الإرشادات التوافقية للمنظمة الاوروبية لعلاج متلازمة الضائقة التنفسية عند الخدع نتيجة متلازمة الغشاء الزجاجى – تحديث ٢٠١٣ معدلة بواسطة الإرشادات التوافقية للمنظمة الاوروبية متلازمة الضائقة التنفسية عند الخدع نتيجة متلازمة الفشاء الزجاجى – تحديث ٢٠١٣ معدلة بواسطة الإرشادات التوافقية للمنظمة المنظمة المصرية لعلاج

الأطفال المستهدفين: كل الأطفال الخدع الذين يعانون من متلازمة الضائقة التنفسية نتيجة متلازمة الغشاء الزجاجى المحجوزين بوحدة الأطفال حديثى الولادة بمستشفى الأطفال الجامعى بأسيوط فى الفترة من ١/١//١١ إلى ١٢/٦/ / ٢٠١٦.

الأطفال المستبعدين نكل الأطفال الخدع الذين يعانون من متلازمة الضائقة التنفسية نتيجة لسبب غير متلازمة الغشاء الزجاجي

النتائج: تتضمن الرسالة ٥٠ حالة يعانون من متلازمة الضائقة التنفسية نتيجة متلازمة الغشاء الزجاجي عدد الذكور ٣٦ وعدد الإناث ١٤ حالة.

وأظهرت بيانات الدراسة أن الأنظمة العلاجية المستخدمة فى علاج متلازمة الضائقة التنفسية نتيجة متلازمة الغشاء الزجاجى اتبعت جزئياً المعايير المرجعية للدراسة.

- تم أستيفاء البيانات الخاصة بالعمر لبجيني والنوع والوزن في ١٠٠٪ من الحالات
- تم أستيفاء البيانات الخاصة بالتقييم الأكلينيكي للحالات في ١٠٠٪ من الحالات.
- تم أستيفاء البيانات الخاصة بتاريخ الحمل ومراعاة ما قبل الولادة فى أغلب الحالات ولكن لم يتم السؤال عن إعطاء مضادات حيوية فى حالات تمزق الأغشية كما لم يتم السؤال عن إذا كانت الأم العرضة للولادة قبل الميعاد قد أخذت أدوية مثبطة للانقباضات الرحمية كما أن النتائج أوضحت أن ٢٨٪فقط من الأمهات قد أخذوا كورتيزون ما قبل الولادة.
- تم أستيفاء البيانات الخاصة بأستقرار الجنين فى غرفة الولادة فى معظم الحالات ولكن لا توجد تسجيلات كتابية تدل على تأخير ربط الحبل السرى للجنين أو إستخدام جهاز الضغط الهوائى الإيجابى المستمر فىغرفة الولادة.
- تم أستيفاء البيانات الخاصة بإستخدام المادة الفعالة للسطح و لكن لم يتم إستخدامه كأجراء وقائى للأطفال الخدع الذين يعانون من متلازمة الضائقة التنفسية نتيجة متلازمة الغشاء الزجاجي.
- تم أستيفاء البيانات الخاصة بالدعم التنفسى للأطفال الخدع بشكل جيد حيث تم وضع كل الحالات على أكسچين لكن لم يتم إستخدام جهاز الضغط الهوائى الإيجابى المستمر أو حقن المادة الفعالة للسطح فى غرفة الولادة.
- تم أستيفاء البيانات الخاصة بالرعاية الداعمة ولكن لا توجد حالات أخذت تغذية معوية وهي متصلة على جهاز التنفس الصناعي.
- تم أستيفاء البيانات الخاصة بنتائج العلاج مما أوضح أن معظم حالات الوفاة كانت لأطفال عمرها الجنينى أقل من ٣٢ أسبوع وأقل من ١كجم٠