

# Surfactant Replacement Therapy for Neonates with Respiratory Distress

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			E.g. New Policy, Formal review, Informal review due to...
1	Apr 2018	Dr S Narayanan	New network guideline
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## Table of Contents

Purpose of the Guideline .....	4
Target Audience.....	4
Introduction & Background .....	4
Definitions.....	4
Prophylactic surfactant therapy: .....	4
Early rescue surfactant therapy:.....	5
LISA/MIST: .....	5
INSURE:.....	5
Indications, preparation and dosage: .....	5
Surfactant administration procedure:.....	7
Adverse effects .....	8
Post procedure care.....	8
Audit standards .....	9
References.....	9

## GLOSSARY OF TERMS

TERM	Gestation $\geq$ 37 completed weeks
PRETERM	Gestation < 37 completed weeks
Moderate/Late PRETERM	Gestation between 32 – 36 weeks
Very PRETERM	Gestation between 28 – 32 weeks
Extreme PRETERM	Gestation < 28 weeks
RDS	Respiratory Distress Syndrome
NICU	Neonatal Intensive Care Unit
LNU	Local Neonatal Unit
SCU	Special Care Unit
PH	Pulmonary Haemorrhage
INSURE	<b>IN</b> tubate, <b>SUR</b> factant and <b>Ext</b> ubate
LISA	<b>L</b> ess <b>I</b> nvasive <b>S</b> urfactant <b>A</b> dministration
MIST	<b>M</b> inimally <b>I</b> nvasive <b>S</b> urfactant <b>T</b> herapy

## Purpose of the Guideline

To provide guidance on surfactant administration in preterm and term neonates who either present with, or are at high risk of developing, surfactant deficiency respiratory distress syndrome. The document outlines the indications, dosage, formulation and procedure for surfactant administration & timing/strategies for surfactant use. It also discusses the post procedure management; complications and sets out audit standards for monitoring compliance.

## Target Audience

This guideline is intended for the use of neonatal medical and nursing staff in birthing environments and neonatal unit. Very rarely for a preterm neonate presenting to Children's Emergency department following an out of hospital birth.

## Introduction & Background

Respiratory distress syndrome (RDS) due to surfactant deficiency (primary deficiency) is a major cause of morbidity and mortality in preterm infants [1]. Secondary surfactant deficiency complicates other respiratory disorders in neonates such as meconium aspiration syndrome, sepsis/pneumonia, and congenital lung malformations.

There is good quality clinical evidence that links early surfactant treatment and reduced respiratory morbidity and improved survival without bronchopulmonary dysplasia [2, 3].

Natural surfactants are more beneficial when compared to synthetic surfactants in the treatment of respiratory distress syndrome. Babies (preterm and term) with respiratory distress where surfactant is indicated should receive a natural surfactant preparation [3].

Preparations available in the UK are:

**Poractant alfa** (Curosurf®), 120 mg / 1.5 mL & 240 mg / 3 mL, Chiesi Pharmaceuticals), a natural surfactant preparation of porcine origin

**Beractant** (Survanta® 200 mg / 8 mL, ABBVIE Ltd), a natural surfactant preparation of bovine origin

## Definitions

### Prophylactic surfactant therapy:

This preventative strategy is no longer recommended. Early rescue treatment is recommended as standard. [1,2,4] In UK, neonatal critical service specifications stipulate LNU/SCUs to transfer babies requiring complex care e.g. gestations below 27 weeks to NICUs and traditionally transport teams have taken a cautious approach and advised intubation and ventilation of these neonates.

Though this might be a pragmatic solution, prophylactic surfactant does not improve outcomes and therefore needs to be carefully considered and individualised on a case by case basis.

### **Early rescue surfactant therapy:**

Early rescue involves surfactant administration very early in the course of respiratory distress syndrome when additional oxygen requirement is relatively low, usually within 1-2 hours of birth. This approach has been reported to be superior to prophylactic treatment in terms of reduction in mortality, air leak, chronic lung disease and/or death.

Early rescue treatment is considered as current standard [2,3]. Babies with RDS should be given surfactant early in the course of the disease. **A suggested protocol would treat at a FiO<sub>2</sub> threshold of 30 % whilst on non-invasive ventilation or when there is increased work of breathing noted.** Infants who need intubation as part of their stabilisation process or in a very preterm infant who is not steroid mature should also receive early surfactant [2].

LISA (Less Invasive Surfactant Administration) / MIST (Minimally Invasive Surfactant Treatment) and INSURE (Intubation-Surfactant-Extubation) are recommended. Emerging evidence suggests that LISA may be beneficial when compared to INSURE.(5) However, LISA technique requires specific equipment, training of healthcare professionals and consideration of modified sedation protocols. Detailed description of LISA/MIST is outside the scope of this guideline, units interested in implementing LISA/MIST should have an agreed policy at unit level.

### **LISA/MIST:**

Less invasive or Minimally Invasive surfactant administration/therapy has been developed in the recent years as an alternative to INSURE. This technique instead uses a thin catheter to deliver surfactant in a spontaneously breathing infant on nasal CPAP. A recent meta-analysis reported that LISA technique reduced the need for MV and reduction in composite outcome of BPD/death and BPD in survivors (5)

### **INSURE:**

**IN**tubate, **SUR**factant, **EX**tubate is the recommended technique with a view to reducing baro/volutrauma and airway inflammation secondary to mechanical ventilation [2]. This technique involves surfactant administration after endotracheal intubation, followed by brief ventilation, and early extubation typically within 1 hour. Consideration should be given towards continuing MV for infants that are being transferred to NICUs.

### **Indications, preparation and dosage:**

#### **Primary surfactant deficiency:**

Preterm neonate 23 – 34 weeks with moderate to severe RDS and/or high risk of developing moderate/severe RDS (no or inadequate antenatal steroids, gestational diabetes etc.) Useful

guide: Preterm babies with RDS on CPAP with  $FiO_2 > 30$  **and/or** increased work of breathing with respiratory and/or mixed acidosis and/or radiographic evidence of neonatal RDS [5]

Late preterm infant/Early term infant with moderate to severe RDS requiring  $FiO_2 > 0.4$  and/or increased work of breathing

Other indications in intubated infants (secondary surfactant deficiency):

- Meconium aspiration syndrome
- Persistent pulmonary hypertension of newborn
- Congenital pneumonia
- Pulmonary haemorrhage (note this can be a rare complication of surfactant therapy in which case it should not be repeated)

Surfactant therapy not useful/not indicated:

- Chronically ventilated preterm/term infants
- Congenital diaphragmatic hernia
- Infants ventilated for non-respiratory indications (e.g. neuromuscular problems, congenital anomalies etc.)

A second or occasionally a third dose of surfactant should be considered if there is persistent high oxygen requirement ( $FiO_2 > 40\%$ ) and continuing need for mechanical ventilation.

- Preparation & storage:

- Poractant alfa (Curosurf)

- **Curosurf 120 mg®** / 1.5 ml vial Intratracheal Suspension
- **Curosurf 240 mg®** / 3 ml vial Intratracheal Suspension
- Stored in refrigerator ( $2 - 8^{\circ} C$ , protected from light)



- Beractant (Survanta)

- 200 mg / 8 mL
- Stored in refrigerator ( $2-8 C$ , protected from light)



- Dose

○ **Curosurf**

- **200 mg/kg/dose** offers significant benefits than 100 mg/kg for treatment of surfactant deficiency RDS [5]
- **100 mg/kg/dose** for secondary surfactant deficiency or for repeat doses

○ **Survanta**

- 100 mg/kg

## Surfactant administration procedure:

(Additional info on surfactant administration available @ <https://youtu.be/noNN2ocs9ZY> )

- Clarify indication with consultant and agree on dose and strategy (INSURE, LISA/MIST or surfactant administration and ventilate)
- Prescribe on drug chart on the once only side. Round up to nearest multiple of 120 mg. E.g. Curosurf .... mg (.... mg/kg/dose) via endotracheal route. Document batch no(s).
- Prepare equipment:
- Neo puff/Tom thumb (preferable) or neonatal self-inflating bag
- Suction catheter and functioning suction apparatus,
- Slowly warm up surfactant vial(s) to room temperature (placing in an incubator or within a closed fist for few minutes)
- Turn the vial upside down a few times to ensure uniform suspension
- Check colour (creamy white), expiry date and batch number
- Pre-cut surfactant administration catheter (1-2 cm short of total endotracheal tube length including adapter)
- 5 ml syringe X 2, (special syringe as surfactant coats normal syringes)
- 18 G needle X 2
- Verbal consent from parents and document in notes, if possible [6]

## Administration in delivery suite:

- Resuscitate/stabilise newborn as per first hour care and/or NLS guideline
- Intubate airway, confirm correct airway placement by ensuring bright yellow colour change on Pedi-cap
- Ensure adequate, symmetrical chest rise, stable heart rate > 100/min and saturations > 90 %
- Fix endotracheal tube at appropriate length at lips
- Ensure that baby is in supine position
- Draw up surfactant and rapidly administer using the administration set
- Withdraw syringe and draw up 1 ml of air and flush the surfactant catheter with air to ensure that surfactant has been administered
- Continue IPPV until heart rate and O<sub>2</sub> saturations are in satisfactory range
- Consider reducing PIP to decrease volutrauma following improved lung compliance after surfactant administration

- Transfer baby as soon as possible to neonatal unit after showing to parents
- Prescribe and sign prescription chart and document in clinical notes
- Complete high cost drug form (appendix A) & leave in nursing folder for consultant signature

#### **Administration in neonatal unit:**

- Draw appropriate dose of surfactant as prescribed
- Ensure proper endotracheal tube placement
- Suction ET tube if necessary
- Optimise ventilation/oxygenation as necessary
- Disconnect from ventilator and administer surfactant
- Connect to ventilator on targeted tidal volume or volume guarantee, alternatively hand ventilate gently on required pressures and reconnect to ventilator
- Once stable, reconnect to ventilator or extubate in case of INSURE
- Sign prescription chart and document procedure in clinical notes

#### **Adverse effects**

- Desaturations/bradycardia during/immediately after administration, usually resolves with IPPV
- Endotracheal tube blockage (if this results in desaturations/decrease in tidal volumes that doesn't improve with IPPV for 2-3 minutes, suction endotracheal tube to remove obstruction)
- Pulmonary haemorrhage

#### **Post procedure care**

- Remain by patient's cot side until you are satisfied that baby's clinical condition is stable
- Check oxygen saturations, tidal volume and ventilator graphics. Lung compliance rapidly improves and ventilator pressures may need to be reduced rapidly (if not on TTV) as guided by tidal volumes to prevent baro/volu-trauma
- Check a blood gas 15 – 30 minutes' post procedure
- Record any deterioration/adverse event or post procedure interventions in clinical records
- Suctioning should ideally be avoided for 1-hour post procedure, however, if surfactant is blocking the endotracheal tube and interfering with ventilation, prompt suctioning should be performed to clear obstruction
- Any unused, unopened but pre-warmed surfactant can be returned for storage in the refrigerator as long as it is within 24 hours- once only

## Audit standards

- i. All preterm infants with RDS who need mechanical ventilation should be given a natural surfactant within 30 minutes of intubation.
- ii. All preterm infants, in whom surfactant treatment is indicated, should receive an initial dose of 200 mg/kg (rounded up to the nearest multiple of 120 mg) of surfactant.

## References

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## Exceptional Circumstances Form

Form to be completed in the **exceptional** circumstances that the Trust is not able to follow ODN approved guidelines.

Details of person completing the form:	
Title:	Organisation:
First name:	Email contact address:
Surname:	Telephone contact number:
Title of document to be excepted from:	
Rationale why Trust is unable to adhere to the document:	
Signature of speciality Clinical Lead:	Signature of Trust Nursing / Medical Director:
Date:	Date:
Hard Copy Received by ODN (date and sign):	Date acknowledgement receipt sent out:

Please email form to: [mandybaker6@nhs.net](mailto:mandybaker6@nhs.net) requesting receipt.

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