Neonatal Hypoglycaemia

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1. Overview

Purpose
The purpose of this guideline is to provide evidence-based recommendations for staff and LMCs caring for a neonate at risk of hypoglycaemia.

Scope
All Maternity clinical staff and Maternity Facility Access Holders, Registered Nurses, and Medical Practitioners working within the maternity and newborn services at WDHB

2. Background information

Hypoglycaemia is a common metabolic problem in neonates. A prolonged period of hypoglycaemia, particularly if associated with clinical signs and symptoms can have an adverse impact on brain development.

The fetus is entirely dependent on transplacental glucose. After birth, the newborn has to maintain blood glucose levels independently. Healthy term babies often feed infrequently in the first 24-48 hours after birth; they manage this without ill effect because they are able to mobilize energy stores through a process known as counter-regulation. Blood glucose concentrations normally fall in the first 1-2 hours after birth, and then begin to rise again as babies mobilise their body stores of fat and glycogen and begin to feed. The
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Physiological fall itself is thought to be necessary to release hormones such as glucagon in order to stimulate glycogenolysis and gluconeogenesis, thereby attaining normal glucose homeostasis.

In some babies, this physiological fall in blood glucose does not correct and, if undetected and untreated, may potentially cause significant hypoglycaemia leading to brain damage.

Some newborns are at risk of developing hypoglycaemia, either because of inadequate supply of glucose (e.g. decreased hepatic glycogen stores) or increased glucose utilization (cold stress, infection, hyperinsulinism). All ‘at risk’ infants need to be monitored closely, regardless of mode of feeding.

3. At risk infants

- Preterm babies (<37 weeks)
- Small babies, <2.5kg or small for gestational age (SGA) with birthweight < 10th centile*
- Large babies, >4.5kg
- Diabetes in pregnancy
- Apgar score <7 at 5 minutes
- Hypothermia
- Neonates with haemolytic disease
- Neonatal syndromes (e.g. Beckwith-Wiedemann syndrome)
- Maternal drug treatment (e.g. Propranolol, Prozac (Fluoxetine), illicit drug abuse)

*To determine Small for Gestational Age (SGA) neonates, it is recommended all babies are assessed using a Customised Growth Centile chart for weight. If customised growth centile chart not available, use WHO gender specific growth charts. [http://www.gestation.net/cc/3/172546.htm](http://www.gestation.net/cc/3/172546.htm)

4. Clinical manifestations

Newborns with hypoglycaemia may be asymptomatic or may present with nonspecific clinical manifestations:

- Jitteriness
- Hypothermia
- Irritability
- Feeding intolerance
- Vomiting
- Changes in levels of consciousness: lethargy, stupor, coma
- Apnoea, cyanotic episodes
- Hypotonia, limpness
- Tremor
- Seizures

Neonatal hypoglycaemia may be an early sign of other significant disease processes requiring further investigation and treatment.

Any baby showing clinical signs of hypoglycaemia must be reviewed by the paediatric team urgently.
5. Definition of hypoglycaemia

- Blood glucose < 2.6 mmol/L, a blood glucose ≥ 2.6 mmol is considered safe for most infants
- An episode of hypoglycaemia is defined from the first blood glucose < 2.6 mmol/L until resolution with a blood glucose ≥ 2.6 mmol/L.

6. Management of infants at risk

6.1 Screening for hypoglycaemia

The following babies should be screened for hypoglycaemia (see flow charts for more details)

- All infants that have clinical signs of hypoglycaemia
- All “at risk” infants
- In SCBU, at least daily blood glucose testing on all infants on intravenous fluids

Screen using the iSTAT analyser, if no analyser is available a laboratory sample must be sent. Follow up blood glucose measurements are always indicated to confirm that the infant can maintain normal glucose levels over several feed-fast cycles.

6.2 Care at birth

- Identify risk factors
- Take steps to avoid hypothermia as this can result in hypoglycaemia. Ensure the birth room is warm
- Dry the baby, encourage immediate skin to skin contact and cover with a warm blanket. Preferably this is with the mother but if not possible, with the support person
- All babies should remain skin to skin for at least an hour, watch for feeding cues and assist mother to breastfeed, if needed
- In at risk infants commence a neonatal observation chart and 3-4 hourly before feeds hourly monitor temperature, respirations and responsiveness for 12 hrs

6.3 Feeding

- Encourage a feed within the first hour after birth. Educate the mother about feeding cues and how to recognise and anticipate cues so baby can be offered the breast when showing signs of hunger, even when this occurs less than 3 hours since last feed
- Express and give EBM (expressed breastmilk) if baby does not initiate feeding
- Aim for frequent feeds at least 8-12 in a 24 hour period
- Assess frequency and effectiveness of feeds and document breastfeeding score on feed chart
- Where feeding at the breast is not achievable, the mother must be encouraged to hand express 2-3 hourly and offer baby EBM

6.4 Monitoring of blood glucose

- Check blood glucose at 1-2 hours of age
- If glucose between 1.2-2.5 mmol/L on first testing (1-2 hours)
  - rub 0.5 ml/kg of 40% dextrose gel into buccal mucosa
  - then feed the baby (breastmilk only)
  - recheck glucose within 30min
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- Check blood glucose subsequently 3-4 hourly, preferably before feeds
- If glucose between 2.0-2.5mmol/L on subsequent testing
  - rub 0.5 ml/kg of 40% dextrose gel into buccal mucosa
  - then feed the baby (breastmilk only)
  - recheck glucose within 30min
- If feeding well, monitor glucose for at least 12 hours
- Any recorded hypoglycaemia, monitor glucose for at least 12 hours after last low level

⚠️ Inform paediatric team immediately if
  - blood glucose <1.2mmol/l at any stage
  - blood glucose < 2.0mmol/L despite one dose of oral dextrose gel
  - blood glucose < 2.6mmol/L despite two doses of oral dextrose gel
  - if feeds are not tolerated

If BMS is required an appropriate volume (5ml/kg/ feed) is offered by spoon/cup in the first instance. The indication for the prescription of BMS must be discussed with the parents when required (e.g. not in advance) and documented.

7. Dextrose 40% Gel - Dose and Administration

- Dextrose 40% gel (40g/100ml) must be prescribed on a medication chart either by a midwife or medical practitioner in the “once only” section
- Blood glucose concentration <2.6mmol/L, but ≥ 1.2mmol/L on first administration and ≥ 2.0mmol/L on subsequent doses AND
  - ≥ 35 weeks’ gestational age AND
  - < 48 hours after birth
- Dose
  - 200mg/kg (0.5 ml/kg)
  - Maximum of 3 doses of dextrose gel to treat one episode of hypoglycaemia (defined as first blood glucose level < 2.6mmol/L until subsequent blood glucose level > 2.6mmol/L)
  - Maximum of 6 doses of dextrose gel in 48 hours
- Administration
  - Draw up dose (0.5mL/kg) of dextrose 40% gel into an oral dose syringe
  - Dry inside of buccal mucosa with gauze
  - Apply dextrose to a gloved finger, and then massage into buccal mucosa until well absorbed
- Storage
  - Store in fridge
  - Dextrose Gel 40% (Biomed Ltd) expires 30 days after opening

8. SCBU

8.1 General considerations

- If bolus feeding consider increasing frequency of feeds, continuous feeds or commencing intravenous (IV) fluids (D10%)
- If low birth weight (<2500g), intravenous 10% dextrose might be indicated, and should be initiated as soon as feasible, and always within the first 2 hours after birth
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- In order to maintain blood glucose, increase fluid volume first, then dextrose concentration
- Maximum IV fluid on Day one SHOULD NOT exceed 90ml/kg/day without first consulting SMO on-call
- Infusions of dextrose of up to 12.5% may be managed with peripheral intravenous access, higher concentrations require central venous access
- Glucagon may be useful in an emergency to mobilise glycogen stores. SGA infants may not respond as well.

**Note:** Glucagon can be used for emergency situations, particularly in situations where there is difficulty starting intravenous glucose infusion

Glucagon Hypokit is available in the SCBU pyxis


8.2 Further management for persistent or severe hypoglycaemia

- In certain at risk infants the threshold may be moved to a higher blood glucose level (e.g. 3.0 or 3.5 mmol/L) in order to provide substrate for brain metabolism
- Calculate glucose requirement > 10mg/kg/min – NW Newborn Clinical Guideline - Glucose Flow Calculator
- Diazoxide needs to be discussed with SMO on-call prior to prescribing
- For persistent or severe hypoglycaemia the following tests are done at the time of hypoglycaemia, including a blood glucose at the same time (= paired blood glucose and insulin samples). Ammonia can be done anytime, regardless of blood glucose level
- A pre-packed specimen bag with micro-containers and a laminated instruction sheet is available in each unit (see 6.7 Tests and investigations)
- Consult with the Paediatric Endocrinology team at Starship Children’s Hospital if necessary

**Note:** Diazoxide can be considered in persistent and severe hypoglycaemia (requiring more than 10mg/kg/min of glucose or lasting longer than 1 week)

Diazoxide 50mg/ml Oral mixture is available in SCBU pyxis machine. This is a Section 29 drug preparation and parents should be informed that it is not registered for oral use in NZ

Consider starting chlorothiazide at the same time


8.3 Tests and investigations

<table>
<thead>
<tr>
<th>Test</th>
<th>Microtainer</th>
<th>Amount of blood required</th>
</tr>
</thead>
<tbody>
<tr>
<td>Glucose</td>
<td>Green top</td>
<td>Fill to the top line = 600micro litres</td>
</tr>
<tr>
<td>Cortisol</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Beta hydroxybutyrate (ketones)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Insulin*</td>
<td>Red top</td>
<td>Fill to the top line = 500micro litres</td>
</tr>
<tr>
<td>Growth hormone</td>
<td>Red top</td>
<td>Fill to the top line = 500micro litres</td>
</tr>
<tr>
<td>Free fatty acids (must be sent on ice)</td>
<td>Purple/EDTA</td>
<td>Fill to the top line = 500micro litres</td>
</tr>
<tr>
<td>Ammonia (must be sent on ice)</td>
<td>Green</td>
<td>Fill to the top line = 600micro litres</td>
</tr>
</tbody>
</table>

!* Consult Paediatric Endocrinology team at Starship Children’s Hospital* to help with interpretation

* Hyperinsulinism – this is a very likely diagnosis, if ketones are negative, and the insulin level inappropriately high for a low blood glucose, e.g. BSL <2.2 mmol/L, insulin > 5 mU/L
9. Documentation

- A care plan must be documented in the clinical records; this may require input from Midwife, Lactation Consultant, Charge Midwife, SCBU Nurse, paediatric medical staff
- Ensure that the infant has a feed chart and a neonatal observation chart
- Ensure that the plan is evaluated and after any treatments are re-evaluated
- Parental consent must also be documented
10. Flowchart 1: Management for at risk infants (>35/40, <48h of age)

Risk factors:
- Preterm babies (<37 weeks)
- Small, <2.5kg or SGA (<10th centile)
- Large, >4.5kg
- Diabetes in pregnancy
- Apgar score <7 at 5 minutes
- Hypothermia
- Neonates with haemolytic disease
- Neonatal syndromes
- Maternal drug treatment

At birth:
- Identify risk factors
- Avoid hypothermia, dry the baby, S2S
- Neonatal assessment
- Observation chart and feeding chart

YES

within first 2h:
- Feed as soon as feasible
- Blood glucose at 1-2h

1.2-2.5mmol/L

- Rub 0.5ml/kg of 40% dextrose gel into buccal mucosa
- Then breastfeed the baby
- Recheck glucose within 30min

<1.2mmol/L

2.0-2.5mmol/L

- Rub 0.5ml/kg of 40% dextrose gel into buccal mucosa
- Then breastfeed the baby
- Recheck glucose within 30min

<2.0mmol/L

≥2.6mmol/L

- Check blood glucose at 4h, then 3-4 hourly, preferably before feeds
- If feeding well, monitor glucose for at least 12 hours
- Any recorded hypoglycaemia, monitor glucose for at least 12h after last low level

<2.6mmol/L

Urgent paediatric review
- (consider administering 0.5ml/kg 40% dextrose gel while arranging admission)

- Max 3 doses of 40% dextrose gel per each hypoglycaemic episode
- Max 6 doses of 40% dextrose gel in 48h

Consider admission to SCBU
Management of Hypoglycaemia  
BSL < 1.7 mmol/L

1. **Blood glucose < 1.7 mmol/L**
   - IV fluids Dex 10% at 60ml/kg/day
   - May continue to feed in addition
   - Repeat blood glucose after one hour

2. **Blood glucose = 1.7 to 2.5**
   - Increase IV Dex 10% to 75ml/kg/day
   - Repeat blood glucose after one hour
   - Monitor blood glucose q3-4h
   - Increase IV Dex 10% to 90 ml/kg/day
   - Repeat blood glucose after one hour
   - Increase concentration to IV Dex 12.5%
   - If glucose requirement > 10mg/kg/min, follow management of persistent hypoglycaemia guidelines, eg blood tests, considered diazoxide/glucagon treatment, consider endocrinology consult

3. **Blood glucose >/= 2.6**
   - Wean IV fluids by approx 10%
   - Increase oral fluids accordingly

4. **Blood glucose >/= 2.6 (x3)**
   - Continue Dex 10% at 75ml/kg/day
   - Monitor blood glucose q3-4h
   - Increase IV Dex 10% to 90 ml/kg/day
   - Repeat blood glucose after one hour
   - Increase concentration to IV Dex 12.5%
   - Consider diazoxide/glucagon treatment
   - Consider endocrinology consult

5. **Blood glucose >/= 2.6 (x3)**
   - Continue Dex 10% at 90ml/kg/day
   - Monitor blood glucose q3-4h

6. **Blood glucose </= 1.7 mmol/L**
   - Consider IV bolus 2ml/kg Dex 10%

7. **Blood glucose = 1.7 to 2.5**
   - See Transition Flow Chart for further management
Management of Hypoglycaemia
BSL = 1.7 mmol/L to 2.5 mmol/L

- **Blood glucose**: 1.7 to 2.5 mmol/L
  - Feeding or IV Dex 10% at 60 ml/kg/day
  - Repeat blood glucose 1h post feed

- **Blood glucose**: 1.7 to 2.5
  - Increase fluids to 75 ml/kg/day
  - Repeat Blood glucose after 1h

- **Blood glucose**: 1.7 to 2.5
  - Increase fluids to 90 ml/kg/day
  - If not already, consider starting IV Dex 10%
  - Repeat blood glucose after one hour

- **Blood glucose**: < 1.7 mmol/L
  - Increase concentration to IV Dex 12.5%
  - If glucose requirement > 10 mg/kg/min, follow management of persistent hypoglycaemia guidelines, e.g., blood tests
  - Consider diazoxide/glucagon treatment
  - Consider endocrinology consult
  - See Flowchart (2)

- **Blood glucose**: > 2.6
  - Wean IV fluids by approx 10%
  - Increase oral fluids accordingly
  - See Transition Flowchart (4)

- **Blood glucose**: > 2.6 (x3)
  - Monitor blood glucose q3-4h, or prefeed

- **Blood glucose**: > 2.6
  - If blood glucose stabilizing
  - See Flowchart (2)
12. Flowchart 4: Transition from intravenous fluids to enteral feeding

Transition from intravenous fluids to enteral feeds

IV fluids
blood glucose > 2.6 (x3)

Give part of total fluid requirement as enteral feed, q2-3h

Blood glucose < 2.6
Increase IV fluids to previous rate, (Continue enteral feeds) Repeat blood glucose prior to next feed

Blood glucose > 2.6
If enteral feed tolerated, after 30 min decrease IV fluid accordingly, repeat blood glucose prior to next feed

Increase enteral feeds

Once on full enteral feeds, needs blood glucose > 2.6 (x3)
# Neonatal Hypoglycaemia

## 13. References

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<tbody>
<tr>
<td>WDHB Guideline</td>
<td>Birthweight Centile Calculator - Customised</td>
</tr>
</tbody>
</table>
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