Exchange Transfusion: Neonatal

Introduction
An exchange transfusion involves removing aliquots of patient blood and replacing with donor blood in order to remove abnormal blood components and circulating toxins whilst maintaining adequate circulating blood volume. It is primarily performed to remove antibodies and excess bilirubin in isoimmune disease, the incidence of exchange transfusion is decreasing secondary to the prevention, and improved prenatal management of alloimmune haemolytic disease and improvements in the management of neonatal hyperbilirubinaemia.

Indications
1. Alloimmune haemolytic disease of the newborn
   - Remove circulating bilirubin to reduce levels and prevent kernicterus
   - Replace antibody-coated red cells with antigen-negative red cells

Severe hyperbilirubinaemia secondary to alloimmune haemolytic disease of the newborn is the most common reason for exchange transfusion in the neonatal intensive care unit.

A total serum bilirubin level at or above the exchange transfusion level should be considered a medical emergency and intensive phototherapy (multiple light) should be commenced immediately. The Consultant Neonatologist on service should be contacted without delay.

2. Significant unconjugated hyperbilirubinaemia with risk of kernicterus due to any cause when intensive phototherapy is unsuccessful
3. Severe anaemia (where there is normal or increased circulating blood volume)
4. Antibodies in maternal autoimmune disease
5. Polycythaemia (to reduce haematocrit, usually accomplished with partial exchange transfusion using normal saline replacement)
6. Severe disturbances of body chemistry
The following guidelines for exchange transfusion levels are based on the American Academy of Pediatric Guidelines and are adapted from the Department of Human Services (Victoria) Neonatal Handbook.

GUIDELINES FOR EXCHANGE TRANSFUSION IN INFANTS 35 OR MORE WEEKS OF GESTATION

<table>
<thead>
<tr>
<th>Age (hrs)</th>
<th>Infants at higher risk</th>
<th>Infants at medium risk</th>
<th>Infants at lower risk</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>35-37⁺⁶ weeks + risk factors</td>
<td>≥38 weeks + risk factors or 35-37⁺⁶ weeks and well</td>
<td>38 weeks and well</td>
</tr>
<tr>
<td></td>
<td>SBR (micromol/L)</td>
<td>SBR (micromol/L)</td>
<td>SBR (micromol/L)</td>
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<tr>
<td>Birth</td>
<td>200</td>
<td>235</td>
<td>270</td>
</tr>
<tr>
<td>12 hours</td>
<td>230</td>
<td>255</td>
<td>295</td>
</tr>
<tr>
<td>24 hours</td>
<td>255</td>
<td>280</td>
<td>320</td>
</tr>
<tr>
<td>48 hours</td>
<td>290</td>
<td>320</td>
<td>375</td>
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<tr>
<td>72 hours</td>
<td>315</td>
<td>360</td>
<td>405</td>
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<tr>
<td>96 hours</td>
<td>320</td>
<td>380</td>
<td>425</td>
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<tr>
<td>5 days</td>
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<tr>
<td>6 days</td>
<td>320</td>
<td>380</td>
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</tr>
<tr>
<td>7 days</td>
<td>320</td>
<td>380</td>
<td>425</td>
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</tbody>
</table>

GUIDELINES FOR EXCHANGE TRANSFUSION IN LOW BIRTHWEIGHT INFANTS BASED ON AGE

<table>
<thead>
<tr>
<th>Age</th>
<th>Wt &lt;1500g</th>
<th>Wt 1550-2000g</th>
<th>Wt &gt;2000g</th>
</tr>
</thead>
<tbody>
<tr>
<td>Hours</td>
<td>SBR (micromol/L)</td>
<td>SBR (micromol/L)</td>
<td>SBR (micromol/L)</td>
</tr>
<tr>
<td>&lt;24</td>
<td>&gt;170-255</td>
<td>&gt;255</td>
<td>&gt;270-310</td>
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<tr>
<td>24-48</td>
<td>&gt;170-255</td>
<td>&gt;255</td>
<td>&gt;270-310</td>
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<tr>
<td>49-72</td>
<td>&gt;170-255</td>
<td>&gt;270</td>
<td>&gt;290-320</td>
</tr>
<tr>
<td>&gt;72</td>
<td>&gt;255</td>
<td>&gt;290</td>
<td>&gt;310-340</td>
</tr>
</tbody>
</table>

Notes
- Levels in the first 24 hours are less certain due to a wide range of clinical circumstances and a range of responses to phototherapy
- Immediate exchange transfusion is recommended in infants showing signs of acute bilirubin encephalopathy or if total serum bilirubin is ≥ 85 micromol/L above these levels
- Risk factors include alloimmune haemolytic disease, G6PD deficiency, asphyxia, significant lethargy, temperature instability, sepsis and acidosis
- Use total serum bilirubin (do not subtract direct acting or conjugated bilirubin)
- If total serum bilirubin does not decrease, or continues to rise despite intensive phototherapy treatment this suggests the presence of haemolysis

The final decision to perform an exchange transfusion will be made by the Consultant Neonatologist on service and will be based on the above guidelines as well as the following:
- Trend of serum bilirubin levels and response to treatment
- Clinical presentation of infant (signs of bilirubin encephalopathy)
- Underlying condition
- Previous treatment at referring hospital if applicable (including in-utero management of underlying condition)
Blood Volumes
The volume of blood for exchange is calculated using an estimate of the neonate’s circulating blood volume:
- Term infants 80ml/kg
- Preterm infants 100ml/kg

Double volume exchange transfusion
- most commonly used for removal of bilirubin and antibodies
- 2 x circulating blood volume (for example, for a term infant 2 x 80ml/kg = 160ml/kg)
- Replaces approximately 85% of the blood volume
- This will cause an approximate reduction of 50% of the pre-exchange bilirubin level (but can be expected to rebound 4 hours post transfusion to approximately two thirds of pre-exchange level)

Single volume exchange transfusion
- 1 x circulating blood volume (for example, for a term infant 80ml/kg)
- Replaces approximately 60% of the blood volume
- Consider when aetiology is not Haemolytic Disease of the Newborn

Partial exchange transfusion for polycythaemia using normal saline
- Where desired haematocrit following exchange transfusion is 0.55, the volume of exchange (mls) can be calculated as follows:

\[
\text{Volume of exchange (mls)} = \frac{(\text{actual Hct} - \text{desired Hct}) \times \text{infant's blood volume (mls)}}{\text{actual Hct}}
\]

Blood Product
- Ensure appropriate samples for pre-transfusion testing are sent to the RCH Blood Bank as early as possible.
- Notify Blood Bank via telephone as soon possible after decision is made to exchange and
- Order appropriate volume of blood for exchange
- Order FFP for transfusion midway through and at completion of exchange (10ml/kg per transfusion)
- Appropriate red cells for exchange will be provided by RCH Blood Bank. Blood for exchange transfusion should meet the following criteria:
  - Have a known haematocrit of 0.5-0.6
  - Appropriate group based on infant and maternal blood group and antibodies
  - Negative for antigens determined by maternal antibodies
  - Leukocyte depleted
  - Irradiated and used within 24 hours of irradiation
  - CMV negative
  - As fresh as possible (ensure at least less than 5 days old)
Complications
The most commonly reported adverse events during or soon after exchange transfusion:
- Catheter related complications; air emboli; thrombosis; haemorrhage
- Haemodynamic (related to excess removal of injection of blood): hypo or hypertension, intraventricular haemorrhage (preterm)
- Hypo or hyperglycaemia
- Hypocalcaemia, hyperkalaemia, acidaemia

Potential complications related to exchange transfusion:
- Arrhythmias
- Bradycardia
- Neutropenia, dilutional coagulopathy
- Feed intolerance, necrotizing enterocolitis
- Septicaemia, blood born infection
- Hypo or hyperthermia

Preparation of the Infant
- Medical staff should discuss the procedure with the parents/guardian and obtain consent
- Advise AUM and Consultant Neonatologist on duty as soon as decision to exchange is made
- Exclusively allocate at least one doctor and one nurse to care of the infant throughout the procedure
- When an exchange transfusion is taking place the Consultant Neonatologist on duty should be present on the unit to provide support and to troubleshoot issues so that the Fellow or Registrar can carry out the procedure without interruption
- Ensure resuscitation equipment and medications are easily accessible
- Nurse infant under radiant warmer for accessibility
- Ensure infant is comfortable and settled – sedation and pain relief are not usually required unless the infant is active and likely to compromise line stability or sterile field
- Ensure full cardio-respiratory monitoring is initiated and document full set of baseline observations (temperature, respiratory and heart rate, blood pressure and oxygenation)
- Infant should be nil orally as soon as decision is made to perform exchange transfusion. Pass oro/nasogastric tube and aspirate stomach contents. Leave tube in-situ and on free drainage for duration of procedure
- Before commencing exchange transfusion collect blood samples for required baseline bloods and any specific testing required. Tests may include (but not be limited to) blood cultures, blood gas, serum bilirubin, blood glucose, FBC, UEC, LFT, newborn screening test, haematological, chromosome or metabolic studies
- Establish vascular access for procedure if not already in-situ (see RCH Clinical Practice Guideline “Central Vascular Access Devices Insertion and Management”) depending on whether the procedure will be performed via arterial and venous access or via single venous access
- Check blood as per RCH Procedure “Blood Transfusion”
Equipment
- Plastic aprons or protective gowns
- Protective eye wear
- Sterile gloves
- Blood warmer – Beigler BW585
- Beigler blood warming extension set
- Blood administration set
- Water feed set
- Urine drainage bag
- Exchange transfusion recording sheet
- Sterile drape
- 3-way taps
- Syringes assorted sizes as required
- Blood gas syringes
- Drawing up needles
- Sleek tape
- Sodium chloride 0.9% and Water for Injection ampoules
- Emergency resuscitation equipment including medications and fluids
  - Calcium gluconate 10%
  - Sodium bicarbonate 8.4%
  - Glucose 10%
  - Frusemide (20mg/2ml)
- Pathology collection tubes as required
- Alcohol swabs
- Sterile gauze and
- Packed red blood cells
- Fresh frozen plasma (ordered but do not collect from Blood Bank until required)

Set-Up
Exchange transfusion should be performed slowly over approximately 2 hours to avoid major fluctuations in blood pressure.

Anticipate the need for increased oxygen requirement during procedure (administer oxygen via nasal cannula in self ventilating babies if required).

Set blood warmer at 41°C.

Blood warming extension set should be threaded onto the blood warming coil while it is not primed. Start at the back of the device and wind anti-clockwise towards the front 8 times (that is 80cm between blood warmer and patient). Line must be completely inserted between the grooves of the blood warming coil.

Check red cells for use as per RCH Procedure “Blood Transfusion”.
Using aseptic technique:

- Connect the blood administration set to the blood warming coil and clamp off the lines
- Insert the administration set spike into the bag of red cells
- Release the clamp and prime the line/coil with blood. Clamp off the lines and maintain the sterility of the end of the line
- Attach the water feed set to the urine drainage bag. Secure with a strip of sleek. Do not remove pin from self-closing clamp on the water feed set. Fasten the urine drainage bag below the cot

Strict aseptic technique should be maintained throughout procedure.

Record baseline observations (infant temperature, heart rate, respiratory rate, blood pressure, oxygen requirement, oxygen saturations, neurological status) prior to commencement of procedure.

**Technique**

Two medical officers will perform an exchange transfusion using the arterial/venous 2 line technique (one of whom is experienced in the procedure). A registered nurse may assist an experienced medical officer if a second medical officer is unavailable.

One medical officer may perform an exchange transfusion using venous 1 line technique. A second medical officer or experienced registered nurse should be available to assist as required.

**Exchange transfusion via arterial and venous lines (2 persons)**

**Arterial line**

- Without contaminating the ends of the lines, stop the infusion and turn 3-way tap so that all ports are occluded
- Disconnect intra-arterial line at 3-way tap and reconnect to uppermost port of the same 3-way tap. This will allow the line to monitor intra-arterial blood pressure during the exchange transfusion
- Do not restart the infusion
- Connect a second 3-way tap (primed with 0.9% sodium chloride) to the remaining port of the first 3-way tap.
- Connect a 10ml or 20ml syringe to the upper port of the second 3-way tap
- Attach the water feed set to the remaining port of the 3-way tap. Do not remove pin from self-closing clamp on the water feed set

**Venous line**

- Without contaminating the ends of the lines, stop the infusion and disconnect line (including PALL filter if in use)
- Attached a second 3-way tap (primed with 0.9% sodium chloride) to the venous line
- Leave 3-way tap nearest to the baby free for medication administration
- Attach a 10ml or 20ml syringe to the uppermost port of the 3-way tap furthest from the baby
- Connect the end of the primed blood warming coil line to the 3-way tap furthest from the baby
Exchange transfusion via venous lines only (1 person)

- Without contaminating the ends of the lines, stop the infusion and disconnect line (including PALL filter if in use)
- Turn the 3-way tap off to the infant
- Attach two 3-way taps (primed with 0.9% sodium chloride) to the 3-way tap already attached to the umbilical venous line
- Leave 3-way tap nearest to the baby free for medication administration
- Connect the end of the primed blood warming coil line to the middle 3-way tap
- Connect a 10ml or 20ml syringe to the uppermost port of the 3-way tap furthest from the baby
- Attach the water feed set the last remaining port of the 3-way tap. **Do not remove pin** from self-closing clamp on the water feed set

Procedure

Exchange transfusion involves the sequential withdrawal and injection of aliquots of blood, through arterial and venous lines, either peripheral or central. Note arterial lines (umbilical or peripheral) should only be used for withdrawal of infant blood, not for injection of donor blood.

<table>
<thead>
<tr>
<th>In</th>
<th>Out</th>
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<tbody>
<tr>
<td>Umbilical</td>
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<tr>
<td>vein</td>
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<tr>
<td>Umbilical</td>
<td>Peripheral</td>
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<tr>
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<td>artery</td>
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<td>Peripheral</td>
<td>Peripheral</td>
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<tr>
<td>vein</td>
<td>artery</td>
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</tbody>
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Aliquots usually tolerated for exchange transfusion:
Less than 1500g – 5ml
1500g – 2500g – 10ml
2500g – 3500g – 15ml
Greater than 3500g – 20ml

Small aliquots exchanged at a slower rate are usually better tolerated by infants with cardiovascular instability. As a guide the time for each in/out should be around 5-10 minutes (withdrawal and infusion of blood should be performed at the same rate). In and out aliquots are repeated sequentially until desired volume for exchanged is reached. In and out aliquots should be called out, so that the nurse recording the procedure can keep an accurate tally,

Always check 3-way tap settings to ensure blood withdrawal and infusion flows in the correct direction to prevent accidental loss or infusion of blood.

Registered nurse should observe the infant throughout the exchange transfusion and record the following on the Exchange Transfusion Record Sheet:
- volume of blood withdrawn and injected at the end of each cycle (aliquot)
- infant temperature, heart rate, respiratory rate, blood pressure, oxygen requirement, oxygen saturations, blood warmer temperature, general condition of infant – **every 15 minutes**
- If phototherapy lights remain on during procedure they should be turned off frequently to assess infant colour and general condition. Observe infant for clinical signs of complications of exchange transfusion (i.e. agitation due to hypocalcaemia, signs of hypoglycaemia)
- Administration of medications as required
- Routine observations should continue to be recorded on the ICU/NNU observation chart (MR100) each hour as per standard procedure.

Blood glucose and blood gas should be monitored at least pre, mid and post exchange and more frequently as indicated to manage metabolic and electrolyte derangements.

After each 100ml of blood exchanged, flush line with 0.9% sodium chloride, administer 1ml of 10% calcium gluconate (diluted with 1ml of Water for Injection) by slow push followed by a 0.9% sodium chloride flush. The calcium gluconate should not come into direct contact with donor red cells or clotting may occur. Monitor for changes in heart rate and rhythm during calcium gluconate administration.

The last withdrawal volume should be saved for post exchange blood tests.

**Cease exchange transfusion if infant's condition suddenly deteriorates.** If the exchange transfusion is ceased for any reason, always leave the infant’s blood volume in balance. Sudden deterioration in infant’s condition may be related to the procedure, underlying condition or an adverse reaction to transfusion. A transfusion adverse reaction report should be made

**Fresh Frozen Plasma Administration**
Midway through the exchange transfusion and at the completion of transfusion, fresh frozen plasma should be administered at 10ml/kg in place of a blood aliquot. Aliquot of FFP should be given over 15-20 mins and patient temperature should be monitored frequently throughout. If patient has a peripheral IVC in place, or second UVC lumen, consider using for FFP transfusion and transfuse according to RCH Procedure “Blood Transfusion”.

Blood bank will require 20 minutes notice for thawing of fresh frozen plasma.

**Partial Exchange Transfusion**
- Maintain aseptic technique during procedure and follow Standard Precautions
- Administer required amount of 0.9% sodium chloride via venous line
- Remove required amount of blood via venous or arterial line
- Ensure administration and removal of fluid occurs at compatible rate

**Post Procedure Care of the Infant**
- Continuously monitor vital signs and record 30 minutely for first 4 hours post procedure. If stable after this time routine observations as per ICU/NNU observation chart (MR100) should be continued
- Perform blood glucose levels immediately post procedure and then hourly until stable
- Measure serum bilirubin levels one hour post exchange transfusion and repeat 6 hourly
• Continue phototherapy until bilirubin levels are within acceptable range. Anticipate rebound increase in serum bilirubin (up to 60% of pre-exchange level) 2-4 hours post procedure
• Observe catheter sites for signs of bleeding
• Keep infant NBM for at least 4 hours post exchange transfusion, or longer at the direction of the medical officer. As exchange transfusion carries a potential risk of necrotizing enterocolitis (especially in the preterm infant) monitor the appearance of the abdomen and the presence of bowel sounds. Observe for signs of feed intolerance when feeding is recommenced
• Measure urea and electrolytes, full blood examination, haematocrit and blood gas on a regular basis until infant stable (as directed by medical officer)
• The medical officer performing the exchange transfusion should document the procedure in the progress notes

See also:
RCH Clinical Practice Guideline: Jaundice in Early Infancy
RCH Procedure RCH0395: Blood Transfusion
RCH Clinical Practice Guideline: Central Vascular Access Devices Insertion and Management
Neonatal Handbook (NETS): Jaundice in the First Two Weeks of Life

Reference Documents


The Royal Women’s Hospital, Melbourne, Exchange Transfusion (total & partial): Neonatal, 2006.