

Trust Guidelines for the use of Blood Products in Newborn Infants

A clinical guideline recommended for use:

| | |
|---|---|
| In: | NICU |
| By: | Medical and Nursing staff |
| For: | Newborns in NICU and SCBU requiring blood product support |
| Division responsible for document: | Women and Children's Services |
| Key words: | Neonate, Blood, CGAP |
| Name of document author: | Dr M P Dyke |
| Job title of document author: | Consultant Neonatologist |
| Name of document author's Line Manager: | Dr H O'Reilly |
| Job title of author's Line Manager: | Consultant Neonatologist |
| Supported by: | Dr R Roy, Dr P Clarke, Dr P Muthukumar, Dr F Walston, Consultant Neonatologists; Dr J Wimperis, Dr H Lyall, Consultant Haematologists |
| Assessed and Approved by the: | Chair's Action Clinical Guidelines Assessment Panel (CGAP) If approved by committee or Governance Lead Chair's Action; tick here <input checked="" type="checkbox"/> |
| Date of approval: | 13/09/2021 |
| Ratified by or reported as approved to (if applicable): | Clinical Safety and Effectiveness Sub-Board |
| To be reviewed before: This document remains current after this date but will be under review | 13/09/2024 |
| To be reviewed by: | Dr M P Dyke |
| Reference and / or Trust Docs ID No: | 1202 |
| Version No: | 6 |
| Compliance links: | None |
| If Yes - does the strategy/policy deviate from the recommendations of NICE? If so why? | N/a |

This guideline has been approved by the Trust's Clinical Guidelines Assessment Panel as an aid to the diagnosis and management of relevant patients and clinical circumstances. Not every patient or situation fits neatly into a standard guideline scenario and the guideline must be interpreted and applied in practice in the light of prevailing clinical circumstances, the diagnostic and treatment options available and the professional judgement, knowledge and expertise of relevant clinicians. It is advised that the rationale for any departure from relevant guidance should be documented in the patient's case notes.

The Trust's guidelines are made publicly available as part of the collective endeavour to continuously improve the quality of healthcare through sharing medical experience and knowledge. The Trust accepts no responsibility for any misunderstanding or misapplication of this document.

Trust Guidelines for the Use of Blood Products in Newborn Infants

Version and Document Control:

| Version No. | Date of Update | Change Description | Author |
|-------------|----------------|--|-------------|
| 6 | 13/09/2021 | Slightly amended thresholds for using platelet transfusions in newborns as a result of recently published trial. | Dr M P Dyke |
| | | | |
| | | | |
| | | | |

This is a Controlled Document

Printed copies of this document may not be up to date. Please check the hospital intranet for the latest version and destroy all previous versions.

Trust Guidelines for the use of Blood Products in Newborn Infants

Contents

| Topic | Page |
|---|-------------|
| 1. Objective | 4 |
| 2. Rationale | 4 |
| 3. Broad Recommendations | 4 |
| 4 Blood Products | |
| 4.1 Red Cell Transfusion | 5 |
| 4.1.1 Indications | |
| 4.1.2 Procedure for cross-matching | |
| 4.1.3 Management of specific situations | |
| 4.1.4 Other practical points | |
| 4.2 Platelets | 8 |
| 4.2.1 Normal range | |
| 4.2.2 Incidence | |
| 4.2.3 Causes of thrombocytopenia | |
| 4.2.4 Risk | |
| 4.2.5 Platelet transfusions | |
| 4.2.6 Management of specific situations | |
| 4.2.6.1 Neonatal Allo-immune thrombocytopenic purpura | |
| 4.2.6.2 Thrombocytopenia from other causes | |
| 4.3 Fresh Frozen Plasma | 12 |
| 4.3.1 Clotting studies | |
| 4.3.2 Specific conditions | |
| 4.3.2.5 Surgery | |
| 4.4 Cryoprecipitate | 14 |
| 4.5 Human Albumin Solution | 14 |
| 4.6 Intravenous Immunoglobulin | 14 |
| 5. Clinical audit standards | 15 |
| 6. Glossary | 15 |
| 7. Summary of development and consultation process | 15 |
| 8. Distribution list/ dissemination method | 16 |
| 9. References/ source documents | 16 |
| Appendix 1 Source and preparation of Neonatal Blood products | 20 |
| Appendix 2 NICU Blood Transfusion form | 21 |
| Appendix 3 NICU FFP Form | 22 |

Trust Guidelines for the Use of Blood Products in Newborn Infants

1. Objective/s

To facilitate the safe use of blood products in the newborn infant admitted to NICU-SCBU

2. Rationale

Newborn infants may suffer from a variety of conditions for which blood products are indicated. In the past blood products have been very widely used in neonatal intensive care, raising concerns about donor exposure and the risks of transmissible agents.

There is published evidence confirming that the use of guidelines restricting transfusion can safely reduce the numbers of transfusions, volumes of blood and donor exposure for each infant.

Short term benefits (such as a reduction, following transfusion, in cardiac output which had become elevated due to anaemia) is only seen when “restrictive” transfusion policies are followed i.e. Hb values allowed to fall significantly before transfusing.

Long term risk-benefit ratios are not well documented but “restrictive” policies have no adverse effects on death or hospital discharge and may be protective of long term neurodevelopment. This guideline draws on US and UK consensus statements on the rational use of blood products and some randomised controlled trial data, to offer a consistent approach by which we might reduce the risks of under- or over-treatment.

3. Broad recommendations

Many infants admitted to NICU develop conditions for which the use of blood products might be considered. The following notes should guide management:

- Infants should **NOT** receive a blood product unless it is absolutely necessary. If in doubt consult a senior colleague.
- Whenever practicable, the planned use of blood products should be discussed with parents in advance.
- All efforts should be made to minimise the necessity for blood products:
 - Delayed cord clamping should be encouraged.
 - Crystalloid infusions may be used instead of colloid or blood in many circumstances.
 - Blood sampling should be kept to the minimum required for safe practice.
 - Blood drawn to “clear” an arterial line should be returned to the patient after sampling.

4. Blood products

4.1 Red Cell Transfusion

4.1.1 Indications for red cell (rbc) transfusion

[this should be recorded in the notes]

Red cell transfusion may be required for the following conditions:

- Blood loss with hypovolaemic shock.
 - This situation requires **urgent fluid volume resuscitation**, usually with crystalloid infusion initially, followed by blood (either type-specific cross-matched or Gp O RhD Neg).
 - **Massive Blood Loss [MBL]** is uncommon in neonatal practice but requires urgent attention to additional measures over and above rbc transfusion:
 - MBL should be considered when blood loss is estimated to be:
 - $\geq 3\text{ml/kg/min}$.
 - $\geq 40\text{ml/kg}$ over 3 hours.
 - $\geq 80\text{ml/kg}$ over 24 hours.
 - Details of the Children's MBL protocol [C-MBL] and how to put it into action are contained in the Trust Guideline for the Management: of Massive Blood Loss in Children (C-MBL) [Trustdocs ID: 9960](#).
- Top-up transfusion for infant requiring respiratory support.
 - The Hb should be maintained Hb $>120\text{g/L}$ (PCV >40) for infants on a ventilator or on nCPAP or Vapotherm with significant oxygen requirement ($\text{FiO}_2 >0.4$).
 - The Hb should be maintained Hb $>100\text{g/L}$ (PCV >35).
 - For infants on ventilator or nCPAP/vapotherm with $\text{FiO}_2 < 0.4$.
 - For infants off respiratory support but with high O_2 requirements (e.g. $>0.75\text{L/min}$ via nasal cannula).
- In infants with other symptoms thought to be due to anaemia (e.g. frequent apnoea requiring resuscitation, poor feeding, poor weight gain, signs of congestive cardiac failure) consider transfusion if Hb $<70\text{g/L}$.
- In an asymptomatic infant with an Hb $<70\text{g/L}$, check reticulocyte count and discuss with consultant (NB our unit policy is NOT to test FBC routinely in asymptomatic infants).

Exchange transfusion for Haemolytic Disease see
Performing Exchange Transfusion in Newborn [Trustdocs id 1304](#)

4.1.2 Procedure for cross-matching

- On admission, take a 1mL EDTA specimen for Group and Save from all infants requiring intensive care.
- Ensure, where possible, that a 7mL EDTA specimen of maternal blood has been sent for Group, antibody screening and cross-match.
- Register all the following details on the ICE request:
 - Patients full name.

Trust Guidelines for the Use of Blood Products in Newborn Infants

- Hospital number.
- Date of birth.
- Any special circumstances (e.g. intra-uterine transfusions, suspicion of NAITP or immuno-deficiency).

4.1.3 Management of Specific situations

4.1.3.1 Top-up transfusion

- Inform Blood bank (NNUHFT) of infant's weight. Blood will be allocated from a **satellite-pack** ie a unit from single donor divided into multiple small volume units which are reserved for one infant to minimise donor exposure.
 - Infants >1.5kg will have a 3-unit pack.
 - Infants <1.5kg will have a 6-unit pack.
- Request the required **volume** of packed red cells from NNUH blood bank (including volume for priming the lines). See box below for details.
- Ensure that the blood supplied is:
 - ABO compatible with mother and infant.
 - RhD compatible with infant.
 - Irradiated if the infant has received intra-uterine transfusion or has congenital defect of cellular immunity eg Di George syndrome (Trust Guideline for the Use of Irradiated Blood and Blood Products, [Trustdocs ID: 1286](#)).
- Except when blood transfusion is required as an emergency, the planned transfusion should be discussed in advance with the parents and consent obtained
- Give packed red cell transfusion using the formula:

1. For infants on a ventilator/ nCPAP/Vapotherm with $FiO_2 >0.4$, aim for an Hb ~160g/L

**Volume of transfusion (mLs) = 0.4 x desired rise in Hb (g/L) x weight (kg)
(usually in range 20-25ml/kg)**

e.g. 0.75kg infant with Hb of 110g/L **Volume = 0.4 x (160-110=50) x 0.75 = 15mLs**

NB Remember to request an additional 4mLs to "prime" the line

2. For infants on ventilator/nCPAP/vapotherm with $FiO_2 \leq 0.4$ OR for infants off respiratory support but with high O_2 requirements (eg $>0.75L/min$ via nasal cannula) aim for an Hb ~120-140 g/dL

Volume of transfusion (mLs) = 0.4 x desired rise in Hb (g/L) x weight (kg)

eg 1.5kg infant with Hb of 80g/L **Volume = 0.4 x (120-80=40) x 1.5 = 24mLs**

NB Remember to request an additional 4mLs to "prime" the line

3. For other infants with symptoms [e.g. apnoea] etc aim for an Hb ~100-120g/dL

Volume of transfusion (mLs) = 0.4 x desired rise in Hb (g/L) x weight (kg)

eg 2kg infant with Hb of 65 g/L **Volume** = 0.4 x (110-65 = 45) x 2 = 36mLs

NB Remember to request an additional 4mLs to “prime” the line

- Infuse over a maximum of 4 hours (shorter duration if treating hypovolaemia).
- **Furosemide** confers no clinical benefit in most infants and is not routinely required but may be given (at Consultant discretion) for an infant who:
 - Is in heart failure.
 - Has severe chronic lung disease (e.g. requiring ventilation).
 - Is ventilated and receiving active treatment for a PDA.[When required, Furosemide dose = 1mg/kg].

4.1.4 Other practical points on rbc transfusion

- The circulating blood volume of an infant is approximately 80-90 mL/kg.
- All cellular blood products are capable of causing graft-versus-host disease. This risk is reduced by leucodepletion and may be reduced further by irradiation of blood products where required.
- Irradiation shortens shelf-life to 14 days in satellite packs used for top up transfusions and to 24 hours in blood used for exchange transfusion. Only request irradiated blood for infants who:
 - Have suspected congenital defects of cellular immunity (e.g. Di George Syndrome).
 - Have received in-utero transfusions.
 - Are receiving directed donation from 1st degree relatives.
 - Are receiving an Exchange Transfusion.
- The supply of irradiated products may introduce a delay before transfusion can commence. In situations where this delay may pose a greater risk to the infant than the use of non-irradiated products, a decision to proceed with non-irradiated products may be in the best interests of the infant. This decision should be made at Consultant level after discussion with the NNUH Consultant Haematologist on-call.
- The preferred route of administration is a peripheral venous cannula or umbilical venous catheter. Transfusion via umbilical arterial catheter carries higher risk and should be used only at Consultant discretion.
- Observations of temperature, heart rate, respiratory rate, BP and Oxygen saturations should be carried out at the start of the transfusion then every 15 minutes for the first hour then hourly throughout the remainder of the transfusion.

Trust Guidelines for the Use of Blood Products in Newborn Infants

4.2 Platelets

4.2.1 Normal range = 150-450x10⁹/L

4.2.2 Incidence of thrombocytopenia is 1-4% of all newborns, 20-35% of all NICU admissions and 40-72% of sick preterm infants.

4.2.3 Causes of thrombocytopenia in newborn are:

- Prematurity.
- Placental disease (associated with pre-eclampsia, small for gestational age infant, maternal HELLP syndrome).
- Sepsis.
- Disseminated Intravascular Coagulopathy (DIC).
- Neonatal Allo-immune thrombocytopenic purpura (NAITP).
- Large Haemangioma (Kasabach-Merritt syndrome).
- Maternal Idiopathic Thrombocytopenic Purpura (ITP).
- Rhesus Haemolytic disease.
- Congenital infection.
- Necrotising entero-colitis (NEC).

4.2.4 Risk

- There is no single “safe platelet count” applicable to all infants – the risk of bleeding may depend on causation, gestation, associated coagulopathy and age at presentation.
- Whilst there is an association between severe thrombocytopenia and episodes of bleeding, a causal relationship remains unproven.
- The role of prophylactic platelet transfusion in preventing new episodes of bleeding and/or the extension of existing haemorrhage is unproven.
- Most thrombocytopenia occurs in preterm (+/- small for gestation) infants.
- Most thrombocytopenia is asymptomatic, mild, of early-onset (<72 hours) and only slowly progressive.
- Late-onset thrombocytopenia (>72 hours) is often more severe and rapidly progressive. It is also more likely to be associated with other pathology such as sepsis or NEC.
- The finding of NAITP carries a considerably higher risk of haemorrhage than other causes of thrombocytopenia and particularly when diagnosed ante-natally.
- NAITP should be considered in any infant with significant thrombocytopenia (see 4.2.6 below) particularly when other risk factors for thrombocytopenia are absent.

4.2.5 Platelet transfusions

- Are required in only the minority of thrombocytopenic infants ie those who have symptoms or severe thrombocytopenia (see sections 4.2.6.1 (NAITP). and 4.2.6.2.2 (others)).
- Should **not** be given without consultant approval.

Trust Guidelines for the Use of Blood Products in Newborn Infants

- Should be given as 20 mL/kg infusion of standard platelet suspension over 30 mins (except in NAITP (see 4.2.6.1 below)).

4.2.6 Management of Specific Conditions

4.2.6.1 Neonatal Allo-immune thrombocytopenic purpura (NAITP)

- NAITP is rare (approximately 1:1000 pregnancies) but may cause severe haemorrhage (fetal intra-cerebral haemorrhage [ICH] rates of 2-20% in an affected pregnancy) and death.
- There are at least 19 recognised platelet-specific antigens of which 14 have been characterised at molecular level. Of these, HPA-1a, HPA-5b and HPA-3a are the most important in the aetiology of NAITP.
- 98% of the population are HPA-1a antigen positive; only 2% HPA-1a antigen negative.
- An HPA-1a negative mother with HPA-1a positive fetus may form anti-HPA-1a antibodies (in a manner similar to that seen with red cells in Rhesus haemolytic disease). However, in practice, only around 1 in 10 do so.
- Anti-HPA IgG antibodies cross the placenta and facilitate platelet destruction. Only about 40% of those with antibodies have thrombocytopenia and, of those, fewer than 50% will have platelet counts $<50 \times 10^9/L$.
- Unlike Rhesus disease, it is not uncommon for first pregnancies to be affected.

4.2.6.1.1 NAITP Clinical presentation

- In-utero:
 - Symptomatic: cerebral haemorrhage, hydrops or hydrocephalus.
 - Asymptomatic: by screening following a previous pregnancy in which NAITP was either confirmed or strongly suspected (eg unexplained 2nd or 3rd trimester IUD or recurrent late miscarriage).
- In newborn:
 - Bleeding, bruising, petechiae.
 - Platelet count performed for other reasons.

4.2.6.1.2 NAITP Investigation

- It is extremely important to make the diagnosis for the management of future pregnancies as well as the immediate clinical problem.
- Discuss platelet antigen investigations with Consultant Neonatologist and/or Haematologist on-call.
- Send to NNUH Blood Bank for Group and Platelet Antigen Screen:
 - Infant – 1mL EDTA specimen (cord blood if antenatally-diagnosed case).
 - Mother – 3x6mL EDTA and 6mL Clotted.
 - Father – 3x6 mL EDTA.

4.2.6.1.3 NAITP Management

- Obstetric management is dealt with in a separate guideline (see NNUHFT Intranet Obstetric Dept Guideline page Ref:AO23)

Known NAITP

- Avoid intramuscular injections (e.g. vitamin K) unless platelet count is known to be $>50 \times 10^9/L$.
- Severe thrombocytopenia (Platelet count $<25 \times 10^9/L$).
 - Give urgent platelet transfusion with **ABO/RhD-compatible HPA1a- and HPA5b-negative platelets** if available (may take up to several hours). If severely haemorrhagic, a transfusion of random donor platelets may be available more quickly and may produce a short-lived increment of clinical benefit.
 - If platelets are unavailable, IVIg 1g/kg/dose given as an infusion over 6 hours on two consecutive days will often raise the platelet count but may take 24-48 hours to be effective. Contact pharmacy and complete the request form for Immunoglobulin available from intranet (<http://intranet/committees/DTMM/index.htm>).
- Moderate thrombocytopenia (Platelet count $25-50 \times 10^9/L$).
 - Asymptomatic – monitor count daily until rising trend is established and continue to monitor until count $>50 \times 10^9/L$. If count falls $<25 \times 10^9/L$ transfuse as above.
 - Symptomatic – transfuse as above.
- Mild thrombocytopenia (Platelet count $>50 \times 10^9/L$).
 - Monitor daily until rising trend established.
 - Transfuse as above if significant bleeding occurs.

Suspected but unproven NAITP

- Severe thrombocytopenia (Platelet count $<30 \times 10^9/L$).
 - Discuss the suspected diagnosis with the NNUH Haematology Consultant and Blood Bank as soon as possible.
 - Perform serology and antigen status on infant and parents (see 4.1.6.1.2) but do not await results before treating.
 - Give urgent platelet transfusion with **ABO/RhD-compatible HPA1a- and HPA5b-negative platelets** if available [may take several hours]. If severely haemorrhagic, a transfusion of random donor platelets may be available more quickly and may produce a short-lived increment.
 - Monitor platelet count daily initially until rising trend established and continue to monitor until count $>50 \times 10^9/L$.
- Moderate thrombocytopenia (Platelet count $25-50 \times 10^9/L$)
 - Asymptomatic – monitor daily initially until rising trend established and continue to monitor until count $>50 \times 10^9/L$. If count falls $<25 \times 10^9/L$ then transfuse as above.

Trust Guidelines for the Use of Blood Products in Newborn Infants

- Symptomatic – transfuse as above.

4.2.6.1.4 NAITP Follow-up

- Recovery of normal platelet counts usually takes 10-14 days but delays of 3-4 weeks are described.
- Monitor platelet count daily until rising trend is established and then less frequently until count $>50 \times 10^9/L$.
- Appropriate counselling should be arranged for the mother with a Consultant Obstetrician and Consultant Haematologist to cover implications for future pregnancies and transfusion requirements.

4.2.6.2 Thrombocytopenia from other causes

4.2.6.2.1 Investigations

- Consider list of causes (4.2.3) and investigate appropriately based on history, examination and time of onset.

| Classification of Thrombocytopenia by most common time of onset | |
|---|--|
| Time of onset | Condition |
| Fetal | Alloimmune Congenital infection Aneuploidy Maternal Autoimmune (ITP, SLE) Severe Rh Disease Congenital/inherited (eg Wiskott-Aldrich) |
| Early-onset <72 hours | Placental insufficiency Perinatal asphyxia DIC Alloimmune Maternal Autoimmune Congenital infection Thrombosis BM replacement eg leukaemia Kasabach-Merritt Congenital/inherited (eg TAR) |
| Late-onset ≥ 72 hours | Sepsis NEC Congenital infection Kasabach-Merritt Congenital/inherited (eg TAR) |

- Clotting studies should be performed:
 - If platelet count $<25 \times 10^9/L$.
 - If there is active bleeding.
 - With signs of severe sepsis or NEC.

4.2.6.2.2 Management

Transfuse 20 ml/kg of type-specific platelets over 30 minutes if:

- Severe thrombocytopenia (Platelet count $<25 \times 10^9/L$).
- Mild ($>50 \times 10^9/L$) or moderate thrombocytopenia ($25-50 \times 10^9/L$) **and** active bleeding.

Consider platelet transfusion if:

- Indometacin or Ibuprofen to be used (e.g. for PDA closure) and platelet count $<50 \times 10^9/L$.
- Invasive procedure required (e.g. surgery, lumbar puncture, suprapubic aspiration of urine) and platelet count $<50 \times 10^9/L$.

If auto-immune disease is suspected (maternal ITP/SLE) **IVIg 1g/kg/dose** for two consecutive days may be effective.

4.3. Fresh Frozen Plasma (for coagulopathy)

4.3.1 Notes on Clotting studies

There is some evidence for “normal range” values for term infants but those for preterm and sick new-borns are not clearly established. Commonly, such infants have some prolongation of PT and APTT; also Fibrinogen levels are lower than in older patients. However, there is no evidence to suggest that underlying derangement of clotting function is implicated in most periventricular or other forms of haemorrhage, including surgical bleeding

4.3.1.1 Indications for clotting studies

Clotting studies are not routinely indicated in well infants, including those who require surgery. They should be performed:

- Where there is clinical evidence of bleeding (pulmonary, umbilical, intraventricular haemorrhage grade 3 or above).
- In clinical situations where the risk of DIC is significant and platelet count is low (see 4.2.6.2.1).
- In acute liver failure.
- Where there is a known inheritable defect of coagulation factors (e.g. Haemophilia).
- After significant peri-partum hypoxia-ischaemia.

NB if fibrinogen levels are required, a separate request must be made on ICE

4.3.1.2 Sampling

- Fill a blue topped bottle with blood exactly to the required mark (approximately 1.8 mLs).

Trust Guidelines for the Use of Blood Products in Newborn Infants

- Ideally the blood should not be drawn from a heparinised line, but if alternative routes are impractical, then draw back approximately 5 mLs to “clear” the line, draw the sample and replace the “cleared” blood. Identify source of sample on ICE request.

4.3.1.3 Normal values

| Test | Gestation in weeks | | | Age |
|-------------|--------------------|------------|------------|-------|
| | 28-31 | 32-36 | term | adult |
| PT (secs) | 23 | 17 (12-21) | 16 (13-20) | 12-14 |
| APTT (secs) | N/A | 70 (58-82) | 55 (45-65) | 44 |
| Fibrinogen | 2.43 (1.5-3.8) | | | |

Nathan & Oski Hematology of Infancy and Childhood 4th Ed: Saunders 1992 Robertson 4th Ed

4.3.2 Specific conditions

4.3.2.1 Vitamin K Deficiency Bleeding (VKDB) is preventable with a single dose of parenteral Vitamin K.

- Vitamin K should be given to all newborn infants admitted to NICU.
- A second dose is rarely required in the first few weeks of life and should not be given unless there is good evidence of deficiency.
- Suspected Vitamin K deficiency should be treated with parenteral vitamin K and, if active bleeding occurs, with FFP 20 mL/kg.
- Where vitamin K deficiency is suspected, this should be confirmed by measuring PIVKA-II levels (elevated in VKDB).

4.3.2.2 Disseminated Intravascular Coagulopathy (DIC)

- Is the most common coagulopathy in preterms and sick newborns.
- Treat underlying cause whenever possible (e.g. sepsis, acidosis, hypoxia).
 - If coagulopathy is severe (INR > 2) give 20mLs/kg fresh frozen plasma (FFP) over 1 hour (**after discussion with consultant**).
 - If fibrinogen falls below 0.8g/L give cryoprecipitate 10ml/kg over 30 minutes.
 - Repeat clotting studies after treatment (with FFP or cryoprecipitate).

4.3.2.3 Severe liver failure is usually accompanied by profound coagulation derangements, including hypo-fibrinogenaemia. These children will need blood product support with cryoprecipitate (if the fibrinogen is less than 0.8 g/l) and FFP, until the liver recovers or the child has a liver transplant.

4.3.2.4 Other coagulation defects are rare and should be discussed with a NNUHFT Consultant Haematologist for advice on appropriate investigation and Factor replacement. FFP may occasionally be needed if active bleeding requires urgent treatment before specific diagnosis or treatment is available.

4.3.2.5 Surgery

- Well infants do not require routine pre-operative testing of clotting studies.
- Clotting studies should be performed in infants with:

Trust Guidelines for the Use of Blood Products in Newborn Infants

- Previous significant bleeding.
- Sepsis.
- Other conditions with high risk of DIC eg severe hypoxia-ischaemia.
- Whilst there is no published evidence on which to base recommendations about “safe” levels of clotting values for infants scheduled for surgery, local consensus [haematology, neonatal, surgical and anaesthetic] suggests the use of FFP should be considered with values around 1.5 x normal ie PT > 24 secs or APTT > 66 secs.

4.4 Cryoprecipitate

- May, occasionally, be required in cases of hypofibrinogenaemia (eg DIC).
- If fibrinogen falls below 0.8g/L give cryoprecipitate 10ml/kg over 30 minutes.

4.5 Human albumin solution

4.5.1 Hypovolaemia: Colloid infusions (human albumin, FFP etc) are not indicated in the resuscitation of infants requiring fluid volume expansion for suspected hypovolaemia. Instead, give 0.9% saline 10mLs/kg infused over 20mins.

4.5.2 Metabolic acidosis requires appropriate steps to identify and treat the cause. Occasionally, hypoperfusion due to hypovolaemia will be the suspected cause and a single infusion of saline (as above) may be justified. Colloids should not be used.

4.5.3 Hypoalbuminaemia:

- Low serum albumin levels and peripheral oedema are extremely common in preterm infants (mean values rise from 19g/dl at 26 weeks GA to 31g/dl at term).
- There is no controlled trial evidence of clinical benefit from albumin infusion in hypoalbuminaemic preterm infants.
- 4.5% human albumin solution (HAS) should not be used as it contains insufficient albumin to effect a rise in serum values.
- 20% HAS (10ml/kg or 2g/kg) may be considered for use when severe hypoalbuminaemia (<15g/dl) is associated with:
 - Pulmonary oedema.
 - Hypovolaemia causing circulatory compromise.
 - Gross tissue oedema compromising wound healing post-surgery.

4.6 Intravenous Immunoglobulin (IVIg)

The use of Intravenous immunoglobulin may be indicated in the following conditions:

- Immune-mediated thrombocytopenia.
- NAITP - see section 4.2.6.1.
- Maternal ITP - see section 4.2.6.2.2.
- Maternal SLE - see section 4.2.6.2.2.

Trust Guidelines for the Use of Blood Products in Newborn Infants

- Haemolytic Disease of the Newborn (see Trust Guideline for the Management of Haemolytic Disease of Newborn including Exchange Transfusion Ref: CA4053 V2).

NB. Supplies of IVIg are variable and usage may be limited at times of shortage.

5. Clinical audit standards

1. All blood products given should have a clear clinical indication recorded in the notes, compatible with advice in this guideline.
2. All infants with thrombocytopenia should have platelet counts monitored at intervals compatible with advice in this guideline.
3. Platelet transfusions should only be given for thrombocytopenia which is either severe or associated with significant bleeding.
4. Where coagulopathy requires treatment with FFP or cryoprecipitate, the clinical indication for treatment should be clearly documented.
5. Where coagulopathy requires treatment with FFP or cryoprecipitate, clotting studies should be repeated after treatment.

6. Glossary

| | |
|------------------|---|
| FiO ₂ | Fractional Inspired Concentration of Oxygen |
| FFP | Fresh Frozen Plasma |
| GA | Gestational age |
| Hb | Haemoglobin |
| HELLP | Haemolysis, Elevated Liver enzymes, Low platelets |
| HPA | Human Platelet Antigen |
| NAITP | Neonatal Alloimmune Thrombocytopenic Purpura |
| PCV | packed cell volume |
| RBC | Red blood cell |
| SLE | Systemic Lupus Erythematosus |

7. Summary of development and consultation process undertaken before registration and dissemination

The original guideline (CA 2045) was drafted by Drs Dyke, Wimperis and Turner on behalf of the Neonatal and Haematology Departments. It was circulated for comments to Neonatal Consultants Specialist Registrars, ANNPs and SHOs and discussed at the Departmental Guidelines meeting where amendments were made to incorporate suggestions and comments. The neonatal unit pharmacist also reviewed it. The main author undertook an audit of practice against this guideline in 2009, reviewed the published literature to inform guideline updates in 2009 and 2012 and has incorporated some new evidence, after which revised versions have been approved [at a Departmental Guideline meeting in Jan 2009 and via circulation [August 2012].

Following a clinical incident review in 2017 a further amendment was made to guidance on clotting studies in surgical cases, based on a literature review and multi-disciplinary consensus meeting [Drs Arora, Dyke, Lyall, Mr England, Mr Mathur]. In 2019 the

Trust Guidelines for the Use of Blood Products in Newborn Infants

document was reloaded to incorporate the hyperlinks to relevant forms. In 2021 a further revision was made to platelet transfusion thresholds based on the publication of RCT evidence

This version has been endorsed by the Clinical Guidelines Assessment Panel.

8. Distribution list/ dissemination method

- Hospital intranet.
- Neonatal Unit.

9. References

Alagappan et al. Impact of transfusion guidelines on neonatal transfusions. J Perinatol 1998;18:92-7

Altuntas N, Yenicesu I, Beken S, Kulali F, Burcu Belen F, Hirfanoglu IM, Onal E, Turkyilmaz C, Ergenekon E, Koc E, Atalay Y. Clinical use of fresh-frozen plasma in neonatal intensive care unit. Transfus Apher Sci. 2012 Aug;47(1):91-4. Epub 2012 May 27

Andrew et al. A Randomised Controlled Trial of platelet transfusions in thrombocytopenic premature infants. J Pediatr 1993;123:285-291

Andrew et al. Clinical impact of neonatal thrombocytopenia. J Pediatr 1987 Mar; 110 (3): 457-64

Andrew M, Vegh P, Caco C, Kirpalani H, Jefferies A, Ohlsson A, Watts J, Saigal S, Milner R, Wang E.
A randomized, controlled trial of platelet transfusions in thrombocytopenic premature infants . J Pediatr. 1993 Aug;123(2):285-91

Balegar V KK, Kluckow M. Furosemide for packed red cell transfusion in preterm infants: a randomized controlled trial. J Pediatr. 2011 Dec;159(6):913-8.e1. Epub 2011 Jul 23

BCSH (2004). Transfusion guidelines for neonates and older children. British Journal of Haematology, 124, 433-453

Beiner et al. Risk factors for neonatal thrombocytopenia in preterm infants. Am J Perinatol 2003;20(1), 49-54

Bussel JB, Primiani A. fetal and neonatal alloimmune thrombocytopenia: progress and ongoing debates. Blood Rev 2008 Jan;22(1):33-52

Calhoun et al. Consistent Approaches to Procedures and Practices in Neonatal Hematology. Clinics in Perinatology 2000;27(3):733-753

Cartlidge PH, Rutter N. Serum albumin concentrations and oedema in the newborn. Arch Dis Child 1986 Jul;61(7):657-60

Chakravorty S, Roberts I. How I manage neonatal thrombocytopenia. Br J Haematol. 2012 Jan;156(2):155-62. doi: 10.1111/j.1365-2141.2011.08892.x. Epub 2011 Sep 27

Trust Guidelines for the Use of Blood Products in Newborn Infants

Chirico G, Beccagutti F, Sorlini A, Motta M, Perrone B. Red blood cell transfusion in preterm infants: restrictive versus liberal policy. Cochrane Database of Systematic Reviews 2011, Issue 11. Art. No.: CD000512. DOI: 10.1002/14651858.CD000512.pub2. Low versus high haemoglobin concentration threshold for blood transfusion for preventing morbidity and mortality in very low birth weight infants. J Matern Fetal Neonatal Med. 2011 Oct;24 Suppl 1:20-2

Christensen RD. Platelet transfusion in the neonatal intensive care unit: benefits, risks, alternatives. Neonatology. 2011;100(3):311-8. Epub 2011 Oct 3

Curley A et al. Randomised Trial of Platelet-Transfusion thresholds in Neonates N Engl J Med 2019 Jan 17;380(3): 242-251

Del Vecchio A, Motta M. Evidence-based platelet transfusion recommendations in neonates. J Matern Fetal Neonatal Med. 2011 Oct;24 Suppl 1:38-40. Epub 2011 Aug 31

De Vos et al. Epidemiology and management of fetal and neonatal alloimmune thrombocytopenia. Transfus Apher Sci 2020;59:102704 Epub 2019 Dec

Dani C, Poggi C, Ceciari F, Bertini G, Pratesi S, Rubaltelli FF. Coagulopathy screening and early plasma treatment for the prevention of intraventricular hemorrhage in preterm infants. Transfusion. 2009 Dec;49(12):2637-44. Epub 2009 Jul 22

Fredrickson LK, Bell EF, Cress GA, Johnson KJ, Zimmerman MB, Mahoney LT, Widness JA, Strauss RG. Acute physiological effects of packed red blood cell transfusion in preterm infants with different degrees of anaemia. Arch Dis Child Fetal Neonatal Ed. 2011 Jul;96(4):F249-53. Epub 2010 Nov 20.

Goldenberg NA, Manco-Johnson MJ. Paediatric hemostasis and use of plasma components. Best Pract Res Clin Haematol 2006;19(1):143-55

Holzauer S, Zieger B. Diagnosis and management of neonatal thrombocytopenia. Semin Fetal Neonatal Med. 2011 Dec;16(6):305-10. Epub 2011 Aug 10.

Jardine LA, Jenkins-Manning S, Davies MW. Albumin infusion for low serum albumin in preterm newborn infants. Cochrane Database Syst Rev 2004;(3):CD004208

Kuperman AA, Kenet G, Papadakis E, Brenner B. Intraventricular hemorrhage in preterm infants: coagulation perspectives. Semin Thromb Hemost. 2011 Oct;37(7):730-6. Epub 2011 Dec 20.

Lippi G et al. Routine coagulation tests in newborns and young infants. J Thromb Thrombolysis 2007 Oct;24(2):153-5

Maier et al. Changing practices of red blood cell transfusions in infants with birth weights less than 1000g. J Pediatr 2000;136:220-4

Motta M, Del Vecchio A, Radicioni M. Clinical use of fresh-frozen plasma and cryoprecipitate in neonatal intensive care unit. J Matern Fetal Neonatal Med. 2011 Oct;24 Suppl 1:129-31.

Murray NA, Roberts JAG. Neonatal transfusion practice. Arch Dis child fetal neonatal Ed 2004; 89: F101-F107.

Trust Guidelines for the Use of Blood Products in Newborn Infants

Murray NA. Evaluation and treatment of thrombocytopenia in the neonatal intensive care unit. *Acta Paediatr Suppl* 2002;91(438):74-81

Murray et al. Platelet transfusion in the management of severe thrombocytopenia in neonatal intensive care unit patients. *Transfusion Medicine*, 2002, 12, 35-41.

Muthukumar P, Venkatesh V, Curley A, Kahan BC, Choo L, Ballard S, Clarke P, Watts Md T, Roberts I, Stanworth S; for the Platelets [Neonatal Transfusion](#) Study Group . Severe thrombocytopenia and patterns of bleeding in neonates: results from a prospective observational study and implications for use of platelet transfusions. *Transfus Med.* 2012 Jun 27;9999(9999). doi: 10.1111/j.1365-3148.2012.01171.x. [Epub ahead of print]

New HV et al on behalf of the British Committee for Standards in Haematology. Guidelines on transfusion for fetuses, neonates and older children *BJH* 2016;175:784–828

Nopoulos PC, Conrad AL, Bell EF, Strauss RG, Widness JA, Magnotta VA, Zimmerman MB, Georgieff MK, Lindgren SD, Richman LC. Long-term outcome of brain structure in premature infants: effects of liberal vs restricted red blood cell transfusions. *Arch Pediatr Adolesc Med.* 2011 May;165(5):443-50. Epub 2011 Jan 3.

Ouwehand WH, Smith G, Ranasinghe E. Management of severe alloimmune thrombocytopenia in the newborn. *Arch Dis Child fetal neonatal Ed* 2000; 82: F173-F175.

Plaisant F Evolution of neonatal transfusion practices: current recommendations. *Transfus Clin Biol.* 2011 Apr;18(2):262-8. Epub 2011 Mar 25.

Poterjoy BS, Josephson CD. Platelets, frozen plasma, and cryoprecipitate: what is the clinical evidence for their use in the neonatal intensive care unit? *Semin Perinatol.* 2009 Feb;33(1):66-74.

Rebulla P, platelet transfusion trigger in difficult patients. *Transfus clin Biol* 2002 Jan;9(1):109.

Reverdiau-Moalic P et al. Evolution of Blood Coagulation Activators and Inhibitors in the Healthy Human Fetus. www.bloodjournal.org July 19, 2017

Roberts IA, Murray NA. Management of thrombocytopenia in Neonates *Br J Haematology* 1999;105:864-870

Roberts IA, Murray NA. Thrombocytopenia in the newborn. *Curr Opin Pediatr* 2003;15(1):17-23

Roberts I, Murray NA. Neonatal thrombocytopenia. *Semin Fetal Neonatal Med.* 2008 Aug;13(4):256-64. Epub 2008 Apr 18.

Simon TL et al. Practice parameter for the use of red blood cell transfusions: developed by the Red Blood Cell Administration Practice Guideline Development Task Force of the College of American Pathologists. *Arch Pathol Lab Med* 1998;122(2):130-8

So KW, Fok TF, Ng PC, Wong WW, Cheung KL. Randomised controlled trial of colloid or crystalloid in hypotensive preterm infants. *Arch Dis Child, Neonatal Ed* 1997;76(1):F43-6

Trust Guidelines for the Use of Blood Products in Newborn Infants

Transfusion Task Force. Amendments and corrections to the 'Transfusion Guidelines for neonates and older children' (BCSH, 2004a); and to the 'Guidelines for the use of fresh frozen plasma, cryoprecipitate and cryosupernatant' (BCSH, 2004b). Br J Haematol. 2007 Feb;136(3):514-6.

von Lindern JS, Khodabux CM, Hack KE, van Haastert IC, Koopman-Esseboom C, van Zwieten PH, Brand A, Walther FJ. Long-term outcome in relationship to neonatal transfusion volume in extremely premature infants: a comparative cohort study. BMC Pediatr. 2011 May 28;11:48.

von Lindern JS, Hulzebos CV, Bos AF, Brand A, Walther FJ, Lopriore E. Thrombocytopaenia and intraventricular haemorrhage in very premature infants: a tale of two cities. Arch Dis Child Fetal Neonatal Ed. 2012 Jan 31. [Epub ahead of print]

Westkamp E et al. Blood transfusion in anemic infants with apnea of prematurity. Biol Neonate. 2002; 82 (4): 228-32.

Williams MD, Chalmers EA, Gibson BE; Haemostasis and Thrombosis Task Force, British Committee for Standards in Haematology. The investigation and management of neonatal haemostasis and thrombosis. Br J Haematol 2002;119(2):295-309

Williamson LM et al. The Natural History of feto-maternal alloimmunisation to the platelet-specific antigen HPA-1a as determined by antenatal screening. Blood 1998;92:2280-7

10 Source documents

Performing Exchange Transfusion in Newborn [Trustdocs id 1304](#)

Trust Guideline for the Management: of Massive Blood Loss in Children (C-MBL)
[Trustdocs ID: 9960](#)

Trust Guideline for the Use of Irradiated Blood and Blood Products, [Trustdocs ID: 1286](#)

Trust Guidelines for the Use of Blood Products in Newborn Infants

Appendix 1

Source and preparation of Neonatal Blood products

Components for neonates are prepared from blood donated by donors who have given at least 1 donation in the past 2 years, and which was negative for all mandatory markers. Ideally the donors should be male.

Red cells should be:

- Preferably suspended in SAG-M (except for those used for exchange transfusion).
- ABO and RhD compatible with infant.
- K negative.
- Free from clinically significant antibodies including high titre anti A and anti B.
- CMV antibody negative.
- HbS screen negative if large volumes are used.
- Irradiated if necessary.

Platelet transfusions should be:

- Free from clinically significant antibodies including high titre anti A and anti B.
- CMV antibody negative.
- Preferably collected by apheresis to limit donor exposure; pooled platelets may be used if apheresis not available.
- Irradiated if necessary.

Fresh frozen plasma should be

- Manufactured from whole blood or apheresis plasma from volunteer male donors.
- Sourced from abroad (currently USA).
- Treated with methylene blue to inactivate viruses.

Cryoprecipitate

- For neonates and infants is derived from US sourced plasma that has been MB treated.

*****For form see

[NICU Blood Transfusion Form \(Trustdocs id 11360\) or click here](#)

Form developed by Karen Hinchley, Neonatal Nurse



Norfolk and Norwich University Hospitals **NHS**
NHS Foundation Trust

D

NICU Blood Transfusion form

Name:.....

Hospital Number:.....

Date/...../20

| | |
|--------------------|---|
| <i>Affix label</i> | Pre transfusion Hb:g/L Target Hb: 160g/L (in most cases) Desired rise in Hb g/L x 0.4 x Wt (Kg) or 20 mL/Kg Amount to be transfused (use greater amount)..... Pack number |
|--------------------|---|

| | |
|--------|---|
| Yes/No | Does the patient need a new cross match for change of name? |
| | Parents informed? |
| | Has blood been ordered and written up? |
| | Do you have a site to give blood through? |
| | Do you need furosemide? |
| | Is there an ID barcode on cont/baby? |
| | Pick up slip from EBTS |
| | Once blood collected |
| | Is the NNST due and |

Example

| | Respirations | Saturations | BP |
|---|--------------|-------------|----|
| Pre transfusion observations | | | |
| Observations 15 minutes into transfusion. | | | |
| Observations 30 minutes into transfusion. | | | |
| Observations 45 minutes into transfusion. | | | |
| Observations 1 hour into transfusion. | | | |
| Observations 2 hours into transfusion. | | | |
| Furosemide administered? Yes/No | | | |
| Observations 3 hours into transfusion. | | | |
| Observations when transfusion complete. | | | |

Pump number:

| Time | Rate | Total | Site |
|------|------|-------|------|
| | | | |
| | | | |
| | | | |
| | | | |

| | |
|-------------|---|
| Yes/No | Time completed documented on drug chart |
| | Any reaction noted? |
| SIGN | Completed transfusion on EBTS (ended) |

For actual document click here [Trustdocs ID 16379](#)



Norfolk and Norwich University Hospitals **NHS**
NHS Foundation Trust

D

NICU FFP form

Name:.....

Hospital Number:.....

Date: dd/mm/yyyy/...../20

Affix label

Circle Doctor / ANNP

Clinical indication for FFP.....
Pre transfusion INR:
Amount to be transfused (20mls/kg).....
Discussed with Consultant/documentated

| Yes / No | Nursing staff |
|----------|---|
| | Does the patient need a new cross match for change of name? |
| | Parents informed? |
| | Has FFP been ordered and written up? |
| | Do you have a site to give FFP through? |
| | Is there an ID barcode on cot/baby? |
| | Pick up slip from EBTS (Electronic Blood Tracking System) |
| | Once FFP collected double check and start transfusion on EBTS |
| | Pack number |

Example

| Nursing Staff | Respirations | Saturations | BP |
|---|--------------|-------------|----|
| Pre transfusion observations | | | |
| Observations 15 minutes into transfusion. | | | |
| Observations 30 minutes into transfusion. | | | |
| Observations 45 minutes into transfusion. | | | |
| Observations 1 hour into transfusion. | | | |
| Observations at end of transfusion | | | |

Pump number:

| Time 24 hour clock | Rate | Total | Site |
|--------------------|------|-------|------|
| | | | |

| | |
|--------------------------|--|
| Yes/No | |
| | Time completed documented on drug chart |
| | Any reaction noted? |
| Print name and Signature | |
| | Completed transfusion on EBTS (ended) |
| Doctor/ANNP | Post FFP transfusion clotting studies performed <input type="checkbox"/> |