

Serious Hazards of Transfusion Paediatric Data

Paula Bolton-Maggs, Medical Director

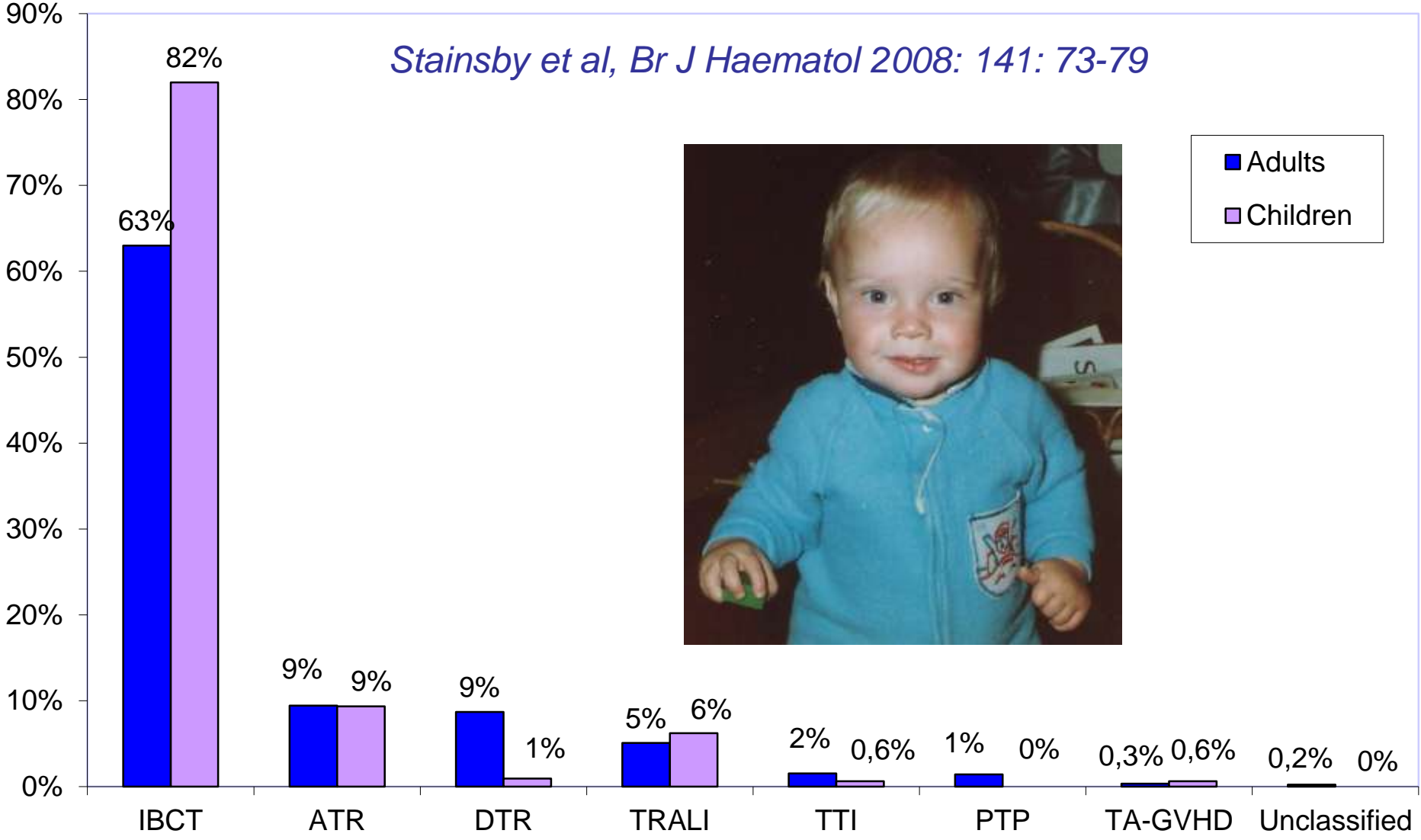
Helen New, Consultant Haematologist

SHOT

IHS April 2012

Adverse reactions and events in adults vs children

Stainsby et al, Br J Haematol 2008: 141: 73-79



■ Adults
■ Children

Risk of adverse outcome of transfusion in children vs adults

Population-based epidemiological study 2004

- 4.2% red cells transfused to patients <18 yr
- 1.7% to infants <12 months

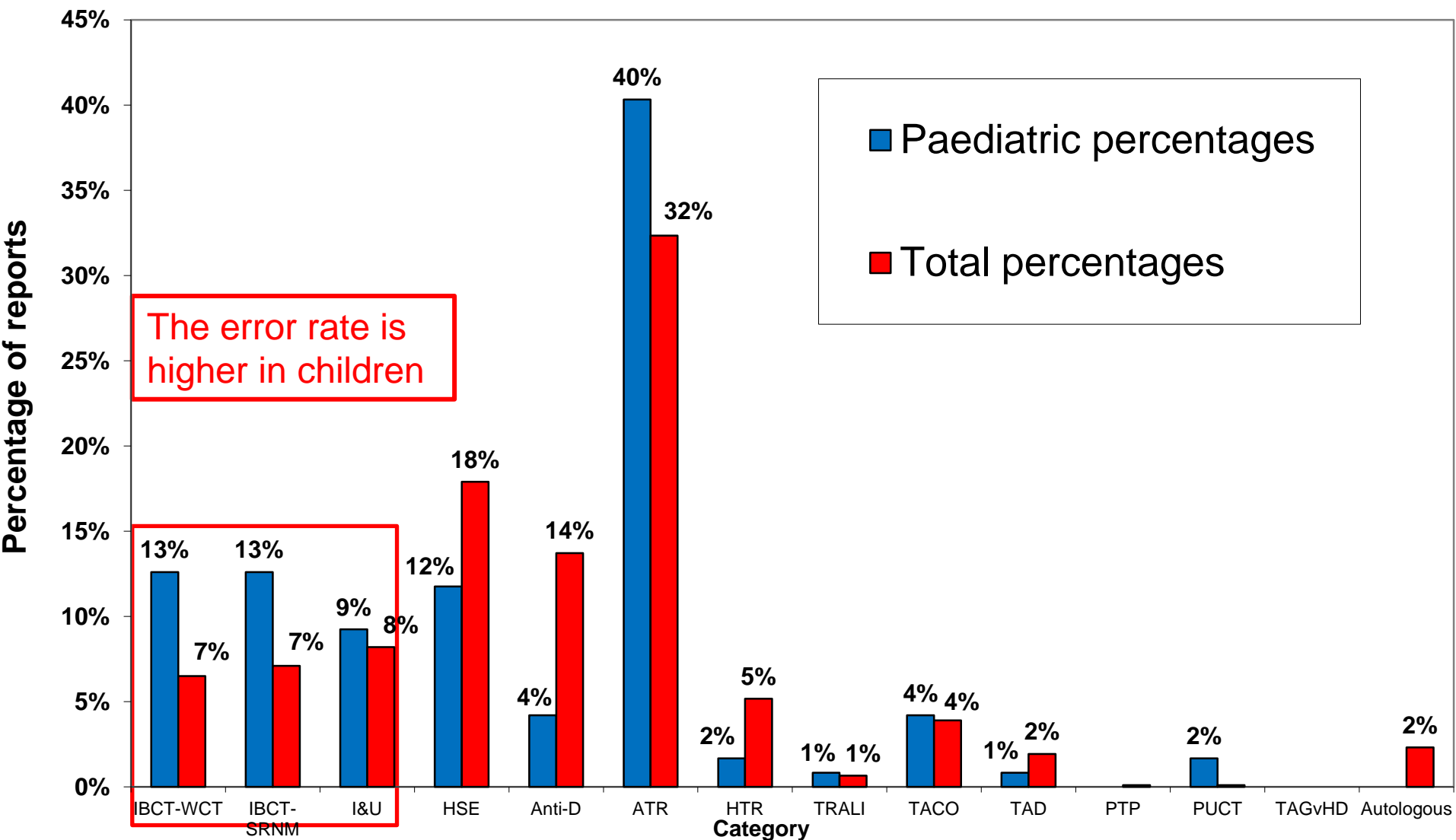
Incidence of adverse outcome of blood transfusion

per 100,000 red cells issued

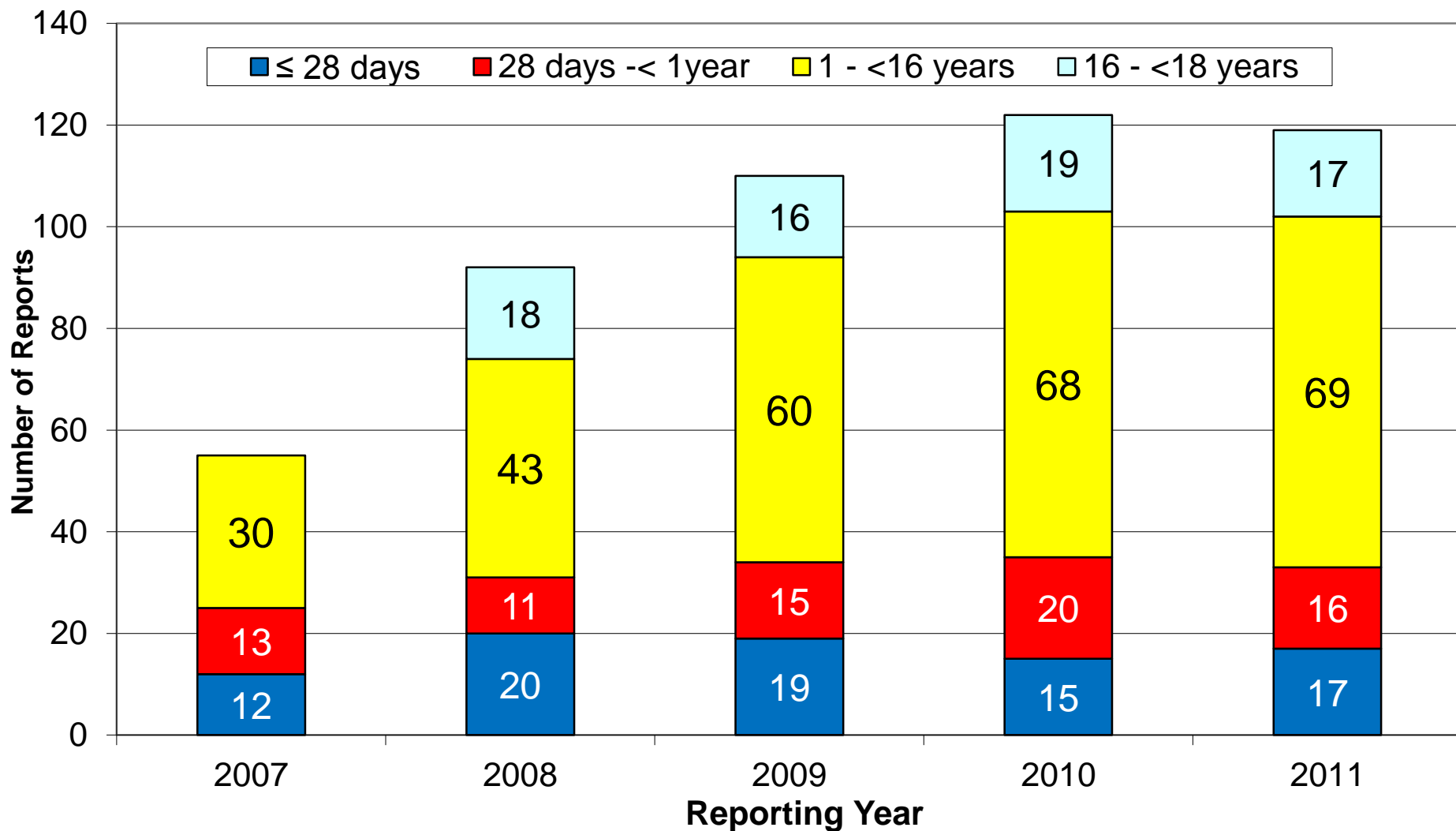
- Children <18 yrs 18
- Infants <12 mths 37
- Adults 13

Stainsby et al, Br J Haematol 2008: 141: 73-79

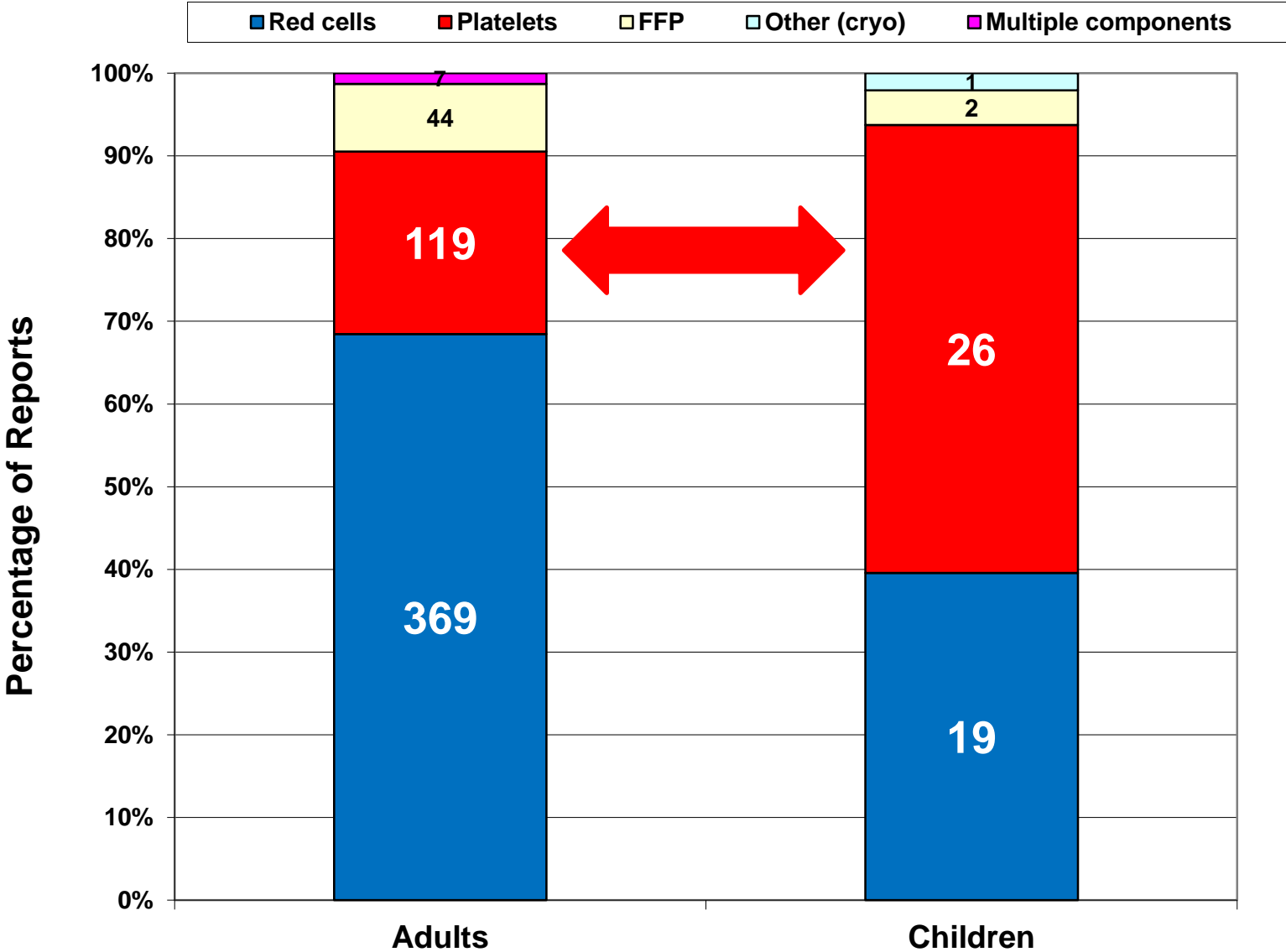
Reports - Paediatrics vs Adults 2011



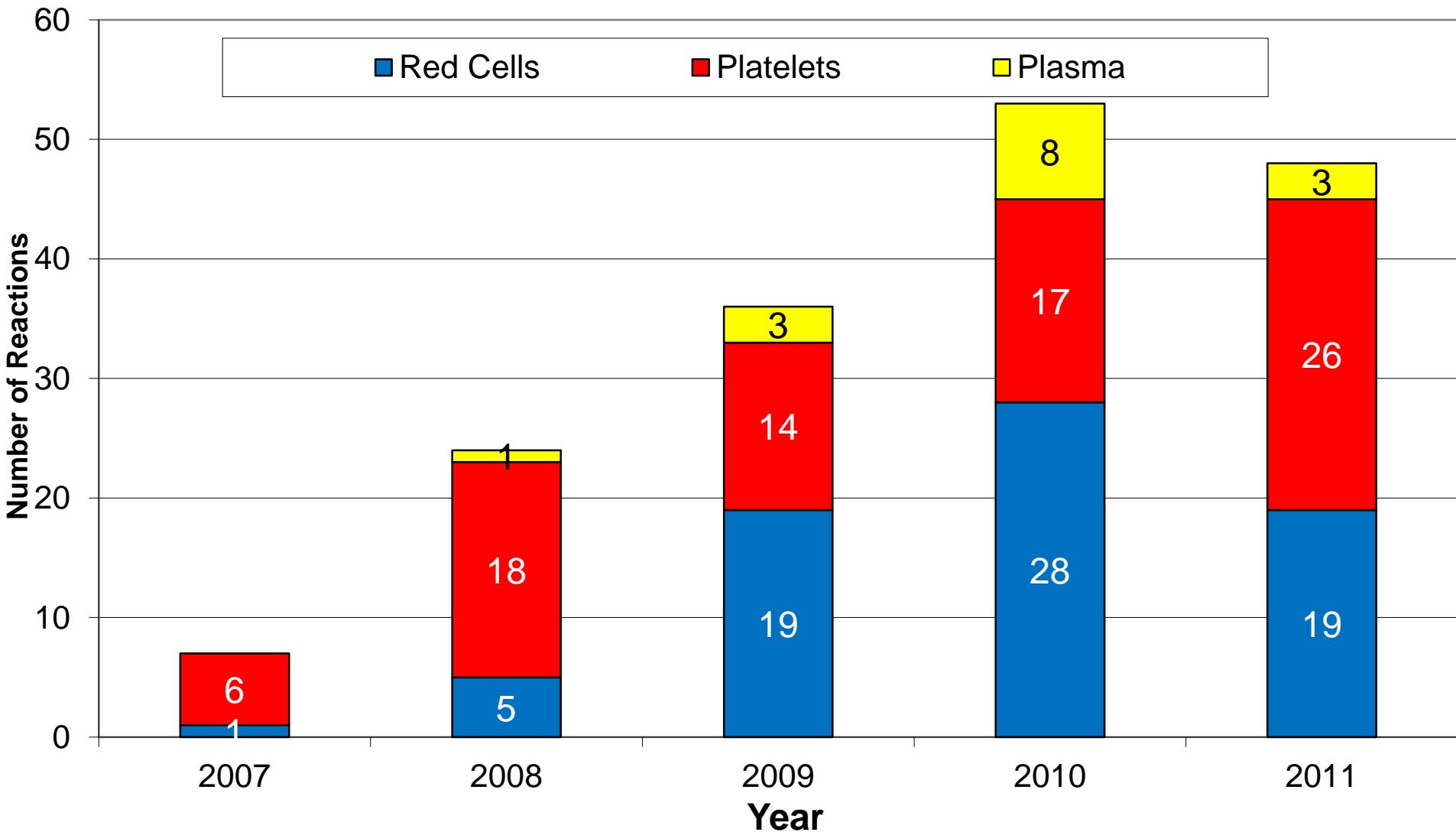
Cumulative paediatric data by age groups 2007-2011



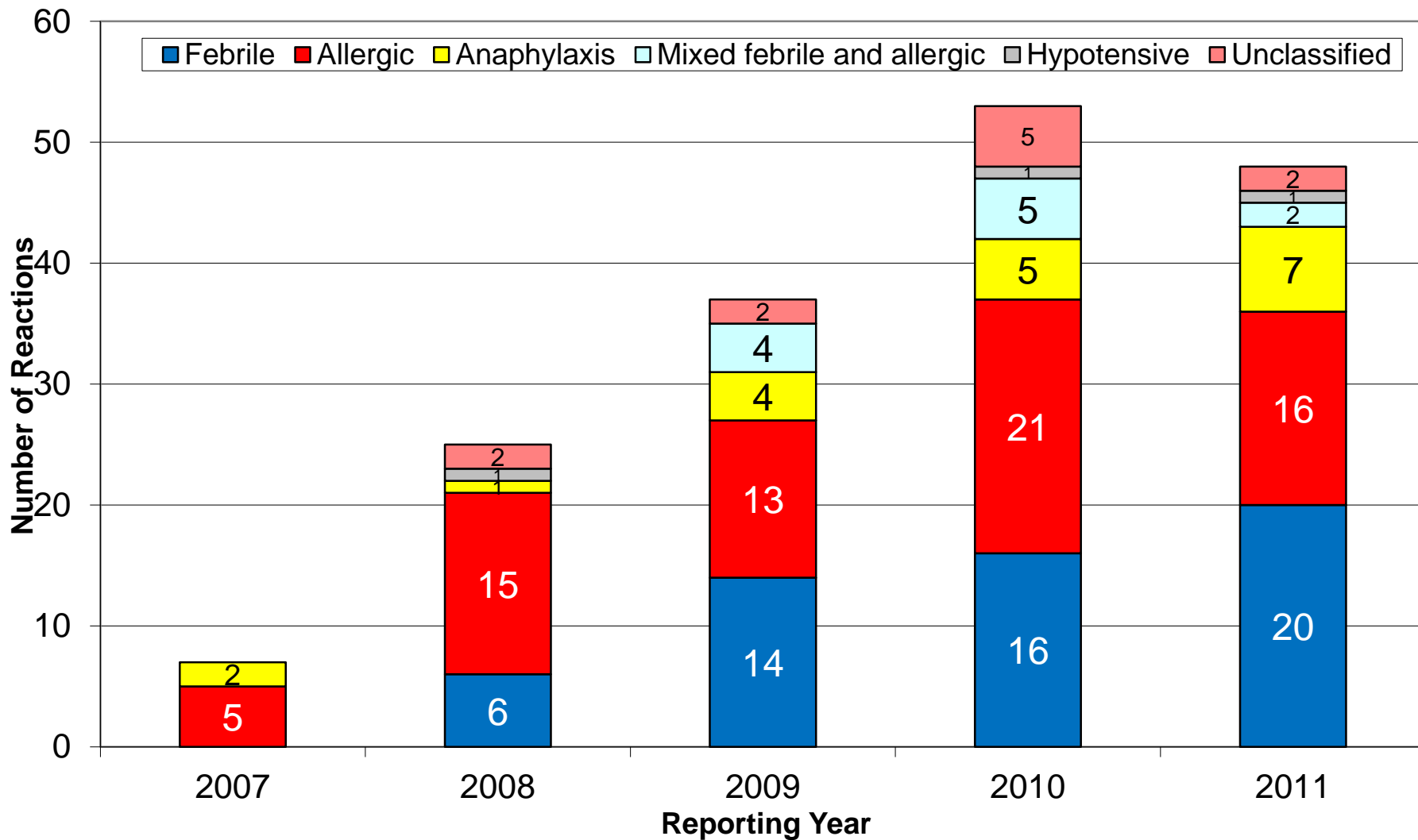
Comparison between adults and children – reactions to different components 2011



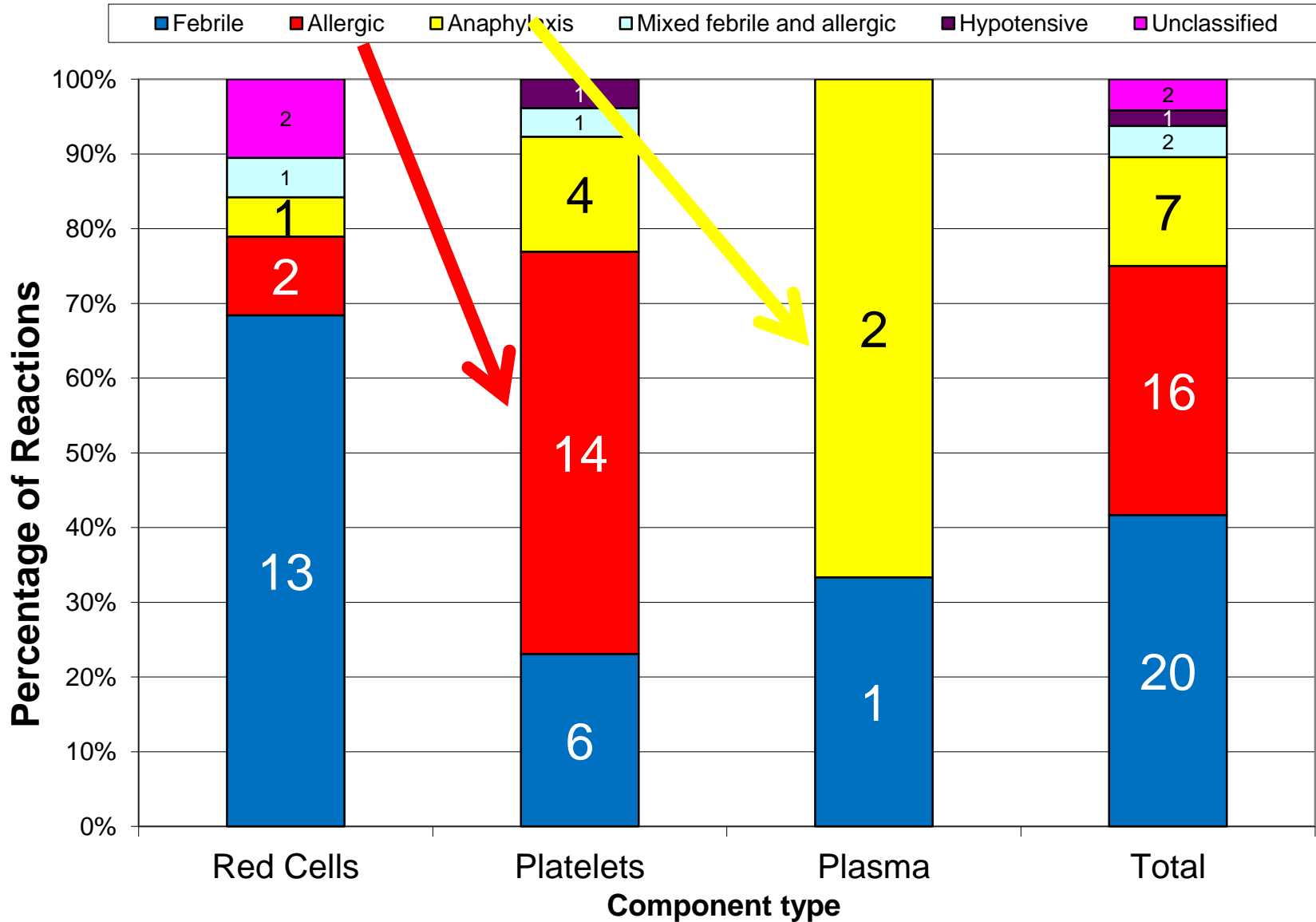
Cumulative paediatric reports by component type 2007-2011 (excluding multiple components)



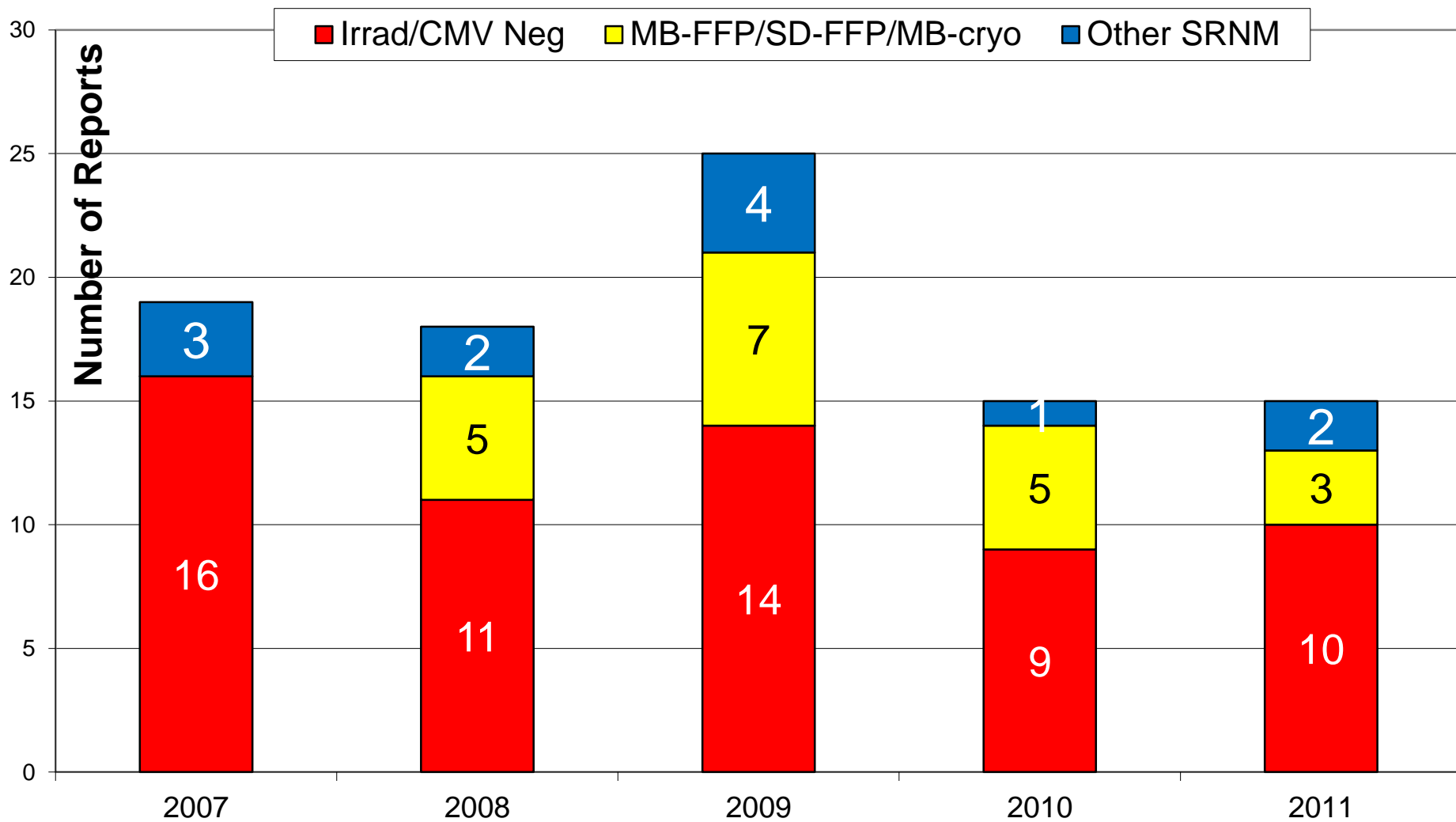
Cumulative paediatric reports by reaction type 2007-2011



Paediatric Reactions by component type 2011



Paediatric cases where the special requirements were not met 2007-2011



New observations in 2011

- Transfusion-associated circulatory overload
 - 5 cases aged from neonatal to 17yrs
- Two cases of necrotising enterocolitis possibly related to transfusion
 - 1 death
 - 1 needed surgery and survived

Some examples

Case study: Reaction to SD-FFP

- *A male infant with a congenital coagulation deficiency received SD-FFP to treat a cerebral bleed, and experienced a severe anaphylactic reaction within 30 minutes of starting the transfusion, with tachycardia, hypoxia and hypotension.*
- *He required intubation and was given adrenaline.*
- *He was subsequently given MB-FFP to treat the continuing bleeding problems. On one occasion, his oxygen saturation dropped again, but otherwise he experienced no problems*
- *He continues to receive MB-FFP without problems. Investigations for the cause of anaphylaxis proved negative.*

Case Study:

Necrotising enterocolitis post transfusion

- *A clinically stable non-ventilated 6 week old preterm infant, born at 26 weeks gestation, was given a red cell transfusion for symptomatic anaemia of prematurity (Hb 9.3 g/dL).*
- *There were no adverse events during the transfusion, and the post Hb was 16.7 g/dL. 4.5 hrs post transfusion the baby developed tachycardia, and over the next 12 hours deteriorated and developed a distended abdomen.*
- *An X-ray was consistent with NEC, the baby continued to deteriorate and died at approximately 36 hrs post transfusion.*

Case study

Lack of awareness of the need for irradiated blood following IUT

- *A baby who had received IUT for HDN was admitted at age 7 wks with Hb 4.4 g/dL and transfused with a non-irradiated paedipack*
- *Neither request form or prescription indicated that irradiated blood was required*
- *The laboratory SOP was unclear and the BMS believed there was no need to irradiate top-ups following IUT*
- *Nursing staff did not notice that irradiated blood was required*

Case study

Doctor unaware of neonatal specification units in satellite refrigerator

- *Premature baby Hb 6.2g/dL following emergency caesarean section delivery*
- *Staff grade doctor borrowed midwife's blood fridge access card*
- *Removed a unit of adult O RhD negative, NOT the paediatric emergency O RhD negative blood that was also present*
- *Baby received 100mL without any adverse reaction*

Case study

Baby with HDN due to anti-c given O RhD negative blood

- *Baby born by emergency caesarean section, suffering from HDN due to high titre maternal anti-c*
- *Removed emergency O RhD negative unit for transfusion without informing the laboratory*
- *Baby suffered an immediate (mild) reaction, which fully resolved*
- *Bilirubin climbed further, requiring exchange, and this may well have been exacerbated by the incompatible transfusion*

Case studies

Clinically significant over-transfusion

- *12-month old child on PICU required 110mL red cells*
 - *Adult unit supplied, and nursing staff transfused all 230mL*
 - *Post-transfusion Hb was 19g/dL, requiring venesection*
-

- *6-month old infant on ICU required 140mL red cells post-op*
- *Nurse asked doctor if she should give '1 unit' and he verbally agreed*
- *Entire unit (257mL) was transfused*
- *No adverse outcome apart from excessive flushing*

Case study

Confusion during the collection process

- *A preterm baby required an emergency transfusion at 6 days of life and should have been given O RhD negative emergency blood from the satellite fridge. The nurse inadvertently collected an adult O RhD negative unit that had been issued for an obstetric patient on the delivery suite.*
- *The blood group and CMV status was checked with another nurse, but **neither** noticed that the tag on the unit had a compatibility label on as opposed to an 'emergency blood' label*

Case study

Incorrect pre-transfusion compatibility testing procedures

- *A G&S/DAT request was received in the laboratory for a newborn baby with a low Hb. Blood was requested later that day and twice more, 2 days later*
- *The first two requests were treated as Electronic Issue, but the third was fully crossmatched*
- *Mother was known to have an antibody, so ALL requests should have been crossmatched*

Case study

Administration error results in over-transfusion

- *A 24-day-old baby was prescribed a transfusion of 14.3mL red cells. It was noted that the baby's Hb rose from 9.7g/dL to 20.0 g/dL*
- *It was noticed that the paedipack was empty, meaning that the baby had received ~50ml blood*
- *The roller clamp in the neonatal Y giving set had not been closed*

Paediatric Commentary 2010-11

- Poor understanding by lab staff of procedures for pre-transfusion compatibility testing
- Confusion among clinical staff as to blood availability for emergency transfusion
- Over-transfusion of children – prescribing in ‘units’ rather than mL
- Ensuring giving sets are fit for purpose and transfusions are monitored throughout
- Paediatric ATR reports are increasing, particularly febrile reactions with red cells

Lessons (1)

- Mistakes happen even in areas where there is 'one-to-one' care
- Specific education of staff in paediatric transfusion practice is crucial
- Wearing and checking of patient ID is essential
- Prescription - needs volume and duration of transfusion written down – no verbal instructions

Lessons (2)

- BCSH guidelines should be followed
- Closely monitor children for reactions
- Care with administration – nursing staff must be skilled and competent in the use of infusion devices, appropriate rates and volumes for transfusion, and special requirements
- Good communication is vital, between lab and clinicians and between institutions sharing care
- Avoid unnecessary transfusion – especially of FFP and platelets

Paediatric Recommendations

- Prescribing only by those with appropriate knowledge and expertise
- Particular care for special requirements, including documentation, and communication – use of specific paediatric prescription charts
- Lab BMSs must be aware of special component requirements for <16s, and routine checking for additional flags based on DOB
- Encourage clinical staff to report reactions

Paediatric Haemovigilance

- Uncertainty about the true nature and extent of adverse transfusion outcomes in children, particularly neonates and infants
- Likely to be under-reporting to SHOT
 - Signs may be more subtle due to immunological immaturity, may be masked by symptoms, or simply not recognised
 - Some reactions may not be clearly defined as relating to transfusions *eg necrotising enterocolitis*
 - Other complications, such as line-associated infections, problems with multiple cannulations or extravasations are not currently reported to SHOT

Neonatal transfusions

- Highly transfused, potential of long life
- Appropriate transfusion triggers not clear
- Mixed evidence on outcomes
 - Association between platelet transfusions and hepatic dysfunction
 - 9-fold increase in bacterial infection in neonates who had received >10 platelet transfusions

Audit of adverse outcomes associated with neonatal transfusion

- Many unknowns
- What are the adverse events?
- What to monitor
- Relationship to NEC
- All transfusions in NICU or NNU in participating centres



Audit Management Group

- Neonatology
 - Anna Curley
 - Vidheya Venkatesh
 - Rizwan Khan
- NHSBT
 - Helen New (Clinical Lead)
 - Simon Stanworth
- SHOT
 - Paula Bolton-Maggs
 - Tony Davies
- Database
 - Debbi Poles (SHOT)
- Statistician
 - Linda Hunt (NHSBT)

Objectives & Audit Plan

- Standardise a Transfusion Assessment Audit Tool (TAT) and use the TAT to;
 - Define the level of conventionally recognised acute clinical adverse outcomes
 - Define previously unrecognised adverse outcomes
 - Capture the level of additional events such as cannulation / extravasation
- Systematically record clinical information already being routinely collected during and 6 hrs post transfusion, with extra time points up to 24hrs post transfusion

Prospective Observational Survey of Clinical Adverse Outcomes related to Transfusion in Neonates version0.15
Questionnaire for Red cell transfusions

XXX 111	Audit number: _____
For office use only:	
Date received: _____	Date entered: _____

PATIENT DETAILS

Date of birth in dd/mm/yy: Gender (please tick): Male Female

Gestation at birth: Weeks Days Birth weight in grams:

TRANSFUSION DETAILS

Date of transfusion in dd/mm/yy:

Transfusion indication:
 Anaemia of prematurity Pulmonary Haemorrhage Suspected NEC Suspected Sepsis
 NEC requiring surgery HDN Acute blood loss Other (please state below)

Transfusion of other blood product in 24 hrs prior to transfusion?: Yes No

If yes, state which blood component(s)
 Platelets FFP Cryoprecipitate
 RBC IVIG Albumin

RESPIRATORY STATUS AT THE START OF TRANSFUSION Please tick all that apply

Mechanical Ventilation Was baby muscle relaxed? Yes No

Non-invasive Ventilation Supplementary Oxygen

Breathing in air without respiratory support

FLUID STATUS

Is infant on regular diuretics? Yes No

Weight (Record weight closest to) Weight in grams DD / MM / YY

Weight 72 hours pre transfusion	<input type="text"/>	<input type="text"/>	<input type="text"/>
Recent Weight recorded pre transfusion	<input type="text"/>	<input type="text"/>	<input type="text"/>
Weight 24 hour Post transfusion	<input type="text"/>	<input type="text"/>	<input type="text"/>

MBP measured by: Indwelling arterial line Peripheral cuff

XXX111

Time transfusion commenced :

Time transfusion completed :

MONITORED VITAL SIGNS DURING PACKED CELL TRANSFUSION

N.B: Please complete the 4 hours prior, 2 hours prior and 0 min as this forms the baseline. For the inotropes column Circle Y if on inotropes. Circle ↑ when dose is increased or another inotrope is added. Circle ↓ when dose decreased. If no change, do not circle either of the arrows. If ventilated record MAP (mean airway pressure. If on noninvasive ventilation record PEEP

Time point	Time 00:00	Temp (C°)	Heart Rate (bpm)	Inotropes				Spontaneous Respiratory Rate (bpm)	MAP cm H2O	PEEP cm H2O	FiO ₂ (%)	MBP mm Hg	Handling	If other product given tick box FFP; Cryo, PLT; RBC
4 hours prior				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
2 hours prior				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
0 mins				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
15 mins				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
30 mins				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
45 mins				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
60 mins				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
1 hr 15 mins				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
1 hr 30 mins				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
1 hr 45 mins				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
2 hrs				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
2 hrs 15 mins				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
2 hrs 30 mins				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
2 hrs 45 mins				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
3 hrs				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
3 hrs 15 mins				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
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5 hrs				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
6 hrs				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
7 hrs				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
8 hrs				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
9 hrs				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
10 hrs				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
12 hrs				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
16 hrs				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
20 hrs				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
24 hrs				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC
28 hrs				Y	N	↑	↓					Y	N	<input type="checkbox"/> FFP <input type="checkbox"/> Cryo <input type="checkbox"/> P LT <input type="checkbox"/> RBC

Was a new cannula replacement required during the transfusion?

Yes No

If so, was there any evidence of extravasation from the transfusion?

Yes No

Please tick all that applies

	During transfusion		In the first 6 hours post transfusion		In the 6-24 hours post transfusion	
Increased frequency of apnea with bradycardia compared with 24 hours pretransfusion	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>
Initiation of a new mode of ventilation	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>
Increase in FiO2 of >10% for ≥ 15 minutes compared with average 0, 2, and 4 hours pretransfusion			Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>
Increase in respiratory rate by > 15 per minute for ≥ 15 minutes compared with average 0, 2, and 4 hours pre transfusion			Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>
Additional diuretic given (apart from any regular diuretics that baby was on pretransfusion)	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>
Rise in temperature by more than 1°C			Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>
Urticaria or hive like rash.	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>
Evidence of severe allergic reaction or anaphylaxis (hypotension with rash, dyspnoea, stridor, wheeze, angioedema)	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>
Development of signs and symptoms of haemolysis (fall in Hb, rise in LDH, rise in bilirubin, positive DAT)	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>
Was a Chest X ray was taken?	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>
If Yes: Were there new bilateral pulmonary infiltrates on Chest X ray?	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>
Was there evidence of cardiomegaly (cardiac silhouette> 60%)?	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>
Were there any other findings on chest X ray?	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>	Yes <input type="checkbox"/>	No <input type="checkbox"/>
If Yes, Please describe:						

Specific Measures of Outcome

- Respiratory deterioration compared to pre-transfusion average readings
 - Increase in FiO₂ of >10% for >15min
 - Increase of mean airway pressure of >2cm H₂O
 - A new mode of ventilation initiated by worsening lung condition
- Temperature change >2°C
- Mean Arterial Pressure >2 cm H₂O

HV Study – the next steps

- Pilot data to be ‘cleaned’ prior to further analysis and specific outcomes refined
- Cambridge team to continue to collect data – aiming for 200 babies
- Analyse preliminary results – perhaps refine definitions of Paediatric categories
- Extend study to other UK centres
- Interest from US and Australia

Dr Tony Davies, Transfusion Liaison Specialist
Dr David Hogg, Consultant Paediatric Haematologist
Dr Sarah-Jane Smith, Paediatric Transfusion Nurse

Why are children so important from a transfusion point of view?

Children are a vulnerable patient group and have special transfusion requirements. However, it is not their age which determines transfusion. The responsibility for children receiving a transfusion should be considered as a clinical activity, not that of an adult.

Children are especially vulnerable to the possible side-effects of transfusion. They have narrower plasma volume and are at risk of developing iron deficiency.

The acute side effects of transfusion may be greater for children than for adults, as a single unit of transfused plasma will be greater in mass than for an equivalent adult. Consideration of the long term side effects of transfusion for children is particularly important, as the majority will live for decades afterwards.

In general, paediatric transfusion is the responsibility of the paediatric haematologist, but common, and life saving, transfusion is the responsibility of the transfusion team.



In adults, 40% of events are BCT related, and in children 85% of severe events are BCT related, a disproportionate high number. Factors for greater components of the right specifications for transfusion are:

What are the transfusion hazards in children?

The British Society of Transfusion (BST) guidelines advise that the majority of transfusions are given to children with a component of iron deficiency. This is due to the high iron requirements of growing children.



Iron deficiency is a common condition in children, and is often associated with iron deficiency anaemia. The British Society of Transfusion (BST) guidelines advise that the majority of transfusions are given to children with a component of iron deficiency. This is due to the high iron requirements of growing children.

What information is available to children and parents about transfusion?

Like all medical treatments, a blood transfusion should only be given if it is needed. Respecting medical professionals will reduce the risk of a transfusion against the risk of not having one, and should explain to parents and older children why a transfusion may be necessary.



Appropriate use of blood and components in children

Appropriate use of blood components should be promoted and discussed. Transfusion is given for other reasons such as anaemia and it is generally difficult to incorporate adult data for neonates. Collection of iron balance and laboratory and clinical evidence is essential to guide the selection of blood component and guide transfusion.

Recent and planned studies to develop an evidence base for paediatric transfusion include:

- Research on cell transfusion
- Tera study: Bell et al, 2002, 'Red cells: Argonoff et al, 2006
- Red cell transfusion in PICU
- 'Blood' study: Lomas et al, 2007
- Neonatal platelet transfusion
- 'Platelet' study
- Neonatal FFP
- Neonatal FFP of thrombocytopenia in PICU
- NE LITe study: paediatric data analysis
- Transfusion Alternatives Program: to reduce cell blood
- TACT: The ongoing, randomised paediatric
- Comparison of FPG
- TACT: Adult ongoing
- Comparison of a Redbank Cardiac Surgery
- 2002, to commence shortly
- Neonatal platelet transfusion study (PACT 2) in preparation
- Paediatric Red cell M.O. trial 2007

Evaluation and discussion around paediatric transfusion

Recent evidence analysis has highlighted the need to improve the practice of transfusion and to ensure that transfusion is given to the right patient at the right time.

The NHSBT Paediatric Transfusion Group works regularly to discuss these and other issues around paediatric transfusion. It reviews guidelines and current practice, and its guidance on transfusion, laboratory and appropriate transfusion is available to all transfusion services.



BCSH guidelines for neonates and older children to be revised

Prospective study on transfusions in neonates is being carried out jointly between NHSBT Paed Group and SHOT



Conclusions 1

- Still too many errors
 - use of adult emergency O neg blood for neonates
 - laboratory errors in neonatal and maternal grouping and antibody screening,
 - failure to recognise the need for irradiated components after intrauterine transfusion
 - prescription and administration errors leading to either overtransfusion or incorrect rate of transfusion.
- Poor communication and lack of checking in I&U cases with poor clinical understanding of the transfusion process in paediatrics, including the need to administer a specific volume rather than an entire unit.

Conclusions 2

- Children developed TACO, illustrating the importance of prescribing the correct volume and rate for small infants and children.
- Two reports of NEC associated with transfusion in 2011. Prospective studies are needed to further investigate this association.
- ATRs occurred following paediatric platelet transfusion, including 4 anaphylactic reactions. As the majority of the platelet transfusions were given for prophylaxis rather than bleeding, we need to ensure that these are given according to guidelines

SHOT / RCA Toolkits

**Lessons for Laboratory
Staff**

RESOURCES

www.shotuk.org

Reports and Summaries

**Lessons for Clinical
Staff**

SHOT Symposium 2012

The Lowry Centre, Salford Quays

Thursday 5th July £69

e-mail shot@nhsbt.nhs.uk



