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**INTERNATIONAL FORUM** 

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# Vox Sanguinis International Forum on paediatric indications for blood component transfusion

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Meghan Delaney

## **United States of America**

Subject 1. Hospital and transfusion service demographics:

Question 1

United States of America.

Question 2

Yes.

Question 3

Both.

#### Question 4

Neonate is defined by transfusion service as <4 months according to US regulations (FDA, AABB) that is built around being able to waive additional cross-matching to avoid iatrogenic phlebotomy by using Group O red blood cells during this time period (birth to chronological 4 months of age). All of the patients at our facility would be considered 'pediatric' since we are a standalone tertiary/quaternary paediatric referral centre. We do not have 'adult' protocols, even though we may treat patients >18 years, they would fall under regular transfusion approach. The only exception to this may be if there is a patient >18 years who is refusing transfusion due to their wishes, if the patient is a legal minor (<18 years) or not (>18 years), the hospital would proceed differently.

#### Question 5

Yes, it is paediatric specialty hospital that has a NICU as well. We have 313 beds.

## Question 6

No, only neonates and paediatrics. We do treat young adults as well when there is a reason for the patient to see a physician with specialty in paediatric geared specialties (paediatric cancers, genetics, congenital heart disease, etc).

Question 7a

Yes.

Question 7b

Yes.

Question 7c

Yes.

Question 7d

r es.

Question 7e

Yes.

Question 7f

Yes.

Question 7g

Yes.

Question 7h

Yes.

Question 7i

Yes.

Question 7j

Yes.

Ouestion 7k

Yes.

Ouestion 71

Yes.

Ouestion 7m

Yes.

**Question** 7n

Yes.

Ouestion 70

Yes.

**Question 7p** 

Yes.

Ouestion 8a

RBCs: 8057 units (in 2017).

#### **Ouestion** 8b

I am not able to get this information without a long wait and IT request.

**Question** 9a

Plasma: 1760 (in 2017).

#### Question 9b

I am not able to get this information without a long wait and IT request.

## Ouestion 10

Platelets: 3525/year (Full-size units. We aliquot them into smaller doses for smaller patients.) (in 2017).

## Subject 2. Transfusion Indications for paediatric and neonatal patients:

## Ouestion 1

The United States does not have a national transfusion policy for neonates or pediatrics (or adults). The United States has never really had national-level guidelines on transfusion. There are initiatives through various networks and professional societies to create guidelines, but these are typically in specific populations or according to certain practices.

For instance, the PALISI network created a set of 10 manuscripts focused on evidence-based review and areas needed for research on transfusion indications and thresholds in paediatric critical care patients only (called the TAXI project). There are other AABB and ASCO guidelines, some of which mention paediatric patients and some do not.

#### Ouestion 2

Our hospital specialties do have transfusion thresholds for neonatal and/or paediatric transfusions based on diagnosis that are kept at the local level. My plan is to collate these empiric approaches into one transfusion policy at the hospital level. I was able to do this at Seattle Children's Hospital when I was Transfusion Medicine Director there.

#### **Question 3**

Our hospital specialties do have transfusion thresholds for ordering RBC/plasma/platelets based on diagnosis that are kept at the local level. My plan is to collate these empiric approaches into one transfusion policy at the hospital level. I was able to do this at Seattle Children's Hospital when I was Transfusion Medicine Director there.

#### Ouestion 4

Yes. Providers that are ordering blood product transfusion must electronically complete the 'Justification for transfusion' to order the blood.

#### **Ouestion** 5

All outpatient visits require a diagnosis, thus this is linked to transfusion in that setting. All transfusions (inpatient and outpatient) require the provider to enter 'Justification for transfusion.'

## **Question 6**

Our hospital specialties do have transfusion thresholds for RBC transfusion based on diagnosis that are kept at the local level. My plan is to collate these empiric approaches into one transfusion policy at the hospital level. I was able to do this at Seattle Children's Hospital when I was Transfusion Medicine Director there.

Our transfusion data card states, 'Increase oxygen- carrying capacity'

## Question 7

Our hospital specialties do have transfusion thresholds for platelet transfusion based on diagnosis that are kept at the local level. My plan is to collate these empiric approaches into one transfusion policy at the hospital level. I was able to do this at Seattle Children's Hospital when I was Transfusion Medicine Director there.

Our transfusion data card states, 'Correct/prevent bleeding due to thrombocytopenia'

## **Question 8**

Somewhat. Our hospital specialties do have transfusion thresholds for plasma transfusion based on diagnosis.

Our hospital specialties do have transfusion thresholds based on diagnosis that are kept at the local level. My plan is to collate these empiric approaches into one transfusion policy at the hospital level. I was able to do this at Seattle Children's Hospital when I was Transfusion Medicine Director there.

Our transfusion data card states, 'Coagulation factor replacement for which there is no specific factor concentrate'

## Subject 3. Product manipulations for paediatric and neonatal patients

#### **Question** 1

Yes, all platelets and RBCs are irradiated. This policy was put in place because of missed irradiation events in our population that is heavily enriched for patients that have a medical indication for irradiation. The only exception is if they are ordering emergency release blood when the provider signs to waive irradiation.

#### *Question 2*

Yes, all red blood cells are leucocyte reduced.

#### Ouestion 3

Yes, this is by transfusion medical director approval and typically only for prevention of severe allergic reactions.

#### Ouestion 4

No, not at this time, but we plan to in the future.

#### Ouestion 5

No, not at this time.

## Ouestion 6

Yes. We have a large and active sickle cell disease treatment programme and provide Rh/K for these patients. We also default to antigen-matched cells when a patient has certain serological findings, such as panagglutinin, but this is on the case-by-case basis by the transfusion medical and technical laboratory specialists (SBB).

## Question 7

No.

#### **Question 8**

Yes, we have multiple infants assigned to one red blood cell parent unit and we draw multiple aliquots off of them. We put up to five babies on Group O, RhD-positive or RhD-negative RBC unit. We select the unit that is  $\leq 10$  days of storage at the time of assignment.

#### **Question 9**

We do use small paediatric tubes whenever we can.

#### Ouestion 10

Yes, this is based on age of RBC product and time from irradiation. In general, this is what we follow:

| Patient age  | Maximum storage after irradiation   | Additional attributes  |
|--|---|--|
| Neonates<br>(<4 months)  | 24 h  | Fresh units ≤10 days old at time of transfusion O positive or negative   |
| Infants and young children (4 months to 5 years)                 | 72 h after day of irradiation (day 0)   |  |
| Older children and<br>adolescents/<br>adults (>5 years)          | 28 days post-<br>irradiation or<br>original expiration<br>date, whichever is<br>first |  |
| Any age patient:<br>ECMO, cardiac<br>surgery pump<br>prime units | Age-dependent<br>Minimize<br>postirradiation<br>storage time<br>whenever possible     | Fresh units <10 days old at<br>time of transfusion<br>Select <7-day-old units for<br>cardiac surgery pump prime<br>whenever inventory allows |

## Subject 4. Blood product dosing

## Ouestion 1

Patients <50 kg and sickle cell pts: 10–15 ml/kg Patients ≥50 kg: 1–2 units (5–10 ml/kg)

#### Question 2

Patients <5 kg: 1 EU.

Patients  $\geq$ 5 kg: 1 EU/5 kg (oncology/BMT/bleeding patient).

1 EU/10 kg.

Apheresis derived: 1 EU = equivalent to 1 random donor platelet or  $5.5 \times 10^{10}$  platelets in 25–50 ml plasma).

#### **Question 3**

Patients <50 kg: 10–15 ml/kg

Patients ≥50 kg: 2 units (5–10 ml/kg)

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## Czech Republic

Subject 1. Hospital and transfusion service demographics

Question 1 Czech Republic.

Question 2 Yes.

Question 3 In our facility.

Question 4 neonate: 30 days old.

paediatric: under 19 years old.

Question 5

A pediatric specialty hospital is a part of University Hospital Brno

**Question** 6

Yes, we treat both adult and paediatric patients and we have 400 inpatient beds.

Question 7a Yes.

Question 7b Yes.

Question 7c Yes.

Question 7d Yes.

Question 7e Yes.

Question 7f Yes.

Question 7g Yes.

Question 7h Yes.

**Question 7i** Yes.

Question 7j Yes.

**Question 7k** Yes.

Question 71 Yes.

Question 7m Yes.

Question 7n Yes.

Question 70 Yes.

Question 7p Yes.

Question 8a 1671 T.U.

Question 8b (a) 216 T.U. (b) 231 T.U. (c) 1224 T.U.

Question 9a

182 T.U. + 61 Octaplas.

**Question 9b** (a) 24 T.U. (b) 29 T.U. (c) 129 T.U.

Question 10a 921 T.D.

#### Question 10b

- (a) 40 T.D.
- (b) 24 T.D.
- (c) 857 T.D.

## Subject 2. Transfusion indications for paediatric and neonatal patients

#### Question 1

No. (There is only a chapter in the national guidelines. There is information about intrauterine or intraumbilical transfusions and doses for children.)

#### **Question 2**

No. (There is only a chapter about children and neonates indications of the blood components in the hospital recommendation).

#### Ouestion 3

No.

#### Ouestion 4

No.

#### Ouestion 5

No.

#### **Question** 6

Yes. The threshold for adults is 70–80 g/l. We have no exact recommendation for children.

## **Question 7**

Yes. Recommendation for adults: prophylactic application: usually under  $20 \times 10^9/l$ , therapeutic application: under  $80{\text -}100 \times 10^9/l$  – massive bleeding, cerebral insurance, under  $50 \times 10^9/l$  – serious bleeding gastrointestinal or urological, under  $30 \times 10^9/l$  – bleeding in muscles or skin. There is no special recommendation for children.

#### Question 8

No. We have a clinical definition.

## Subject 3. Product manipulations for paediatric and neonatal patients

## Question 1

Yes. Children under 1 year old, haemato oncology patients, immunodeficits.

### **Question 2**

Yes. All RBCs for paediatric and neonatal patients are leucodepleted.

## Question 3

Yes. IgA selective immunodeficit, serious allergic adverse events after application of blood components.

#### Question 4

No.

#### Ouestion 5

No.

### **Question** 6

Yes. haematoonkology patients.

## **Question 7**

No.

## Question 8

No.

#### Question 9

Yes.

#### **Question 10**

Yes. Polytraumatic patients maximally 14 days old, neonates 5 days old, paediatric 14 days old.

## Subject 4. Blood product dosing

## Question 1

5-10 ml/kg.

## **Question 2**

Standard paediatric dose (about 80-100 ml). Children under 15 kg - 10-20 ml/kg.

## Question 3

10–15 ml/kg. We applicate preferentially Octaplas to paediatrics and neonates.

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## Russia - Children's City Clinical Hospital of St. Vladimir

Subject 1. Hospital and transfusion service demographics

Ouestion 1 Russia, Moscow

Ouestion 2 No.

Ouestion 3

We prepare blood in our own facility.

**Question 4a** Yes.

**Question** 5

Our hospital is a paediatric specialty hospital. (a) 574 (24 reanimatology).

**Question 6** No.

Ouestion 7a

Yes.

Ouestion 7b Yes.

Ouestion 7c Yes.

**Question 7d** No.

**Question 7e** Yes.

Question f No.

Question 7q Yes.

Question 7h No.

**Question 7i** 

Yes.

Question 7j

Ouestion 7k

Yes.

Ouestion 71 Yes.

**Question 7m** Yes.

**Question** 7n Yes.

**Question 70** Yes.

Question 7p Yes.

Question 8 RBC (a) 96.9 l, 651 packages.

Ouestion 9

Yes, One packages from 70 to 100 ml.

**Question 10** Platelet. (a) 5.5 l, 64 packages.

## Subject 2. Transfusion indications for paediatric and neonatal patients

## Question 1b

Yes (There are restrictions on the time storages from the moment of donation: for children under 1 month not more than 10 days, for a replacement transfusion no more than 5 days, 100% individual selection taking into account 10 red cell antigens, recommended leucofiltration, irradiation, virus inactivation and selection of CMVnegative donors. Analysing haemoglobin level in children different age.)

**Question 2b** 

(a) In accordance with national principles.

**Question 3b** 

Yes.

(a) See application in attached table.

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Question 4b

Yes.

(a) No.

Question 5

Yes.

Ouestion 6b

Yes.

(a) See application in attached table.

**Question 7b** 

Yes.

(a) See application in attached table.

**Question 8b** 

Yes.

(a) See application in attached table.

Subject 3. Product manipulations for paediatric and neonatal patients

Question 1a

No.

Question 2b

Yes.

(a) Leucofiltration in the preparation of whole blood.

Question 3b

Yes.

(a) With transfusions of another blood group, an allergic anamnesis.

Question 4a

No.

Question 5a

No.

Question 6a

No.

Question 6b

Yes.

(a) Since 1999, we have been selecting erythrocytes with antigens – A, B, C, Cw, c, E, e, K, k.

Question 7

Yes. We make individual selection of red blood cells, thromboelastography, thrombodynamics.

Question 8

Yes. We do a division of a large dose into several paediatric packages.

Question 9

Yes.

Ouestion 10b

Yes.

(a) Erythrocytes can be used for children under 1 month for not more than 10 days of storage, for a transfusion no more than 5 days of storage.

Subject 4. Blood product dosing

**Question** 1

10-15 ml/kg.

Ouestion 2

5-10 ml/kg.

Question 3

10-15 ml/kg.

Formation of indications for blood transfusions

Erythrocyte-containing media

| Nosology                                  | Newborns and children up to 4 months  | Older age  |
|---|---|--|
| Routine transfusi                         | ions  |  |
| Anaemia<br>without signs<br>of bleeding   | Up to 14 days: Hb < 100 g/l, Ht < 30%, RBC < $3.0 \times 10^6$ 14–30 days: Hb < 96 g/l, Ht < 25%, RBC < $2.5 \times 10^6$ 1–4 months: Hb < 85 g/l, Ht < 25% | Hb < 70<br>Calculation of the<br>volume of erythrocyte-<br>containing media:<br>$V = \text{(Hb desired - Hb true)} \times 0.6 \times \text{m/2}$ |
| Anaemia with signs of respiratory failure | Up to 1 month: Hb < 120 g/ I, Ht < 40%, RBC < $3.9 \times 10^6$   | Hb < 80 g/l  |
| Anaemia and preparation for surgery       | Up to 1 month: Ht < 40%<br>1–4 months: Ht < 30%   | Hb < 100 g/l,<br>PLT < 100 $\times$ 10 $^{9}$  |
| Emergency transf                          | <sup>F</sup> usion  |  |
| Anaemia in critical conditions            | Of ventilation: $FiO_2 > 0.4$ $Hb < 110 \text{ g/l}, \text{ Ht} < 35\%$ $Auxiliary ventilation:$ $FiO_2 < 0.4$ $Hb < 100 \text{ g/l}, \text{ Ht} < 30\%$    | Taking into account transport O <sub>2</sub>   |

#### (Continued)

| Nosology                  | Newborns and children up to 4 months  | Older age   |
|---------------------------|---|---|
|                           | Self-breathing:<br>breathing rate 85/min 24 h<br>Hb < 80 g/l, Ht < 25%  |   |
| Intraoperative blood loss | In children under 4 months, the loss of more than 10% volume of circulating blood is replenished first of all by erythrocytes | Loss >15% volume of circulating blood (VCB): erythrocyte 1 dose + plasma 10% volume of circulating plasma loss >25% volume of circulating blood: erythrocyte 2 doses + plasma 50% volume of circulating plasma + platelet 5–6 doses loss >40% volume of circulating blood: erythrocyte - 30% VCB + plasma - 100% volume of circulating plasma + platelet 5–6 doses + albumin 20% 1 g/kg body mass |

## Thrombose concentrate

| Platelet level                        | Indications  |
|---------------------------------------|--|
| Any baseline<br>level of<br>platelets | Acute massive blood loss (absolute indications, included in the list of mandatory components for replenishment of blood loss). |
| >100 × 10 <sup>9</sup>                | Ratio of erythrocytes/plasma/platelets = 1/1/1 Transfusion not shown   |
| >100 X 10                             | It is possible to make a decision in favour of transfusion with functional insufficiency of platelets (thrombocytopathy)       |
| 50-                                   | Intraventricular haemorrhage III–IV st,  |
| $100 \times 10^{9}$                   | Continued bleeding,  |
|                                       | Surgical interventions of a large volume, including neurosurgical  |
| 20–49 × 10 <sup>9</sup>               | Minimally invasive interventions and operations (biopsy, puncture, epidural anaesthesia, replacement blood transfusion, etc.)  |
|                                       | The first 7 days after birth,  |
|                                       | Premature babies with extremely low body weight (less than 1000 g),  |
|                                       | Concomitant coagulopathy without marked bleeding,  |

#### (Continued)

| Platelet level        | Indications  |
|-----------------------|--|
| <20 × 10 <sup>9</sup> | Ongoing bleeding, skin-haemorrhagic syndrome<br>Septicaemia<br>Absolute indications, even without clinical<br>manifestations |

Contraindications to the use of thrombocyte concentrate:

- Immune thrombocytopaenia,
- Heparin-induced thrombocytopaenia,
- Thrombotic thrombocytopaenic purpura,
- Haemolytic-uraemic syndrome

## Freshly frozen plasma (FFP)

- · Multifactorial coagulation deficiency, confirmed laboratory and associated with the following:
  - o cutaneous haemorrhagic and haemorrhagic manifestations and ICE (Caution: DIC without bleeding is not an absolute indication for transfusion, nor is transfusion indicated for prophylactic purposes),
  - o liver disease (liver failure),
  - o haemolytic disease of newborns up to 20 ml/kg, with simultaneous administration of vitamin K,
  - o intraoperative haemodilution (dilution coagulopa-
- · Hereditary deficiency of coagulation factors, in the absence of a virus-safe preparation (including atypical haemolytic-uraemic syndrome),
- Thrombotic thrombocytopaenic purpura (plasma exchange),
- Elimination of the effect of warfarin, in case of bleeding,
- Operational haemorrhage of more than 15% volume of circulating blood

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Russia – Russian National Centre for Pediatric Hematology, Oncology and Immunology

Subject 1. Hospital and transfusion service demographics

Question 1

Hospital located in Moscow, Russian Federation.

Question 2

Yes, an academic hospital.

Question 3

No, all blood components provided by hospital-based blood bank.

Question 4a

Neonate - less than 28 days old.

Question 4b

Paediatric - less than 18 years old.

Question 5 Yes.

(a) Hospital has 220 inpatient beds.

Question 6

No, only paediatric patients.

Question 7a Yes.

Question 7b

No.

Question 7c

No.

**Question 7d** 

No.

Question 7e

No.

Question 7f

Yes.

Question 7g

No.

Question 7h

Yes.

**Question 7i** 

No.

Question 7j

Yes.

**Question 7k** 

No.

**Question 71** 

No.

**Question 7m** 

Yes.

Question 7n

No.

Question 70

No.

Question 7p

No.

Question 8a

6000 RBC transfusions per year.

Question 8b

(a) 30 transf.

(b) 278 transf.

(c) 5692 transf.

Question 9a

1700 units of FFP per year.

Question 9b

(a) 10 un.

(b) 104 un.

(c) 1586 un.

Question 10a

6300 units per year (1 unit =  $2 \times 10^{11}$  plt).

**Question 10b** 

(a) 29 un.

(b) 69 un.

(c) 6202 un.

## Subject 2. Transfusion indications for paediatric and neonatal patients

#### Ouestion 1b

Yes. Main differences are restrictions of the storages period for RBC: for children under 1 month not more than 10 days, for a replacement transfusion no more than 5 days, match for 10 red cell antigens, recommended leucofiltration.

#### **Question 2b**

Yes. In accordance with national guideline (see above).

#### **Question 3a**

No.

#### **Question 4b**

Yes.

(a) The indication is entered electronically.

#### Question 5

Yes.

#### Question 6b

Yes.

(a) For neonatal Hb <10 g/dl, Ht <29%; for paediatric Hb <7 g/dl, Ht <20%.

## **Question 7b**

Yes.

(a) For neonatal PLT <40, for paediatric PLT <10 or <40 before invasive procedure or <100 before neurosurgery.

## **Ouestion** 8b

Yes

(a) Multifactorial coagulation deficiency, confirmed laboratory.

## Subject 3. Product manipulations for paediatric and neonatal patients

## Ouestion 1b

Yes.

(a) All patients receive only irradiated RBC and platelets.

## **Question 2b**

Yes.

(a) Uniform leucodepletion of all transfused products.

### **Question 3b**

Yes.

(a) Patients with severe repeated PTR, patients with positive DAT, patients with unidentified haemolysis.

#### **Question 4a**

Nο

#### **Question 5b**

(a) Plasma, platelets, and whole blood are inactivated with MIRASOL PRT (Terumo).

#### Ouestion 6b

Yes.

(a) For all patients: k and Cw.

#### Ouestion 7

We are providing CMV-neg blood components upon request.

#### Ouestion 8

No.

#### **Question 9**

Yes.

#### **Question 10b**

(a) For neonatal patients' maximum storage period of RBC do not exceed 5 days.

## Subject 4. Blood product dosing

## Question 1

Millilitres/kg.

#### Question 2

Millilitres/kg.

## Question 3

Millilitres/kg.

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#### Eugene Zhiburt

## Russia – Pirogov National Medical Surgical Center

Subject 1. Hospital and transfusion service demographics

Question 1

Russia.

Question 2

Yes, my hospital is a teaching/academic hospital.

**Question 3** 

We receive the blood components from 3 blood centres.

Question 4a

Neonate - less than 30 days old.

**Question 4b** 

Paediatric - less than 18 years old.

**Question** 5

No, a paediatric specialty works only for outpatients.

**Question** 6

NA.

Question 7

NA.

Question 8

NA.

Question 9

NA.

Ouestion 10

NA.

Subject 2. Transfusion indications for paediatric and neonatal patients

Question 1

Yes.

(a) Reverse ABO typing starts since 4 months of age. Typing of C, c, E, e, Cw, K and k is mandatory. Newborn blood tube is labelled with mother surname and initials. Threshold for RBC transfusions: 85 g/l – before 1 year, 70 g/l – >1 year. For newborns: (i) RBCs should be stored <10 days; (ii) volume of transfused RBCs is 10–15 ml/kg; (iii) rate of RBCs

transfusion is 5 ml/kg/h; and (iv) transfused product should be warmed to 36–37°C. For intrauterine transfusion, RBC should be 0 RhD-negative with a period of storage no more than 5 days.

Ouestion 2

NA.

**Question 3b** 

Yes.

(a) NA.

**Question 4b** 

Yes.

(a) No.

Question 5

Yes.

Question 6b

Yes.

(a) NA.

Question 7b

Yes.

(a) NA.

**Question 8b** 

Yes.

(a) NA.

Subject 3. Product manipulations for paediatric and neonatal patients:

Question 1b

Yes.

(a) Allogeneic stem cell transplantation.

Question 2b

Yes.

(a) 100%.

Question 3a

No. Not necessary.

Question 4a

No.

Question 5b

Yes.

(a) MB-plasma and amotosalen platelets.

Question 6

NA.

Ouestion 7

NA.

**Ouestion 8** 

Jumbo plasma apheresis bags

**Question 9** 

No.

Question 10

No.

Subject 4. Blood product dosing

Question 1

For neonatal patients - 10-15 ml/kg.

Ouestion 2

 $50-70 \times 10^9$  platelets per 10 kg.

Question 3 15 ml/kg

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## The Netherlands

Subject 1. Hospital and transfusion service demographics

Ouestion 1

The Netherlands.

**Question 2** 

Academic.

Ouestion 3

We receive them from the national blood centre (Sanquin).

**Question 4** 

Neonate is less than 28 days, but for transfusion, we 'stretch' this till the age of 3 months.

Paediatric is indeed <18 years.

Question 5

It is a university paediatric hospital, a university adult hospital and a specialized adult oncology hospital. Total amount of paediatric beds = ?

Ouestion 6

See question 5.

Question 7a

Yes.

**Question 7b** 

Yes.

Ouestion 7c

Yes.

Ouestion 7d

Yes.

Ouestion 7e

Yes.

Question 7f

Yes.

Question 7g

Yes.

**Question 7h** 

Until 1st of June yes, from 1-6-2018 only shared care paediatric oncology. Paediatric (benign) haematology still is available. But paediatric stem cell transplantations occur in another academic hospital, only pre- and posttransplantation care occurs again in our hospital.

Ouestion 7i

Yes.

Question 7j

Yes, both. ECMO, LVAD also.

Ouestion 7k

Yes.

**Question 71** 

Yes.

**Question 7m** 

Yes.

**Question** 7n

Yes.

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#### Question 70

Yes

#### Question 7p

Yes and chronic ventilation care.

#### **Question 8a**

The total for both is 1468.

#### **Question 8b**

- (a) 31, younger than 1 month.
- (b) 50, 1-4 months.
- (c) 1387, older than 4 months.

#### **Question 9a**

The total for both is 551.

#### **Ouestion 9b**

- (a) 21, younger than 1 month.
- (b) 21, 1-4 months.
- (c) 509, older than 4 months.

#### Ouestion 10a

The total of both is 816.

## Question 10b

- (a) 26, younger than 1 month.
- (b) 19, 1-4 months.
- (c) 771, older than 4 months.

## Subject 2. Transfusion indications for paediatric and neonatal patients

#### Ouestion 1

Yes, there is (2011, update expected 2019/20), it is embedded in the general (adult) guideline with special chapters if needed. Extra attention is given to:

- Intrauterine blood transfusions
- HPA (e.g. 1a) negative platelets if indicated in FNAIT
- Exchange transfusion (e.g. with hyperbilirubinemia)
- Specific blood transfusions (e.g. apheresis for neonates) or regulations (e.g. irradiated if <32 weeks GA, HLA matched, and PvB19)</li>
- Laboratory tests (for neonates test mother and child)
- Thresholds differ for transfusions per age group

#### Question 2

No, we follow the national guideline. Rarely exceptions are made, for example FF-plasma extended till age of 1 year in case of large cardiothoracic surgery with ECMO.

#### Ouestion 3

Yes, there is. Indications

- Acute or chronic anaemia (threshold depends on clinical situation/ventilation etc), or during sickle cell crisis with complications, sporadic prophylactic in sickle cell disease.
- Platelets if (prophylactic in neonates <20, <10 in paediatric haemato-oncology, preoperative if regenerative thrombocytopaenia or functional platelet disorder), otherwise indications depend on stability of clinical state/correction of bleeding in thrombocytopaenic/-pathic patients
- Plasma, when bleeding tendency on ECMO, when F VI deficiency, when F XI deficiency, when plasmapheresis

#### Ouestion 4a

No, in general, but yes when a specific product is needed (PvB19/irradiated/HLA identical platelets, etc, this is only once at the beginning of the specification via paper- + electronic order. After that, it is automatically continued, also in electronic patient file, unless physician confirms that indication is no longer needed).

## Question 4b

Yes, see 4.a

#### **Question** 5

Only when it means that there are specific laboratory or product requirements. For example, sickle cell (extended blood group typing, use of ATG, SCTx or MDS  $\rightarrow$  irradiation, etc).

### Question 6

Yes.

#### Neonates.

- <7 days and born <32 weeks and/or <1500 g and with respiratory or circulatory support: Hb 8 mmol/ l = 12.8 g/dl
- <7 days and born <32 weeks and/or <1500 g and no respiratory or circulatory support: Hb 7 mmol/ l = 11.2 g/dl
- <7 days, born >32 weeks and/or >1500 gram and with respiratory or circulatory support: Hb 7 mmol/ l = 11.2 g/dl
- <7 days, born >32 weeks and/or >1500 g and no respiratory or circulatory support: Hb 6 mmol/ l = 9.6 g/dl
- >7 days, regardless of gestational age or birthweight and with respiratory or circulatory support: Hb
   5 mmol/l = 8 g/dl
- >7 days, regardless of gestational age or birthweight and no respiratory or circulatory support: Hb 4 mmol/l = 6.4 g/dl

#### Children.

- In need of ventilator or circulatory support: Hb 6 mmol/l = 9.6 g/dl
- In need of oxygen or otherwise critically ill but no (not yet) ventilator or circulatory support: Hb 5 mmol/l = 8 g/dl
- Stable clinical condition and anaemia secondary to regeneration (chemo/bone marrow failure: Hb 4.3 mmol/l = 7 g/dl
- Stable clinical condition, able to compensate for anaemia and acute or chronic anaemia (for other reasons than regeneration, e.g. iron deficiency): Hb 3.5-4 mmol/l = 5.6-6.4 g/dl
- Stable condition, but autoimmune haemolytic anaemia, able to compensate: Hb 3.5 mmol/l = 5.6 g/dl
- Presurgery and stable condition Hb 4 mmol/ l = 6.4 g/dl

## Question 7

Yes

#### Neonates.

- < or >7 days independent for gestational age or birthweight and with major bleeding (e.g. IVH/pulmonary haemorrhage): platelets >50, first 48 h try >100
- <7 days and born <32 weeks and/or <1500 g and with respiratory or circulatory support or with major bleeding (e.g. IVH/pulmonary haemorrhage): platelets >50
- < or >7 days independent for gestational age or birthweight but with no respiratory or circulatory support: platelets >20
- Presurgery (large surgery) independent for gestational age or birthweight: platelets >50
- In case of maternal IgG auto-antibodies (maternal ITP): platelets >20, preferably with IVIG, otherwise in first 5 days of life with platelet transfusion

#### Children.

- In need of major bleedings (e.g. intracranial and trauma): platelet >100
- In need of ventilator or circulatory support: platelets
- In need of ECMO or LVAD and bleeding: platelets
- In need of ECMO or LVAD and not bleeding: platelets >50
- In need of regenerative thrombocytopaenia and with bleeding and/or major surgery and/or: platelets >50

- In need of regenerative thrombocytopaenia and no bleeding nor major surgery nor anticoagulative therapy: platelets >10 if also on chemotherapy, >5 if secondary to bone marrow failure
- In case of surgery (preventive) or in case of bleeding in patients with thrombocytopathy: 20 ml/kg, with maximum of 1 unit
- Stable condition, but autoimmune thrombocytopaenia (ITP): none unless acute, major bleeding

#### **Question 8**

No. indications to diffuse.

Subject 3. Product manipulations for paediatric and neonatal patients

#### Ouestion 1

Yes.

- Neonates <32 weeks or <1500 g BW for 6 months
- Intrauterine transfusions, for 6 months postnatally
- Autologous stem cell transplantation
- Allogenic stem cell transplantation
- Use of ATG/Fludarabin/Campath
- Severe combined immune deficiency or selective severe T-cell defects elevating the risk on TA-GvH

#### Ouestion 2

Yes, all blood component transfusions are LD; this is the standard of the Dutch Blood Supply Company Sanguin

#### **Question 3**

Yes, rarely used, but possible indications are severe allergic reactions, pneumococcal-mediated HUS (T-antibody involvement) and only in PNH when routine blood products clearly lead to aggravation of haemolysis, not routinely.

### Ouestion 4

No, we do not, it will take a minimum of 30-45 min before it is thawed.

## Ouestion 5

No.

#### **Question 6**

Yes.

All girls and patients with MDS and auto-antibodies against red blood cells: also cEK

All haemoglobinopathies: cEK, Fy a/b, Jk a/b, MNSs.

## **Question 7**

Yes, but this is not specific for our hospital laboratory, it is arranged but the Dutch blood Supply Company Sanquin: routinely in case of intrauterine blood transfusion and on indication for individual needs (e.g. critically sick child with SCID pretransplantations, still CMV negative).

## Subject 4. Blood product dosing

## Question 1

10-15 ml/kg, maximum 2 units.

#### Question 2

15-20 ml/kg, maximum 1 unit.

#### **Question 3**

10–15 ml/kg, amount of units depend, usually maximum one, but for the children on ECMO, we sometimes provide continuously for 24 h ( $4 \times 6$  h).

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## **Brazil**

Subject 1. Hospital and transfusion service demographics

## Question 1

We are a tertiary care facility located in São Paulo city, SP, Brazil.

#### Question 2

We are a teaching/academic hospital with a Nursing college, a Medical College, a Research Institute, Medical Internship programmes and a Graduate Program in Health Sciences.

### **Question 3**

All the blood components are collected, tested and prepared at our own facility.

#### Ouestion 4

The following age thresholds are used to define our paediatric population:

- (a) Neonate patient: less than 28 days old.
- (b) Paediatric patient: 29 days old to less than 18 years old.

#### Ouestion 6

We are a 650-bed hospital for both adult and paediatric patients.

A total of 106 inpatient beds are available for paediatric patients with the following distribution:

- (a) Paediatric Ward: 20 beds;
- (b) Paediatric Intensive Care Unit: 18 beds;
- (c) Neonatal Ward: 46 beds;
- (d) Neonatal Intensive Care Unit: 22 beds.

Question 7a

Yes.

**Question 7b** 

Yes.

Question 7c

Yes.

Question 7d

Yes.

Question 7e

Yes.

Question 7f

Yes.

Question 7g

Yes.

Question 7h

Yes.

Question 7i

Yes.

Question 7j

Yes.

 $Question \ 7k$ 

Yes.

Question 71

Yes.

Ouestion 7m

Yes.

Ouestion 7n

Yes.

Ouestion 70

Yes.

Question 7p

Additionally, we provide specialties for paediatric and neonatal patients for Orthopedics and Fetal Medicine.

#### **Question 8a**

We define paediatric transfusion dose according to a patient's body weight.

Seven hundred and eight RBCs transfusions are done per year in both paediatric and neonatal patients at our facility.

#### **Ouestion 8b**

The transfused RBCs distribution according to patient's age is described below:

- (a) Younger than 1 month: 114 transfusions
- (b) 1-4 months: 98 transfusions
- (c) Older than 4 months: 496 transfusions (4 months to 1 year old: 64 transfusions; >1 year old: 432 transfusions)

## **Question** 9a

Eighty-six FFP transfusions are done per year in both paediatric and neonatal patients.

#### **Question 9b**

The transfused FFP distribution according to patient's age is described below:

- (a) Younger than 1 month: 28 transfusions
- (b) 1-4 months: 17 transfusions
- (c) Older than 4 months: 41 transfusions (4 months to 1 year old: 10 transfusions; >1 year old: 31 transfusions)

#### Ouestion 10a

Five hundred and four platelet transfusions are done per year in both paediatric and neonatal patients.

## **Question 10b**

The transfused platelet distribution according to patient's age is described below:

- (a) Younger than 1 month: 58 transfusions
- (b) 1-4 months: 39 transfusions
- (c) Older than 4 months: 407 transfusions (4 months to 1 year old: 32 transfusions; >1 year old: 375 transfusions)

## Subject 2. Transfusion indications for paediatric and neonatal patients

#### Ouestion 1

Brazil does not have specific guidelines for neonatal and paediatric populations regarding transfusion thresholds. However, we do have specific recommendations for irradiation and leukorredution of red blood cell units for the paediatric population, as mentioned in Subject 3, questions 1 and 2.

#### Ouestions 2 and 3

Our hospital has a list of indications for RBC, plasma and platelet transfusion [1-4], as follows:

- (a) RBC:
  - (i) Infants younger than 4 months:

Haematocrit <20% with low reticulocyte count and symptomatic anaemia (tachycardia, tachypnea, poor feeding)

Haematocrit <30% and either of the following: Tachypnea (respiratory rate >80 beats/min), tachycardia (heart rate >180 beats/min) for at least 24 h Significant bradycardia or apnoea

Oxygen support by nasal cannula

Continuous positive airway pressure support or mandatory ventilation on mechanical ventilation with mean airway pressure under 6 cm of water

Low weight gain (<10 g/day observed over 4 days, provided that there is adequate calories intake) On <35% oxygen hood

Haematocrit <35% and either of the following Continuous positive airway pressure support or mandatory ventilation on mechanical ventilation with mean airway pressure >6 cm of water On >35% oxygen hood

Haematocrit <45% and either of the following: Congenital cyanotic heart disease

Extracorporeal membrane oxygenation support

(ii) Paediatric patients:

Haemoglobin<7 g/dl (Haematocrit <21%) in: Stable, noncyanotic patients. Unstable children will be transfused at their physician's discretion Active bleeding with evidence of inadequate oxygen tissue delivery

Haematocrit <24% and either of the following: Symptomatic anaemia

Under chemotherapy or radiotherapy

Acute blood loss nonresponsive to volume replace-

Haematocrit <40% and either of the following: Severe pulmonary disease

Extracorporeal membrane oxygenation support

#### (b) FFP:

(i) Neonates and children:

Clinically significant bleeding or prior to invasive procedures with a significant bleeding risk with abnormal coagulation profile defined by APTT or PT above the normal gestational and postnatal age-related reference range.

## (c) Cryoprecipitate:

(i) Neonates and children:

Fibrinogen <1 g/l for surgery with significant bleeding risk or at critical sites;

Dysfibrinogenaemia with active bleeding or undergoing invasive procedures.

Factor XIII deficiency with active bleeding or while undergoing invasive procedures in the absence of factor XIII concentrate

Patients on ECMO with fibrinogen levels < 0.25 g/l

#### (d) Platelets:

(i) Prophylactic transfusion:

Platelet count  $<10-20 \times 10^9/l$  in stable paediatric patients

Platelet count <30  $\times$  10<sup>9</sup>/l in stable neonates Platelet count <50  $\times$  10<sup>9</sup>/l before invasive procedure Platelet count <100  $\times$  10<sup>9</sup>/l before neurosurgery

(ii) Therapeutic transfusion:

Platelet count  $<50 \times 10^9/l$  and active bleeding in stable neonates

Platelet count  $<100\times10^9/l$  in neonate with disseminated intravascular coagulation or unstable premature neonates

Platelet dysfunction

Patients on ECMO support with active bleeding or platelet count  $<50\times10^9/l$  or  $<100\times10^9/l$  if intracranial bleeding

#### Ouestion 4

It is not mandatory that the physician ordering the transfusion provides the indication by the time the blood products are ordered. However, transfusion staff does perform a critical analysis before preparing blood components to check compliance with our guidelines. If the transfusion is considered not appropriate, a Blood Bank physician will contact the physician who prescribed the blood component so that the latter can provide the reasons why the transfusion was ordered.

#### **Question** 5

It is mandatory that the person ordering transfusion provides patient's diagnosis when blood products are prescribed.

#### **Question 6**

Our hospital has thresholds for RBC transfusion, as described in question 3 above.

#### Question 7

Our hospital has thresholds for platelet transfusion, as described in question 3 above.

#### **Ouestion 8**

Our hospital has thresholds for plasma transfusion, as described in question 3 above.

Subject 3 Product manipulations for paediatric and neonatal patients

#### **Question 1**

Our transfusion service provides irradiated blood components which are recommended in the following conditions:

- (a) Low birthweight newborn (<1200 g)
- (b) Premature newborn (<28 weeks of gestation)
- (c) Neonate and paediatric patients with severe congenital immunodeficiencies
- (d) Transfusion of blood components from relatives and HLA compatible donors
- (e) Intrauterine transfusion
- (f) Massive transfusion. RBCs must be transfused up to 24 h postirradiation

#### Question 2

Our transfusion service provides leucodepleted blood components which are recommended in the following conditions:

- (a) HLA alloimmunization prevention
- (b) Febrile nonhaemolytic transfusion reaction prevention
- (c) Chronic transfusion regimen (i.e. haemoglobinopathies and myelodysplastic syndrome)
- (d) Prestorage leucodepleted blood components are accepted as surrogate for CMV-negative products in our country. Therefore, transfusion of CMV-negative or prestorage leucodepleted blood components is recommended in the following conditions:
  - (i) Low birthweight newborn (<1200 g)
  - (ii) Neonate whose mother is CMV negative or unknown CMV status
  - (iii) Intrauterine transfusion
  - (iv) Bone marrow transplant recipient with CMV-negative donor
  - (v) Solid organ transplant recipient with CMV-negative donor

#### **Question 3**

Our transfusion service provides washed RBCs which are recommended for patients who have had severe allergic or anaphylactic reactions in previous transfusion.

#### Ouestion 4

Our transfusion service does not maintain a stock of thawed plasma for immediate use.

#### Question 5

Our transfusion service does not provide pathogen-inactivated blood products.

#### Ouestion 6

Our transfusion service provides prophylactically matched RBC units for Rh, Kell, Duffy, Kidd and MNS antigen systems for chronically transfused patients with hereditary haemoglobinopathies and myelodysplastic syndrome.

#### **Question 7**

We do not provide other specific tests for paediatric patients.

#### **Ouestion 8**

Our transfusion service does have a procedure to minimize the number of donors the patients are exposed to. Once a RBC unit is selected to a neonate or paediatric patient, we try to transfuse aliquots of the same unit whenever feasible.

## Ouestion 9

Our transfusion service uses small-volume test-tubes (i.e. 1.2 ml) to draw blood samples from neonates and paediatric patients.

## **Question 10**

RBC units for massive transfusion, RBC exchange in neonates or intrauterine transfusion must be transfused up to 5 days after collection.

## Subject 4. Blood product dosing

#### Ouestion 1

We usually prepare 10-15 ml/kg body weight (b.w) of RBC for paediatric/neonatal patients.

## Ouestion 2

We usually prepare 10-15 ml/kg b.w. of platelets for paediatric/neonatal patients.

## **Question 3**

We usually prepare 10-15 ml/kg b.w. of plasma for paediatric/neonatal patients.

## References

- 1 New HV, Berryman J, Bolton-Maggs PH, et al.: Guidelines on transfusion for fetuses, neonates and older children. Br J Hematol 2016: 175:784-828
- 2 AABB. Technical Manual, 19th edn. Bethesda, MD: AABB Press, 2017
- 3 Parker RI: Transfusion in critically ill children: indications, risks and challenges. Crit Care Med 2014; 42:675-690
- 4 Yuan S, Tsukahara E, De La Cruz K, et al.: How we provide transfusion support for neonatal and pediatric patients on extracorporeal membrane oxygenation. Transfusion 2013; 53:1157-1165

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## Spain

Subject 1. Hospital and transfusión service demographics

Ouestion 1 Spain

#### Ouestion 2

Yes, it is a university hospital.

## **Question 3**

We receive the blood components from the Regional Blood Center (Centro de Transfusion Comunidad de Madrid, CTCM). They provide the blood components, and we prepare them for transfusion.

Question 4a

Neonate: less than 30 days old.

Question 4b

Paediatric: less than 18 years old.

Question 5

Yes, it is a monographic paediatric hospital constituted

by 180 inpatient beds.

**Question** 6

No, it is only for paediatric patients.

**Question 7a** 

There are Allergy and Rheumatology services, but not

Immunology.

Question 7b

Yes

**Question 7c** 

Yes

Ouestion 7d

Yes

**Question 7e** 

Yes

Question 7f

No

Question 7g

Yes

Question 7h

Yes

Question 7i

No

Question 7j

Yes

Question 7k

Yes

Question 71

Yes

Question 7m

General surgery, neurosurgery, traumatology, plastic sur-

gery, urology.

Question 7n

Yes

Ouestion 70

Yes

Question 7p

Yes

*Ouestion 8a* 

1800-2000 RBCs.

**Question 8b** 

All of them have been older than 4 months. There is not a neonatal department, so neonatal patients are not admitted.

Question 9a

150-250 plasma units.

**Question** 9b

All of them have been older than 4 months (no neonatal

department).

Question 10a

1300-1500 platelet units.

**Question 10b** 

All of them have been older than 4 months (no neonatal

department).

Subject 2. Transfusion indications for paediatric

and neonatal patients

**Question 1b** 

Yes. Guide for the transfusion of blood components and plasma derivatives, 5th edition, SETS (Spanish Society of

Blood Transfusion and Cellular Therapy).

(a) Criteria vary according to age: preterm infants, under

4 months, older than 4 months.

Question 2b

Yes; elaborated by transfusion Committee based on the guide for the transfusion of blood components and plasma derivatives of the Spanish Society of Blood Transfusion and Cellular Therapy: Blood Derivatives Transfusion Guide (Code: CTR-DC-002).

(a) They differ according to the weight and age.

#### Question 3b

Yes. Elaborated by transfusión Committee based on the guide for the transfusion of blood components and plasma derivatives of the Spanish Society of Blood Transfusion and Cellular Therapy: Blood Derivatives Transfusion Guide (Code: CTR-DC-002). See the indications on question 6, 7 and 8.

#### Question 4b

Yes, they must provide the indication for the transfusion. (a) The indication is entered electronically.

#### **Question** 5

Yes, they must provide the patient's diagnosis at the time of ordering blood products.

## Question 6b

Yes. Elaborated by transfusion Committee based on the guide for the transfusion of blood components and plasma derivatives of the Spanish Society of Blood Transfusion and Cellular Therapy: Blood Derivatives Transfusion Guide (Code: CTR-DC-002) [1].

(a) Thresholds for paediatrics:

Paediatrics <4 months:

- Hb <7 g/dl with low reticulocytes and anaemia symptoms.
- Hb <9 g/dl if:
  - o Oxygen requirements with CPAP or mechanic ventilation with FiO2 30-35% or high flow rate with more than 1 l/kg and FiO<sub>2</sub> 30-35% or oxygen glasses with >2 l/min and saturation ≤92%.
  - o Signs of apnoea, bradycardia, tachycardia or tachypnoea and low weight gain (≤10 g/day for 4 days receiving ≥100 kcal/kg/day).
- Hb <9 g/dl if:
  - o Oxygen requirements with CPAP or mechanic ventilation with  $FiO_2 \ge 35\%$  or high flow rate with more than 1 1/kg and  $FiO_2 \ge 35\%$  and saturation  $\le 92\%$ .
  - o Preoperative anaemia
- Hb <15 g/dl with: congenital cyanotic cardiopathy or oxygenation with extracorporeal membrane.

## Paediatrics >4 months:

- Hb <7 g/dl with chronic anaemia and/or not responding to specific and symptomatic treatment.
- Hb 7–10 g/dl according to the clinical situation:

- Acute loss of >25% of blood volume.
- o Preoperative Hb <8 g/dl or losses above 15% of the blood volume.
- o Hb <8 g/dl and treatment with chemotherapy and/ or radiotherapy.
- Hb <13 g/dl and severe pulmonary disease, cyanotic cardiopathy or oxygenation with extracorporeal membrane with descent of  $O_2$  saturation.

#### Ouestion 7

Yes. Elaborated by transfusión Committee based on the guide for the transfusion of blood components and plasma derivatives of the Spanish Society of Blood Transfusion and Cellular Therapy: Blood Derivatives Transfusion Guide (Code: CTR-DC-002) [1].

(a) Thresholds for paediatrics and neonates:

They should be transfused below the following figures:

- Platelets  $<20 \times 10^9/l$  and asymptomatic
- Platelets between 30 and  $50 \times 10^9$ /l in unstable patient:
  - Premature in their first week of life and/or <1 kg.
  - o Premature with bleeding Grade III or IV
  - Newborns with signs of bleeding
  - o Need for invasive procedures or administration of drugs that alter platelets
  - $\circ$  Platelets <50  $\times$  10<sup>9</sup>/l and
- Need to perform an invasive procedure or exchange transfusion
- Platelets  $<100 \times 10^9/l$  before a major surgery

## **Ouestion 8**

Yes. Elaborated by transfusión Committee based on the guide for the transfusion of blood components and plasma derivatives of the Spanish Society of Blood Transfusion and Cellular Therapy: Blood Derivatives Transfusion Guide (Code: CTR-DC-002) [1].

- (a) Thresholds for paediatrics and neonates:
  - (i) Prior to invasive procedures or surgery if INR (international normalized ratio) >1.5 or 2 times above normal value.
  - (ii) Replacement treatment for disseminated intravascular coagulation or massive transfusion.
  - (iii) Deficit of coagulation factors, with haemorrhage prior to invasive procedures or surgery, if the specific coagulation factor is not available for administration.
  - (iv) Thrombotic thrombocytopenic purpura.
  - (v) In the case of fulminant purpura of the newborn due to protein C and S deficiency, if specific concentrates of these factors are not available for administration.

- (vi) For reconstitution of the erythrocytes concentrate, when total blood is not available for the realization of exchange transfusion.
- (vii) If is necessary to revert the effect of the vitamin K-dependent anticoagulants, we recommend to use vitamin K in the first place. In situations with vital risk of bleeding or urgent invasive procedure, Complex Prothrombin concentrate will be administered, if is not available, fresh frozen plasma will be used.

## Subject 3. Product manipulation for paediatric and neonatal patients

#### **Question 1b**

Yes. Elaborated by transfusión Committee based on the guide for the transfusion of blood components and plasma derivatives of the Spanish Society of Blood Transfusion and Cellular Therapy: Blood Derivatives Transfusion Guide (Code: CTR-DC-002) [2].

Indications for irradiation

- Severe immunodeficiency syndromes.
- Receptors of Allogeneic Hematopoietic Stem Cell Transplantation (HSCT) from the beginning of the conditioning until the patient is receiving prophylaxis for graft-versus-host disease.
- Patients with Hodgkin lymphoma.
- Patients treated with purine analogs (fludarabine, cladribine and deoxycoformycin).
- Exchange transfusion in intrauterine transfusion receptors.
- Receivers of intrauterine transfusions (UTI) up to 6 months after the probable date of birth (40 weeks of gestation) or if the donation comes from a first or second family degree.
- Patients undergoing bone marrow or peripheral blood progenitor extraction for autologous reinfusion from 7 days before and during the collection.
- Patients undergoing transplantation of autologous haematopoietic progenitors (HSCT) since the beginning of the conditioning up to 3 months after transplantation (6 months if total body irradiation was used in conditioning).
- Patients treated with alemtuzumab (anti-CD52).
- Patients with aplastic anaemia treated with antithymocyte gamma globulin.

#### Ouestion 2b

Yes. All of them are leucodepleted according to national requirements

## Question 3b

Yes, provided by the regional blood centre.

(a) Ig A deficit.

#### Ouestion 3a

No, we do not maintain a stock of thawed plasma units.

#### Ouestion 4b

Yes; all plasma units are inactivated according to the CTCM standard procedure. The inactivation protocol varies annually according to a public assignment procedure, being more frequent inactivation with blue methylene in the last years. Also, a small proportion of platelet units are inactivated with Amotosalen.

#### **Question 5b**

In girls, we match the following antigens Rh and Kell, as referred in SOP SET-PT-035.

#### **Question** 6

Yes, we matched blood components for CMV in haematopoietic stem cell transplantation receptors.

We carry out studies such as direct Coombs test, detection of irregular antibodies by indirect Coombs test, identification of them by panels or extended phenotype studies; the rest of the extended or complementary studies are provided by Regional Blood Center.

#### **Question 7**

Yes, we aliquot the units and prioritize the use of apheresis platelets instead of pools.

#### **Ouestion 8**

No.

## Question 9a

No.

#### Subject 4. Blood product dosing

#### Question 1

Units or half unit according to the weight.

## Question 2

Five unit pools or apheresis products for patients over 30 kg, and half apheresis for those less than 30 kg.

## Question 3

10-20 ml/kg adjusted to plasma units volume (250 cc).

## References

1 New H, Berryman J, Bolton-Maggs P, et al.: Guidelines on transfusion for fetuses, neonates and older children. *British Journal of Haematology* 2016; 175:784–828 2 Treleaven J, Gennery A, Marsh J, et al.: Guidelines on the use of irradiated blood components. British Committee for Standards in Haematology blood transfusion task force. Br JHaematol 2011; 152:35-51

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## Japan

Subject 1. Hospital and transfusion service demographics

**Question** 1 Japan.

Question 2 Yes.

Question 3

All the allogenic blood products are from the Japanese Red Cross Blood Center, and autologous blood products are prepared in our hospital.

Question 4a

Less than 30 days old.

**Question 4b** 

Between 30 days old and 15 years.

**Question** 5

No.

Question 6

Yes. About 20 beds.

**Ouestion** 7a

Ouestion 7b Five physicians.

Question 7c One physician.

**Question 7d** Two physicians.

**Question 7e** No.

Question 7f No.

Question 7g Yes.

Ouestion 7h Seven physicians.

**Question 7i** No.

Question 7j Four physicians.

Question 7k Three physicians.

Question 71 One physician.

Question 7m Five physicians.

Question 7n Four physicians.

**Question 70** 

Yes (at the Dept. of Otolaryngology).

Question 7p

Yes (at the Dept. of Palliative Care).

#### **Question 8a**

1260·24 units (in Japan, 1 unit is defined as that obtained from 200 ml of whole blood).

#### Question 8b

- (a) 240.74 units.
- (b) 79.5 units.
- (c) 940 units.

#### Ouestion 9a

406 units (in Japan, 1 unit is defined as 120 ml of plasma).

#### **Question 9b**

- (a) 134 units.
- (b) 40 units.
- (c) 232 units.

#### Question 10a

8067 units (in Japan, 1 unit is defined as  $2 \times 10^{10}$  counts/bag)

#### Question 10b

- (a) 647 units.
- (b) 90 units.
- (c) 7330 units.

## Subject 2. Transfusion indications for paediatric and neonatal patients

## Question 1

Yes.

- (a) We have just revised the guideline for neonatal and paediatric (up to 4 months) transfusions. The threshold for the administration of each labile blood for neonates is somehow different from that for adults. The indications for neonatal and paediatric transfusion are as follows:
  - (i) RBC transfusion will be considered; (1) if the patients' Hb is equal to or lower than 7 g/dl when the patients are in a stable condition; (2) if the patients' Hb is equal to or lower than 11 g/dl when the patients are chronically dependent on oxygen administration; and (3) if the patients' Hb is equal to or lower than 12 g/dl when the patients are the neonates within 24 h after birth or neonates under intensive care treatment. RBC storage less than 2 weeks from donation is preferable, and 10-20 ml/kg of RBC will be given at a speed of 1-2 ml/kg/h if there are no signs of congestive heart failure in the patients. If there are signs of congestive heart failure, RBC transfusion will be discussed separately. Because transfusion of RBC using the needle thinner than 24 gauge with pressure pump

- may cause haemolysis, careful attentions need to be paid to this kind of haemolysis. RBC transfusion should be finished within 6 h after opening the bag. If the duration of transfusion exceeds 6 h, RBC should be aseptically divided into small bags and stored at  $2-6^{\circ}$ C until use.
- (ii) (1) Platelet concentrate transfusion should be considered if the platelet count is lower than 2- $3 \times 10^3/\mu l$  when the patients are stable and have no signs of bleeding. Higher platelet counts are recommended for the premature infants within a few days after birth. (2) Platelet concentrate administration will be considered when the platelet count is lower than  $3 \times 10^3$ ul if the patients are diagnosed as NAIT (neonatal alloimmune thrombocytopenia). (3) Platelet count should be maintained at more than  $5 \times 10^3/\mu l$  when the patients are infants with very low birthweight, with any signs of bleeding or undergoing invasive treatments. (4) Platelet count should be maintained at  $5-10 \times 10^3$ ul when the patients are developing DIC or undergoing major surgeries.
- (iii) FFP should be administered; (1) when there is elongation of PT or APTT despite an administration of vitamin K, and there is bleeding or is undergoing invasive treatments; (2) when RBC transfusion exceeds half of the amount of circulating blood; and (3) when the patients are diagnosed Upshaw-Schulman syndrome (congenital thrombotic thrombocytopenic purpura). FFP should be administered at 10-20 ml/ kg every 12-24 h (1) and (2). In the case of (3), FFP should be administered at 10 ml/kg every 2-3 weeks. FFP can be substituted by saline solution in the case of the partial exchange transfusion for neonatal polycythemia.

Question 2 No.

Question 3

Question 4
Yes.
(a) Yes.

#### **Question** 5

No. However, columns are available to enter the diagnoses of the patients, and physicians can input them on voluntary bases.

#### Question 6

Yes.

(a) For adult patients only.

#### Question 7

(a) For adult patients only.

#### Question 8

Yes.

(a) For adult patients only.

## Subject 3. Product manipulations for paediatric and neonatal patients

#### **Question** 1

Yes.

(a) All RBC and platelets are irradiated.

#### Question 2

Yes.

(a) All the labile blood products are leucodepleted and supplied by the Japanese Red Cross.

#### Question 3

Yes.

(a) Washed RBCs can be purchased from Japanese Red Cross, and we usually do not prepare washed RBC in our hospital. We have experience of preparing washed RBC only once for a patient with paroxysmal nocturnal haematuria.

#### **Question 4**

No.

## Question 5

No.

#### Ouestion 6

No.

## Question 7

No.

## **Question 8**

Yes.

(a) One unit RBC is provided split into up to 3 bags.

## **Question 9**

No.

#### Ouestion 10

No.

## Subject 4. Blood product dosing

#### Ouestion 1

0.5-2 units (in Japan, 1 unit is defined as that obtained from 200 ml of whole blood).

#### **Question 2**

5-20 units (in Japan, 1 unit is defined as 120 ml of plasma)

#### **Question 3**

1–3 units (in Japan, 1 unit is defined as  $2 \times 10^{10}$  counts/ bag)

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## Italy

## Subject 1. Hospital and transfusion service demographics

#### **Question** 1

Our hospital is located in North West Italy in the city of Torino.

#### Ouestion 2

It is a teaching academic hospital.

#### **Question 3**

The Blood Establishment, from which we receive blood components, is part of our facility.

#### **Question 4**

At our hospital, an age thresholds of less than 30 days old is used to define neonate while a thresholds of less than 18 years old define paediatric patients.

#### Question 5

No.

#### **Question** 6

Our hospital treats both adult and paediatric patients and has nearly 300 inpatient beds for paediatric and neonatal patients.

#### **Question 7a**

Yes.

#### Question 7b

Yes.

#### Ouestion 7c

Yes.

#### **Question 7d**

Yes.

#### **Question 7e**

Yes.

## Question 7f

Yes.

## Question 7q

Yes.

## Ouestion 7h

Yes.

#### **Question 7i**

Yes.

## Question 7j

Yes.

#### **Question 7k**

Yes.

## Question 71

Yes.

#### Question 7m

Yes.

#### **Question 7n**

Yes.

#### Question 70

Yes.

## Question 7p

Yes.

Additionally, our hospital provides solid organ transplants for paediatric and neonatal patients.

#### Question 8a

Around 3000 RBCs are transfused to paediatric and neonatal patients every year.

#### Ouestion 8b

- (a) Nearly 300 RBCs to neonates younger than 1 month old.
- (b) 180 RBCs to paediatric patients between 1 and 4 months old.
- (c) 2500 RBCs to paediatric patients between 4 months and 18 years old.

#### **Question** 9a

Around 1300 plasma units are transfused to paediatric and neonatal patients each year.

## **Question** 9b

- (a) 240 plasma units to neonates younger than 1 month
- (b) 90 plasma units to paediatric patients between 1 and 4 months old.
- (c) 1000 plasma units to paediatric patients between 4 months old and 18 years old.

#### Ouestion 10a

Around 2000 platelet concentrates are transfused to paediatric and neonatal patients each year.

## Ouestion 10b

- (a) 50 platelet concentrates to neonates younger than 1 month old.
- (b) 15 platelet concentrates to paediatric patients between 1 and 4 months old.
- (c) 1900 platelet concentrates to paediatric patients between 4 months old and 18 years old.

## Subject 2. Transfusion indications for paediatric and neonatal patients

#### Ouestion 1

In Italy, there is a national recommendation for Neonatal transfusion [1] (last revision in 2014), made by Italian Society of Transfusion Medicine (SIMTI), through a working group of medical experts, including transfusion medicine, foetal medicine, neonatology, paediatric intensive care and haematologist. The recommendation is shared by the Italian society of neonatology (SI). The main differences from adult guidelines concern:

- Blood component selection: from regular donors only, irradiated, then 'fresh' (before the end of Day 5 following donation) and with a 24-h shelf-life postirradiation for intrauterine transfusions (IUT) and exchange blood transfusion (EBT). In order to reduce donor exposure, small-volume splits of single donor (pedi-packs) are used for neonatal transfusions; blood components have to be prepared following the standards of The Guide to the Preparation, use and quality assurance of blood components, 19th Edition, European Directorate for the Quality of Medicines & HealthCare of the Council of Europe [2]
- Small-volume transfusions (15 ml/kg in nonbleeding neonates):
- Suggestions of restrictive RBCs transfusion thresholds in stable and nonbleeding patient. However, the final decision for transfusion is made by clinicians based on their clinical judgement.

### **Question 2**

Our hospital has local policy that is available for clinicians on local network. In addition to the indications suggested at national level, our policy recommends the following:

- Blood component selection: from regular donors who have given at least two donations per year within the previous 2 years;
- Small-volume splits of single donor (pedi-packs, mean volume 70-80 ml) are stored for the same patient at least until T&S expiry date (72 h);
- Suggestions of restrictive RBCs transfusion thresholds in stable and nonbleeding patient, avoiding for very low birthweight (VLBW) and extremely low birthweight (EVLBW) haemoglobin levels below the lower limits tested in the Cochrane review (Table 3) [3].

### **Question 3**

Our hospital has a list of indications for ordering RBC plasma and platelets; for paediatric transfusion, indication refers to BSH Guideline [4]. These indications include IUT, EBT, haemolytic disease of newborn (HDN), preterm anaemia and neonatal cardiac surgery.

#### Ouestion 4

In our hospital it is mandatory, for the person ordering the transfusion, to provide the indication for the transfusion at the time that the blood products are ordered, at the moment the indication is chosen from a list present in the emocomponent order paper. By the end of 2019, an electronic order will be available.

#### Question 5

It is also mandatory for the person ordering the transfusion to provide the patient's diagnosis when blood products are ordered.

#### Ouestion 6b

Our hospital has guidelines with threshold for RBC, plasma and platelets transfusion for adult patients as well as for paediatric and neonatal pts.

(a) See Table 1 for RBCs thresholds in stable nonbleeding neonatal patients. Indications do not exclude clinical evaluation by clinicians who eventually decide for transfusion.

Table 1. Haemoglobin threshold levels (g/l) (Haematocrit %) triggering RBC transfusion (modified by Whyte et al. [3])

| Age in days    | Blood sampling | Respiratory support   | No respiratory support |
|----------------|----------------|-----------------------|------------------------|
| 1–7            | Capillary      | <u>&lt;</u> 115 (35%) | <u>&lt;</u> 100 (30%)  |
| 8-14           | Capillary      | <100 (30%)            | <85 (25%)              |
| <u>&gt;</u> 15 | Capillary      | <u>&lt;</u> 85 (25%)  | <u>&lt;</u> 75 (23%)   |

## Question 7b

Thresholds for platelet transfusion in neonates are also suggested and are shown in Table 2. Indications do not exclude clinical evaluation by clinicians who eventually decide for transfusion.

Table 2. Platelets threshold levels (109/l) triggering platelet transfusion [1, 4]

| Platelets count | Indication for transfusion   |  |
|-----------------|--|--|
| <30             | Always indication for transfusion  |  |
| 30–49           | Neonates with bleeding, current coagulopathy,<br>before surgery, previous intracranial haemorrhage |  |
| 50-99           | Major bleeding or requiring major surgery  |  |
| >100            | Usually no indication for transfusion  |  |

#### **Ouestion 8**

Our hospital does not have thresholds for plasma transfusion in paediatric and neonatal patients but suggests plasma use in neonates with clinically significant bleeding (including massive blood loss), prior to invasive procedures with a risk of significant bleeding, who have an abnormal coagulation profile, defined as a PT or APTT significantly above the normal gestational and postnatal age-related reference range and neonates who have congenital bleeding disorders, with no specific factor concentrate available. Our policy discourages the routine use of plasma to correct abnormalities of the coagulation screen alone in nonbleeding neonates and for prevention of intraventricular haemorrhage.

## Subject 3. Product manipulations for paediatric and neonatal patients

#### Ouestion 1

Our transfusion department offers irradiated products. Indications for irradiation are as follows: prevention of graft-versus-host disease and particularly in IUT, EBT and for all neonatal VLBW and ELBW patients; in paediatric patients with haematological malignancies and in liver transplantation patients younger than 4 years old.

#### **Question 2**

Since 2017, Italy has introduced universal leucodepletion so RBCs and platelets are all leucodepleted. Even before our department has always issued leucodepleted RBCs and platelets for IUT, EBT, neonatal transfusions and paediatric oncologic and haematologic patients.

#### **Question 3**

We also have the chance to offer washed RBCs. Indications are as follows: EBT and all patients with repeated serious allergic or anaphylactic or anaphylactoid reaction or patients considered to be at higher risk of an anaphylactic or anaphylactoid reaction due to selective severe IgA deficiency with or without anti-IgA antibodies. Washed RBCs are also delivered in operating room during liver transplant for patients younger than 8 months old.

#### Question 4

We do not maintain a stock of thawed plasma units for immediate issue.

#### **Question** 5

In our transfusion centre, we can offer pathogen-inactivated plasma, cryoprecipitate and cryoprecipitate-depleted plasma. Methods for inactivation are as follows: solvent/detergent and riboflavin/UV.

#### Question 6

Our transfusion department prophylactically matches, for all neonates and paediatric patients, RBCs C, c, E, e and Kell antigens other than ABO and RhD.

#### Ouestion 7

Usually, we do not provide other specific tests for neonatal/paediatric patients, unless some peculiar condition as CMV-negative stem cell transplantation candidate where CMV-negative blood units are issued.

#### **Question 8**

In order to minimize number of donors, the patients are exposed to, for neonatal transfusion, we use small-volume splits of single donor (pedi-packs, mean volume 70–80 ml) and store them for the same patient at least until T&tS expiry date (72 h to 3 days). During liver transplant, RBC units divided into two paediatric units are delivered coupled to be transfused one after the other in case of needing before transfusion of any other RBC.

#### Ouestion 9

To minimize the volume of the samples drawn for laboratory testing, only small-volume test-tubes for collecting samples are used. A policy to reduce when possible frequency of sampling is applied in all departments.

## Question 10

Irradiated RBCs with no more 5 days shelf-life postirradiation are used for all neonatal patients and paediatric patients younger than 6 months old. Five-day storage age RBCs are issued during liver transplant, for IUT, EBT and neonatal cardiac surgery. RBCs with a 24-h shelf-life postirradiation are used for IUT, EBT, neonatal cardiac surgery and liver transplant younger than 4 years old.

## Subject 4. Blood product dosing

#### Question 1

For stable paediatric and/or neonatal patients, 5–10 ml/kg of RBCs is usually prescribed.

#### **Question 2**

For stable paediatric and/or neonatal patients, 10 ml/kg of platelet concentrate is usually prescribed.

## Question 3

For stable paediatric and/or neonatal patients, 15–20 ml/kg of plasma [1, 4] is usually prescribed.

## References

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- 2 The Guide to the Preparation, Use and Quality Assurance of Blood Components, 19th edn. www.edqm.eu: European Directorate for the Quality of Medicines & HealthCare of the Council of Europe (EDQM).
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- 4 British Committee for Standards in Haematology. British Committee for Standards in Haematology Clinical Guideline: Transfusion for Fetuses, Neonates and Older Children. London: BCSH, 2016.

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## India

Subject 1. Hospital and transfusion service demographics

#### Question 1

Our hospital is located in India.

## Ouestion 2

Ours is a major academic teaching hospital offering the widest possible selection of services. It is a large public tertiary healthcare institute and a referral centre for five provinces of north India. It conducts postgraduate and doctoral courses in all the major medical and surgical specialties and superspecialities. It also conducts graduate and postgraduate courses in nursing and medical laboratory technology. About 1.2-1.3 million patients visit the outpatient services (OPD) of our institute every year. It has inpatient bed strength of 2200 with an annual inpatient admission numbers ranging from 80 000 to 90 000.

#### Ouestion 3

The institute has a Department of Transfusion Medicine, a hospital-based academic blood centre with postgraduate and doctoral programmes in the speciality. The Department collects around 60 000 units of blood, majority (>90%) from voluntary source, prepares blood components and performs mandatory screening for infectious markers (HIV-18t2, HBV, HCV, syphilis and malaria) and compatibility testing. The requisite blood components are issued to both adult and paediatric patients in emergency as well as for routine transfusions. The Dept. also has an active apheresis unit performing plateletpheresis, therapeutic plasma exchange and stem cell harvesting and cryopreservation.

#### Question 4a

A less than 30-day-old child is defined as a neonate in our Institute.

#### Question 4b

A less than 13 years old is defined as a paediatric patient in our Institute.

## Ouestion 5

Our Institute provides treatment for both paediatric and adult patients.

#### **Question 6**

Our Institute provides treatment for both paediatric and adult patients. In our institute, we have an Advanced Pediatric Care Centre with bed strength of 283 exclusively for paediatric patients in the same campus.

Ouestion 7a

Yes.

Ouestion 7b

Yes.

Ouestion 7c

Yes.

**Question 7d** 

Yes.

**Question 7e** 

Yes

Question 7f

Yes.

Question 7g

Yes.

Question 7h

Yes.

Question 7i

Yes.

**Question 7**j

Yes.

**Question 7k** 

Yes.

Question 71

Yes.

Ouestion 7m

Yes.

Ouestion 7n

Yes.

**Question 70** 

Yes.

Question 7p

Yes

#### **Question 8a**

On an average, 10 000–12 000 packed red blood cells (PRBC) units are transfused annually to paediatric and neonatal patients in our institute and in the year 2017–2018 (1 July 2017 to 30 June 2018), 12 144 units of PRBC were transfused.

## Question 8b

- (a) A total of 2754 PRBC units were transfused to the patients younger than 1 month, in the year 2017–2018 (1 July 2017 to 30 June 2018).
- (b) A total of 1140 PRBC units were transfused to patients of 1–4 months, in the year 2017–2018 (1 July 2017 to 30 June 2018).
- (c) A total of 8250 PRBC units were transfused to patients older than 4 months, in the year 2017–2018 (1 July 2017 to 30 June 2018).

#### Ouestion 9a

On an average, 3000–3500 fresh-frozen plasma (FFP) units and 100–150 cryoprecipitate units are transfused annually to paediatric and neonatal patients in our institute and in the year 2017–2018 (1 July 2017 to 30 June 2018), and 3322 units of FFP and 91 units of cryoprecipitate were transfused.

#### **Question 9b**

- (a) A total of 715 FFP units and 2 cryoprecipitate units were transfused to patients of younger than 1 month, in the year 2017–2018 (1 July 2017 to 30 June 2018).
- (b) A total of 212 FFP units and 2 cryoprecipitate units were transfused to patients of 1–4 months, in the year 2017–2018 (1 July 2017 to 30 June 2018).
- (c) A total of 2395 FFP units and 87 cryoprecipitate units were transfused to patients older than 4 months, in the year 2017–2018 (1 July 2017 to 30 June 2018).

#### **Question 10a**

On an average, 8000–8500 whole blood-derived random donor platelets (RDP) and 200–250 single-donor apheresis platelet (SDAP) units (split into aliquots) are transfused annually to paediatric and neonatal patients in our institute and in the year 2017–2018 (1 July 2017 to 30 June 2018), and 8055 RDPs and 227 SDAP were transfused.

## Question 10b

- (a) A total of 2386 RDPs and 16 SDAPs (split into aliquots) were transfused to patients of younger than 1 month, in the year 2017–2018 (1 July 2017 to 30 June 2018).
- (b) A total of 320 RDPs and 8 SDAPs (split into aliquots units) were transfused to patients of 1–4 months, in the year 2017–2018 (1 July 2017 to 30 June 2018).
- (c) A total of 5349 RDPs and 203 SDAPs (split into aliquots) units were transfused to patients older than 4 months, in the year 2017–2018 (1 July 2017 to 30 June 2018).

## Subject 2. Transfusion indications for paediatric and neonatal patients

#### Question 1

There are evidence-based clinical practice guidelines developed by the National Neonatology Forum of India, October 2010, and are available to guide neonatal transfusion practice [1]. For paediatric transfusions, standard textbook and current international published guidelines are followed.

However, there are no national published guidelines for adult transfusion for comparison.

#### Ouestion 2

Our institute has local protocols for paediatric and neonatal transfusions based on the current national and international guidelines. However, there are no published local guidelines for adult transfusion for comparison.

#### Question 3b

The neonatologists follow the National Neonatology Forum of India, October 2010, guidelines for ordering blood components for transfusion. For the paediatric age group; standard national and international guidelines are followed. On review of the blood requisitions for the last one year (1 July 2017 to 30 June 2018), the indications for blood component ordering are listed below:

Indications for RBC transfusion - (percentage of total RBC transfusions)

| Neonates    | Exchange transfusion for neonatal sepsis (50%) and                               |
|-------------|--|
| Medical     | haemolytic disease of newborn (20%)  |
| (60%)       | Anaemia developing due to various other neonatal                                 |
|             | problems such as anaemia of prematurity and                                      |
|             | haemorrhagic disease of newborn (30%)  |
| Neonates    | Congenital malformations such as anorectal                                       |
| Surgical    | malformations, posterior urethral valves and cardiac                             |
| (40%)       | anomalies such as Tetralogy of Fallot  |
| Paediatrics | Haematological malignancies (36%).   |
| Medical     | Genetic disorders (26%) – thalassaemia.  |
| (69%):      | Anaemia due to   |
|             | (a) Acute infections (20%): pneumonia, meningitis, pyrexi                        |
|             | of unknown origin with anaemia, acute respiratory distress syndrome              |
|             | (b) Chronic infections (8%): tubercular meningitis.                              |
|             | (c) Other indications (10%): haemolytic-uraemic syndromo                         |
|             | chronic kidney disease and autoimmune haemolytic anaemias                        |
| Paediatrics | Abdominal mass, subacute intestinal obstruction, acute                           |
| Surgical    | appendicitis.  |
| (31%)       | Congenital malformations – patent ductus arteriosus,                             |
|             | ventricular septal defects, hydrocephalus, pelvic-ureteric junction obstruction. |
|             | Childhood malignancies and tumours such as Ewing's                               |
|             | sarcoma, teratoma, germ cell tumours.  |
|             | Orthopaedic indications such as limb and spine fractures                         |
|             |  |

Indications for plasma: (percentage of total plasma transfusions)

| Neonate – Surgical | Congenital malformations — tracheo-oesophageal    |
|--------------------|---|
| (74%)              | fistula (83·7%), intestinal atresias (16·3%)      |
| Neonate – Medical  | Double volume exchange transfusions (7.6%) Sepsis |
| (26%)              | (92·4%)   |
| Paediatric –       | Congenital malformations (83·7%) $-$ intestinal   |
| Surgical (37%)     | atresias, malrotation, pelvic-ureteric junction   |
|                    | obstruction, subacute intestinal obstruction      |

Trauma (16·3%) – blunt trauma abdomen, fractures, limb injury Paediatric -Bleeding disorders (6.3%) - haemophilia, protein C Medical (63%) and S deficiency, factor deficiency. Acute infections (41.2%) - liver abscess, brain abscess, septic shock, postoperative sepsis, disseminated intravascular coagulopathy Chronic infections (6.3%) - tubercular meningitis, meningoencephalitis Haematological disorders (4.7%) - haemolyticuraemic syndrome Haematological malignancies (41.5%) – acute lymphoblastic leukaemia, acute myeloid leukaemia, lymphoma

Indications for platelets: (percentage of total platelet transfusions)

| Neonate –      | Congenital malformations – such as tracheo-          |  |  |
|----------------|--|--|--|
| Surgical (81%) | oesophageal fistula                                  |  |  |
| Medical (19%)  | Acute infections – early onset of neonatal sepsis,   |  |  |
|                | necrotizing enterocolitis                            |  |  |
| Paediatric –   | Haematological malignancies (75⋅8%) – such as acute  |  |  |
| Medical (73%)  | lymphoblastic leukaemia, acute myeloid leukaemia     |  |  |
|                | and aplastic anaemia.                                |  |  |
|                | Acute infections (24·2%) – pneumonia, meningitis,    |  |  |
|                | parvo virus infection                                |  |  |
| Surgical (27%) | Congenital malformations (98%) – such as tracheo-    |  |  |
|                | oesophageal fistula, Tetralogy of Fallot, intestinal |  |  |
|                | atresia.   |  |  |
|                | Trauma (2%) — limb injury, blunt trauma abdomen      |  |  |

#### **Question 4**

It is not mandatory for the person ordering transfusion to provide the indication for the transfusion at the time the blood products are ordered. However, there is a provision for conveying the specific indication for blood component transfusion in the blood requisition form, which is subsequently entered electronically in the Hospital Information System.

## Ouestion 5

It is not mandatory for the person ordering transfusion to provide the diagnosis at the time the blood products are ordered. However, there is a provision for conveying the specific indication/diagnosis for blood component transfusion in the blood requisition form, which is subsequently entered electronically in the Hospital Information System. In addition, the importance of mentioning an indication/diagnosis on the blood order is also discussed in the hospital transfusion committee to provide necessary instructions to the user departments for compliance at regular intervals.

#### *Question 6b*

- (a) The thresholds for RBC transfusion in neonates in our institute are given as below:
  - (i) PCV <36% and requiring
    - >35% supplemental oxygen
    - MAP ≥6-8 cm H2O by CPAP or IMV
  - (ii) PCV <31% and requiring

    - >9 episodes of apnoea and bradycardia in 12 h or 2 episodes in 24 h requiring bag and mask ventilation while on adequate methylxanthine therapy
  - (iii) PCV <40% and
    - HR >180/min or RR >80/min persisting for 24 h
    - Wt gain <10 g/day for 4 days while on 100 Cal/kd/day
    - Undergoing surgery
  - (iv) PCV <21% and
    - Asymptomatic with reticulocytes <100 000/µl (2%)

The thresholds for RBC transfusion for paediatric patients in our institute are given as below:

- Hb <10 g/dl in haemodynamically unstable patients.
- Hb <7 g/dl in haemodynamically stable patients.

## Question 7b

Yes

- (a) The thresholds for platelet transfusion in neonates in our institute are given as below:
  - (i) Platelet count <50 000/μl in any preterm infant (<33 weeks) in 1st week of life – clinically unstable term infants in 1st week of life – any neonate undergoing invasive procedure (e.g. ventricular tap) – preterm neonate who has to be started ibuprofen or indomethacin or who has recentonset grade III/IV IVH
  - (ii) Platelet count  $<20~000/\mu l$  in all stable infants beyond 1st wk of life without active bleeding.
  - (iii) Platelet count <100 000/μl in the presence of alloimmune thrombocytopaenia
  - (iv) Platelet count <100 000/ $\mu l$  in the presence of active major bleeding
- (b) The thresholds for platelet transfusion for paediatric patients in our institute are given as below:
  - (i) Platelet count  $<50 000/\mu l$  in case of active bleeding other than ecchymoses or petechiae.
  - (ii) Platelet count <100 000/ $\mu$ l in intracranial bleeding or there is a condition that impairs platelet adhesion.
  - (iii) Platelet count <10 000/ $\mu$ l when there is no evidence of active bleeding other than ecchymotic, petechial

haemorrhages, and there is thrombocytopaenia from bone marrow suppression or severe sepsis.

#### **Question 8b**

- (a) The thresholds for plasma transfusion for neonates in our institute are given as below:
  - i Abnormal prothrombin time and activated partial thromboplastin time (PT  $\geq$ 17 s is abnormal in both term and preterm. APTT of >45 s in term and >55 s in preterm).
- (b) The thresholds for plasma transfusion for paediatric patients in our institute are given as below:
  - i FFP transfusion is indicated in critically ill patients with coagulopathy associated with active bleed. International normalized ratio (INR) >1.5.

## Subject 3. Product manipulations for paediatric and neonatal patients

#### Question 1b

Yes, the department of transfusion medicine has the facility for irradiation of blood components.

- (a) Following are the main indications for irradiation:
  - (i) Acute leukaemias (acute myeloid leukaemia, acute lymphocytic leukaemia)
  - (ii) Intrauterine transfusions
  - (iii) Patients undergoing stem cell transplantation.

## Question 2b

Yes, the department of transfusion medicine provides buffy coat-depleted (1 log leucoreduced) RBCs to thalassaemia and haemato-oncology patients, and these units are further leucoreduced at bedside of the patient by using third-generation leucofilters.

## **Question 3b**

Yes, the department of transfusion medicine offers washed RBCs for thalassaemia/multiply transfused patients who report recurrent severe allergic reactions to RBC transfusion.

## Ouestion 4a

No, the department of transfusion medicine does not stock thawed plasma units, but offer thawed plasma only on patient-specific demand.

## **Question** 5a

No, the department of transfusion medicine does not has facility for pathogen inactivation.

## Question 6b

Yes, the department of transfusion medicine prophylactically matches extended Rh blood group antigens (E, e, C,

c) and Kell antigen other than ABO and RhD for thalassaemia patients.

#### Ouestion 7

Department of transfusion medicine does not screen the donor units, other than the infectious markers mentioned above in reply to question 3 (subject: 1), or CMV, due to its high prevalence in general population and it is difficult get a CMV-negative units.

#### Ouestion 8

Department of transfusion medicine prepares Pedi bags from adult red cell units and split the single-donor apheresis platelet products for paediatric patients to minimize the donor exposure and optimize the product usage.

#### Ouestion 9

Smaller blood volume samples are taken from neonates and younger paediatric patients for laboratory testing and they are run the dedicated equipments for their analysis.

## Ouestion 10

Department of transfusion medicine has a policy to reserve RBCs for maximum of 72 h for a particular patient after which it can be released for other patients for optimum usage of the product.

## Subject 4. Blood product dosing

## **Question** 1

- The usually prescribed red cell dose for neonates is 10 ml/kg over 4 h.
- The usually prescribed red cell dose for paediatrics-10-15 ml/kg.

## **Question 2**

- The usually prescribed platelet dose for neonates is 10 ml/kg of PC over 1 h.
- The usually prescribed platelet dose for paediatrics is 10 ml/kg.

## Question 3

- The usually prescribed plasma dose for neonates is 10 ml/kg over 2-3 h
- The usually prescribed platelet dose for paediatrics is 10-20 ml/kg

## Reference

1 https://www.scribd.com/document/357493116/Nnf-Guideline s-2011 [Last accessed 10 July 2018].

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Agneta Wikman & Emöke Deschmann

## Sweden

Subject 1. Hospital and transfusion service demographics

## **Question** 1

Sweden.

## Question 2

Karolinska University Hospital is a teaching/academic hospital in Stockholm.

## Question 3

Blood is collected in our own collection centres and components are prepared in our own facility, a hospital blood bank.

## **Question** 4a

Neonate is defined as less than 1 month old, corrected for gestational age.

#### **Question 4b**

Paediatric patients are less than 18 years old.

## Question 5

No.

#### **Question** 6

At the hospital, both adult and paediatric patients are treated.

(a) The hospital has in total 212 paediatric inpatient beds. The paediatric departments are organized in themes, with 72 beds in neonatology, 28 beds in high specialty medicine (infections, allergy and respiratory disorders, obesitas, investigations NUD) 84 beds in surgery, paediatric intensive care unit, haematology, oncology, 28 beds in orthopaedics, neurology and habilitation.

## **Question 7**

The hospital has all specialties listed, with reservation for heart transplantations that are not done at Karolinska. In addition, the hospital has an ECMO unit, where both neonatal and paediatric patients are treated.

#### Ouestion 8a

3683 RBC transfusions were given to paediatric patients during 2017.

#### **Question 8b**

867 RBC transfusions were given in the Neonatology Department.

Information on the age of the recipients is not available.

## Question 9a

1339 plasma transfusions were given to paediatric patients during 2017.

#### **Question 9b**

375 plasma transfusions were given in the Neonatology Department.

Information on the age of the recipients is not available.

#### Ouestion 10a

1876 platelet transfusions were given to paediatric patients during 2017.

## Ouestion 10b

217 platelet transfusions were given in the Neonatology Department.

Information on the age of the recipients is not available.

Subject 2. Transfusion indications for paediatric and neonatal patients

#### **Question** 1

There are no Swedish national guidelines for neonatal and/or paediatric transfusions. Ongoing work for neonatal transfusions is not implemented yet.

#### Ouestion 2

There are local guidelines at the hospital, specific for neonatal transfusions and more general for paediatric transfusions. The guidelines differ from adult guidelines, regarding the thresholds, indications and considerations.

#### **Question 3**

The hospital has no list of indications for ordering RBC/plasma/platelet transfusions to paediatric patients.

#### Ouestion 4

It is not routine to provide the indication for the transfusion when ordering blood.

#### **Question** 5

No.

#### Ouestion 6

The local guidelines have haemoglobin thresholds for neonatal RBC transfusions [1], but the general paediatric guidelines have not.

*Neonates.* Different thresholds, from Hb  $\leq$ 120 to  $\leq$ 85 g/l, EVF  $\leq$ 0·35 to  $\leq$ 0·25 depending of the age (day 1–15) and respiratory support or not.

*Paediatrics*. There are no strict general transfusion thresholds. Indication is based on status, circulation, cause of anaemia.

#### Ouestion 7

The local guidelines have platelet count thresholds for neonatal platelet transfusions [1], but the general paediatric guidelines have not.

*Neonates*. Stable, platelet count  $<30 \times 10^9/l$ 

BW <1500 g and <7 days old, coagulation disorder  $<\!30\text{--}49\times10^9/l$ 

Bleeding, FNAIT with ICH, neurosurgery  $<50-100 \times 10^9/l$ 

Paediatrics. No strict general platelet count thresholds. Indication based on bleeding tendency. Platelet

transfusion is not indicated in stable patients with platelet count >20 ×  $10^9/l$ .

#### **Ouestion 8**

The local guidelines give no strict thresholds for plasma transfusion. Indications are coagulation disorder, DIC in septicaemia.

## Subject 3. Product manipulations for paediatric and neonatal patients

#### **Ouestion** 1

The transfusion department offers irradiated products, when requested. All neonates receive routinely irradiated products. In addition, stem cell transplanted patients and the majority of paediatric patients with malignant disorders receive irradiated products. Patients with nonmalignant haematological disorders such as sickle cell disease and thalassaemia do not receive irradiated products.

#### Ouestion 2

The transfusion department provides universal leucodepleted blood products, since almost 20 years.

#### Ouestion 3

Washed RBCs are offered to those with a previous severe allergic reaction.

## **Question 4**

The transfusion department maintains a stock of thawed plasma units for immediate issue.

4 units of blood group AB plasma are always available at the blood bank, and also, thawed plasma of other blood groups is normally available. In addition, the trauma unit has 4 units of thawed AB plasma in an external fridge.

## **Question** 5

All platelet units are pathogen-inactivated with Intercept® (Cerus) technology. Solvent-detergent plasma (Octaplas<sup>®</sup>, Octapharma AG, Austria) is available on special indications (e.g. regular plasma transfusions due to a coagulation factor deficiency, severe allergic reactions and plasmapheresis).

#### Question 6

RBCs are prophylactically matched for Rh, K, Kidd, Duffy for patients with chronic transfusion requirements (thalassaemia, sickle cell, etc) [2, 3]. In malignant disorders, RBCs are not routinely antigen-matched.

## **Question 7**

Other than mandatory testing is done only on specific indications. For example, CMV is tested if granulocyte transfusions are given. In other blood components leucodepletion is considered sufficient not to transmit CMV, also in intrauterine blood transfusions.

#### **Ouestion 8**

Procedures to minimize the number of donors the patients are exposed to are not routinely used, fresh blood components are normally prioritized.

#### Ouestion 9

The neonatal ward has a policy to minimize the volume and frequency of the samples drawn for laboratory testing; small-volume test-tubes and priority of analyses have been implemented.

#### Ouestion 10

The transfusion department has policies for the maximum storage age of RBCs that can be issued to specific patients. RBC paediatric units, 70 ml, is stored maximum 14 days after the day of collection and 24 h after irradiation, RBC units for paediatric patients and to all who requires chronic transfusions is less than 14 days. RBC units used for the preparation of blood exchange and prime blood are less than 5 days and maximum 24 h after irradiation [4].

#### Subject 4. Blood product dosing

#### Ouestion 1

The dose of RBCs prescribed for stable patients is to neonates 15-20 ml/kgBW and to paediatric patients 10 ml/ kgBW.

#### **Question 2**

The dose of platelets prescribed for stable patients is to neonates 10-15 ml/kgBW and to paediatric patients 10 ml/kgBW.

#### Ouestion 3

The dose of plasma prescribed for stable neonates is 10 ml/kgBW. In the local paediatric guidelines, the plasma dosage is not mentioned.

## References

- 1 Neonatal hematology and transfusion medicine. Clin Perinatol 2015; 42:469-684
- 2 Matteocci A, Perelli L: Red blood cell alloimmunization in sickle-cell disease and in thalassemia: current status, future perspectives and potential role of molecular typing. Vox Sang 2014; 106:197-208
- 3 La Salle-Williams M, Nuss R, Le T, et al.: Extended red blood cell matching for transfusion in sickle-cell disease: a review of 14 years experience from a single center. Transfusion 2011; 51:1732-1739

4 Whole Blood Leukocyte-Depleted For Exchange Transfusion, p 418, EDQM, 19th edn, 2017.

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## Singapore

Subject 1. Hospital and transfusion service demographics

**Question 1** Singapore.

## Ouestion 2

We are not a hospital, but we serve as Singapore's national blood service (Blood Services Group (BSG), Health Sciences Authority (HSA).

There are three main public hospitals in Singapore with neonate and paediatric services, and these are all teaching/academic hospitals. These hospitals are as follows:

- 1 KK Women's and Children's Hospital (KKH)
- 2 National University Hospital (NUH)
- 3 Singapore General Hospital (SGH primarily neonates) Blood Services Group (BSG), Health Sciences Authority (HSA), which serves as the national blood service of Singapore, supplies blood to the above hospitals. In addition, BSG also supplies blood and provide pretransfusion test services to a few private hospitals which may contribute to a small pool of the neonate/paediatric population who receives transfusion.

## Ouestion 3

See reply to Question 2.

Ouestion 4a

<1 month.

## **Question 4b**

<16 years old in KKH, <18 years old in NUH.

#### **Question** 5a

KKH is a paediatric and women specialty hospital with 365 beds for the paediatric specialty. The other hospitals are not paediatric specialty hospitals.

#### **Question** 6

All the hospitals stated in question 2 treat both adult and paediatric patients.

#### **Question** 6a

For SGH: neonatal intensive care unit 10, high depen-

dency 18, nursery 20 (neonates only).

For NUH: 100 paediatric beds.

## Question 7a

Yes at KKH and NUH.

## Ouestion 7b

Yes at KKH and NUH.

#### Ouestion 7c

Yes at KKH and NUH.

## **Question 7d**

Yes at KKH and NUH.

## **Question 7e**

Yes at KKH and NUH.

## Question 7f

Yes at KKH and NUH.

#### Question 7q

Yes at KKH and NUH.

## Ouestion 7h

Yes at KKH and NUH.

## **Question 7i**

Yes at KKH and NUH.

## **Question 7**i

Yes at KKH and NUH.

#### Question 7k

Yes at KKH and NUH.

#### Ouestion 71

Yes at KKH and NUH.

#### Ouestion 7m

Yes at KKH and NUH.

## **Question** 7n

Yes at KKH and NUH.

#### **Question 70**

Yes at KKH and NUH.

## Question 7p

Yes at KKH and NUH.

#### **Ouestion 8**

The figures stated in Questions 8-10 refer to the number of blood product units supplied to neonates and paediatric patients from BSG, HSA across the various hospitals in Singapore over the last 1 year.

## Question 8a

4394.

## Question 8b

- (a) 794.
- (b) 530.
- (c) 3070.

## Question 9a

1063.

## **Question 9b**

- (a) 320.
- (b) 106.
- (c) 637.

## **Question 10a**

2241.

## **Question 10b**

- (a) 310.
- (b) 274.
- (c) 1656.

## Subject 2. Transfusion indications for paediatric and neonatal patients

#### Ouestion 1

No.

## Question 2

There are hospital-specific guidelines.

#### Ouestion 2b

- (a) Transfusion triggers.
- (b) Provision of manipulated blood products for special groups: intrauterine transfusion, neonatl exchange transfusion, paediatric extracorporeal membrane oxygenation (ECMO), cardiac surgery.

## Question 3b

(a) Please see replies to questions 6-8 below for the indications.

## Question 4b

Yes.

- (a) KKH: yes.
- (b) NUH: yes.
- (c) SGH: yes.

## Question 5

- (a) KKH: no.
- (b) NUH: no.
- (c) SGH: no.

## Question 6b

Below are the RBC transfusion thresholds for the three major paediatric hospitals.

## Question 6b

## KKH thresholds.

## RBC transfusion thresholds for neonates

|  | Suggested transfusion threshold |   |                   |  |
|--|---------------------------------|---|-------------------|--|
| Postnatal age  | Ventilated                      | On oxygen or<br>noninvasive<br>positive pressure<br>ventilation | Off<br>oxygen     |  |
| First 24 h   | <12                             | <12   | <10               |  |
| <week (day="" 1="" 1–7)<="" td=""><td>&lt;12</td><td>&lt;10</td><td>&lt;10</td></week> | <12                             | <10   | <10               |  |
| Week 2 (day 8–14)>Week 3   | <10                             | <9.5  | <7.5 <sup>a</sup> |  |
| (day 15 onwards)   |                                 | <8.5  |                   |  |

#### RBC transfusion thresholds for neonates

|   | Suggested transfusion threshold |   |               |  |
|---|---------------------------------|---|---------------|--|
| Postnatal age   | Ventilated                      | On oxygen or<br>noninvasive<br>positive pressure<br>ventilation | Off<br>oxygen |  |
| Cumulative blood loss in 1st week of life in extremely low birthweight (Body weight <1 kg) infants requiring intensive care | 10% blood                       | volume  |               |  |
| Acute blood loss  |                                 | volume or estimated<br>BC increases Hb by                       |               |  |

 $^{a}\text{It}$  is accepted that clinicians may use up to 8.5 g/dl depending on clinical situation

#### RBC transfusion threshold for paediatrics

| Clinical situation   | Hb transfusion |
|--|----------------|
| Patients undergoing chemotherapy                             |                |
| Stable   | 8 g/dl         |
| Unstable/febrile   | 9 g/dl         |
| Patients with chronic anaemia on regular transfusion therapy | 9 g/dl         |
| Symptomatic patients or patients with ongoing blood loss     | 7 g/dl         |

#### NUH paediatric RBC transfusion thresholds.

- 1 Actively bleeding patient with haemodynamic compromise.
- 2 Symptomatic anaemia.
- 3 Regular transfusion for thalassaemia patients.
- 4 Hb <7 g/dl even if asymptomatic.
- 5 Hb <10 g/dl in hypoxic patients or patients with sepsis.

# SGH neonate RBC transfusion thresholds.

- 1 Transfuse infants at haematocrit <25% (or Hb <8 g/dl) if the neonate is asymptomatic with reticulocytes <100 000/ ul, at the discretion of the attending physician.
- 2 Transfuse infants at haematocrit less than 30% if:
  - There are significant apnoea and bradycardia (defined as >9 episodes in 12 h or 2 episodes in 24 h requiring bag-mask ventilation while receiving therapeutic doses of methylxanthines).
  - The patient requires nasal CPAP of 6 cm water or less (FiO<sub>2</sub> <35% by hood or nasal cannula).
  - The patient has persistent tachycardia (heart rate >180/min) or tachypnoea (respiratory rate of >80

- breaths/min), persisting without other explanation for 24 h.
- Weight gain of patient is deemed unacceptable (defined as weight gain of <10 g/day observed over 4 days) in the light of adequate caloric intake (at least 100 kcal/kg/day) without other explanation, such as known increases in metabolic demands or known losses in metabolic demands (malabsorption).</li>
- If the patient is scheduled for surgery, transfuse in consultation with the surgical team.
- 3 Transfuse for haematocrit levels of less than 35% in the following situations:
  - Infant with severe pulmonary disease [defined as requiring; FiO<sub>2</sub> >35% supplemental hood oxygen or continuous positive airway pressure (CPAP) or NIPPV or mechanical ventilation with a mean airway pressure of >6 cm water].
  - Infant with hypotension requiring pressor support or is critically ill.
  - Infant in whom anaemia may be contributing to heart failure.
- 4 Transfuse if there is a history of massive blood loss (blood volume of 80 ml/kg), at the discretion of the attending physician.
- 5 Transfuse if haematocrit levels less than 45% for babies on ECMO/cyanotic congenital heart disease.

## Question 7b

Below are the platelet transfusion thresholds for the three major public hospitals with paediatric and/or neonatal service.

#### KKH.

### Platelet transfusion thresholds for neonates

| Indications for platelets transfusion                   | Platelet count<br>(×10 <sup>9</sup> /I) |
|---|---|
| Neonates with no bleeding                               | <25                                     |
| including neonates with NAIT if no bleeding and no      |   |
| family history of ICH                                   |   |
| Neonates with bleeding                                  | <50                                     |
| with current coagulopathy, before surgery, or infants   |   |
| with NAIT if previously affected sibling with ICH       |   |
| Neonates with major bleeding or requiring major surgery | <100                                    |

ICH, intracranial haemorrhage; NAIT, neonatal alloimmune thrombocytopaenia.

#### Platelet transfusion thresholds for paediatrics

| Indications for platelet transfusion      | Platelet counts<br>(×10 <sup>9</sup> /l) |  |
|---|--|--|
| Bone marrow failure (eg aplastic anaemia) | 10–20                                    |  |

#### Platelet transfusion thresholds for paediatrics

| Indications for platelet transfusion | Platelet counts<br>(×10 <sup>9</sup> /l) |
|--------------------------------------|--|
| Patients undergoing chemotherapy     |  |
| Stable                               | 20                                       |
| Unstable/febrile                     | 30                                       |
| Prior to lumbar puncture             | 50                                       |
| Prior to surgical procedures         | 50-80                                    |
| Prior to neurosurgical procedures    | 100                                      |

# NUH Paediatric platelet transfusion thresholds.

- 1 Bleeding patient with thrombocytopaenia to maintain platelet count more than  $50 \times 10^9/l$  or  $100 \times 10^9/l$  ( critical site bleeding).
- 2 Severe thrombocytopaenia platelet  $<10 \times 10^9/l$  (even if patient is not bleeding; exceptions: immune/idiopathic thrombocytopaenic purpura in which platelet transfusion alone will not bring up the platelet count) or platelet  $<20 \times 10^9/l$  (depending on assessment of bleeding risk, e.g. sepsis).
- 3 Before invasive procedure which may cause bleeding if platelet count  $<50 \times 10^9/l$ .
- 4 Before surgery if platelet counts less  $<100 \times 10^9/l$  to reduce risk of bleeding.

# SGH neonate platelet transfusion thresholds

| Indications for platelet transfusion   | Platelet counts<br>(×10 <sup>9</sup> /I) |
|--|--|
| Well and stable babies   | 50                                       |
| Active bleeding, definite threat of bleeding from an   | 100                                      |
| invasive procedure or surgery, major haemorrhage   |  |
| (Grade 3-4 IVH, evolving IVH, pulmonary  |  |
| haemorrhage),  |  |
| Fulminant NEC, critically ill baby, NSAID therapy, sepsis and preterm babies $<$ 1500 g (in the first 2 weeks) |  |

IVH, intraventricular haemorrhage; NEC, necrotizing enterocolitis.

#### **Question 8b**

Below are the plasma transfusion thresholds for the three major paediatric hospitals.

KKH neonate and paediatric plasma transfusion thresholds.

- 1 Haemorrhagic disease of the newborn.
- 2 Significant coagulopathy, for example PTT >63 s or PT >25 s (1.5× longer than upper limit of normal range) with risk of significant bleeding or prior to invasive procedure.

- 3 Reversal of warfarin overdose, in combination with prothrombin complex concentrate (PCC) and vitamin K.
- 4 Clotting factor deficiencies for which no alternative clotting factor concentrates are available, for example factor XI deficiency.
- 5 Massive transfusion.

## NUH paediatric plasma transfusion thresholds.

- 1 Bleeding patient with coagulopathy (defined as PT/ aPTT >1.5 times upper limit of normal).
- 2 Prophylactic in patients with coagulopathy going for invasive procedure or surgery.
- 3 Plasma exchange.

SGH neonate plasma transfusion thresholds. Coagulopathy (PT >16 s PTT >60 s) with bleeding or risk of significant bleeding.

Subject 3. Product manipulations for paediatric and neonatal patients

## Ouestion 1b

Yes.

- (a) Following are the indications which BSG follows in the supply of irradiated and leucoreduced blood products for patients of all ages:
  - (i) Haematopoietic stem cell transplant recipients (autologous or allogenic), from the time of conditioning chemotherapy onwards;
  - (ii) Intrauterine transfusions;
  - (iii) Neonatal exchange transfusion subsequent to intrauterine transfusions;
  - (iv) Patients currently or previously treated with the following medications:
    - Purine analogue drugs such as fludarabine, cladribine, deoxycoformycin and clofarabine.
    - Bendamustine
    - Campath (alemtuzumab)
    - Antithymocyte globulin (ATG);
  - (v) Patients with suspected or confirmed congenital T-cell immune deficiency disorders;
  - (vi) Recipients of donor units known to be from a 1st- or 2nd-degree blood relative;
  - (vii) Human leucocyte antigen (HLA) compatible blood components and
  - (viii) All granulocyte products.

## **Question 2b**

Yes.

(a) Following are the indications which BSG follows in the supply of leucoreduced blood products for patients of all ages:

- (i) Patients who have developed febrile nonhaemolytic transfusion reactions on two or more occasions.
- (ii) CMV-seronegative recipients at risk of CMV transmission via transfusion, including:
  - Patients undergoing bone marrow transplants (leucoreduced and irradiated)
  - Neonates (1st 28 days of life)
  - Premature infants and/or infants weighing less than 1200 g at birth
  - Intrauterine transfusions or in neonates who previously received IUT (leucoreduced and irradiated).
- (iii) Nonhepatic solid organ transplant organ candidates to reduce the rate of human leucocyte (HLA) alloimmunization.
- (iv) Haematology patients likely to require regular transfusions of blood and blood components to reduce the rate of human leucocyte (HLA) alloimmunization.
- (v) All neonates undergoing cardiac surgery.
- (vi) All paediatric ECMO patients.
- (b) Following are the indications which BSG follows in the supply of leucoreduced blood products for patients of all ages:
  - (i) Patients who have developed febrile nonhaemolytic transfusion reactions on two or more occasions.
  - (ii) CMV-seronegative recipients at risk of CMV transmission via transfusion, including:
    - Patients undergoing bone marrow transplants (leucoreduced and irradiated)
    - Neonates (1st 28 days of life)
    - Premature infants and/or infants weighing less than 1200 g at birth
    - Intrauterine transfusions or in neonates who previously received IUT (leucoreduced and irradiated).
  - (iii) Nonhepatic solid organ transplant organ candidates to reduce the rate of human leucocyte (HLA) alloimmunization.
  - (iv) Haematology patients likely to require regular transfusions of blood and blood components to reduce the rate of human leucocyte (HLA) alloimmunization.
  - (v) All neonates undergoing cardiac surgery.
  - (vi) All paediatric ECMO patients.

# Question 3b

Yes.

- (a) Following are the indications which BSG follows in the supply of washed RBCs for patients of all ages:
  - (i) IgA deficiency.
  - (ii) Severe allergic/anaphylactic transfusion reactions.

Question 4a

No.

**Question** 5a

No.

Ouestion 6a

No.

#### Question 7

No. Leucoreduced blood products are provided as a measure for minimizing transfusion-transmitted CMV infection.

Question 8

Nο

#### **Question 9**

Yes, please refer below:

*KKH*. Neonates aged below 4 months do not require routine cross-matching if the following criteria are met: BW <1.5 kg, pretransfusion ABO/Rh available, DCT negative, maternal ABO/Rh available, negative maternal antibody screen.

*NUH*. Paediatric blood tubes are used for group and cross-match and other blood investigations.

# SGH (neonates).

- using microtubes and taking the bare minimum needed to run the sample.
- trending electrolytes using the microelectrolyte assay from the Arterial Blood Gas analyser in the neonatal intensive care unit.
- using transcutaneous CO<sub>2</sub> & O<sub>2</sub> monitors to minimize arterial blood gas sampling.
- Plan to also start transcutaneous serum bilirubin measurements to minimize blood taking.

## Question 10a

No for NUH paediatrics.

# Question 10b

Yes for KKH and SGH. Refer below.

## Question 10c

*SGH* (*neonates*). Yes – Preference for as fresh as possible RBCs (within 3 days of collection), but have used up to 5-day-old RBCs when there are supply issues.

## BSG recommends the following.

For intrauterine transfusion and exchange transfusion: red cells less than 5 days from date of collection.

- Paediatric ECMO: red cells less than 7 days from date of collection.
- Neonates of infants <5 kg undergoing cardiac surgery: red cells less than 7 days from date of collection.

Subject 4. Blood product dosing

Question 1 10-20 ml/kg.

Ouestion 2

10 ml/kg (10-20 ml/kg).

Question 3

10-15 ml/kg (10-20 ml/kg).

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# **United Kingdom**

Subject 1. Hospital and transfusion service demographics

Ouestion 1

United Kingdom.

Question 2

Yes.

**Question 3** 

Blood Centre.

**Question 4a** 

First 28 days following birth.

Question 4b

Less than 18 years old.

Ouestion 5

Yes, we are a paediatric specialty hospital with 350 inpa-

tient beds.

**Question** 6

N/A.

**Question 7a** 

Yes.

**Question 7b** 

Yes.

Ouestion 7c

No.

Question 7d

Yes.

Ouestion 7e

Yes.

Question 7f

Yes.

Question 7g

No.

**Question 7h** 

Yes.

Ouestion 7i

Yes.

Question 7j

Yes.

**Question 7k** 

Yes.

**Question 71** 

Yes.

**Question 7m** 

Yes.

**Question** 7n

No.

Question 70

Yes.

Question 7p

Yes.

#### Ouestion 8a

5250 RBCs are transfused to paediatric and neonatal patients per year.

## Question 8b

Information not available.

## **Question** 9a

1700 plasma units are transfused to paediatric and neonatal patients per year.

# Question 9b

Information not available.

#### Ouestion 10a

4457 platelet units are transfused to paediatric and neonatal patients per year.

#### Ouestion 10b

Information not available.

# Subject 2. Transfusion indications for paediatric and neonatal patients

#### **Question** 1

Yes (New HV *et al.*, on behalf of the British Society for Haematology. Guidelines on transfusion for fetuses, neonates and older children. *Br J Haematol* 2016; 175:784–828).

The aim of the guidelines is to provide healthcare professionals with clear guidance on the management of all aspects of paediatric transfusion, available all in one place. Unlike the UK adult transfusion guidelines, which mostly focus on specific areas of transfusion, the guidelines have a wide scope, including areas such as cardiac surgery and major haemorrhage. They review the paediatric-specific literature and make recommendations vs consensus-based 'key practice points' depending on the quality of evidence available. They include both clinical and laboratory sections, covering neonatal/paediatric transfusion indications, administration, pretransfusion testing and component selection, and details of specialized blood components for fetuses/neonates/infants.

#### Ouestion 2

Yes, we have local guidelines specific for neonatal and paediatric transfusions. We have no adult guidelines.

#### **Question 3**

Yes, we have a list of indications for ordering RBC/plasma/platelets, but within the current 40-page blood transfusion policy and not succinct.

#### Question 4

It is currently not mandatory, but we will be introducing a new electronic patient record system in 2019 that will make it mandatory.

#### **Question** 5

No, it is not mandatory but is considered best practice to do so.

#### Ouestion 6

For RBC transfusions, our thresholds are based on British Society for Haematology (BSH) guidelines (New *et al.*, 2016).

# Question 7

Our platelet transfusions are based on BSH guidelines (New *et al.*, 2016).

#### Question 8

Our plasma transfusions are based on BSH guidelines (New *et al.*, 2016).

# Subject 3. Product manipulations for paediatric and neonatal patients

# Question 1

Yes, we offer irradiated products. Please see attached document 'Blood Transfusion Special Requirements'.

#### Ouestion 2

Yes, we offer leucodepleted RBCs. They are standard issue in the UK (universal prestorage leucodepletion).

#### **Question 3**

Yes, we offer washed RBCs, based on consultant decision postrecurrent transfusion reactions.

#### Question 4

Yes, we maintain a stock of thawed plasma. We keep 1 unit of AB solvent-detergent FFP (SDFFP).

#### Question 5

Yes, we offer SDFFP and methylene-blue-treated cryoprecipitate.

#### **Question** 6

Yes, we match for the following:

- Transfusion-dependent patient with Rh-phenotyped units.
- Patient with clinically significant antibodies with units Rh-phenotyped and negative for their clinically significant antibody based on current clinical guidelines.

## Ouestion 7

Yes, CMV testing is done at NHSBT.

#### Ouestion 8

Yes, we minimize exposure by the use of paedipacks (split packs for neonatal red cells and neonatal platelets as provided by NHSBT).

## Question 9

Yes.

# Ouestion 10

Yes, our policy follows:

- Patients under 1 year requiring large volume neonatal pack transfusions (LVT) <5 days.
- Cardiac surgical patients over 1-year units <14 days.
- Sickle cell disease patients units <14 days.
- Irradiated units <14 days (all patients except neonatal/infant large volume transfusion and all cardiac

surgery where irradiated blood is used within 24 h postirradiation).

## Subject 4. Blood product dosing

#### Ouestion 1

This is the recommendation for a target Hb:

Volume to be transfused =

Desired Hb (g/l) – actual Hb (g/l)  $\times$  weight (kg)  $\times$  4\*

\*It is reasonable to use a Factor of 4 (practice varies between 3 and 5), but this should be assessed on an individual patient basis.

With a local volume recommendation of:

- Neonates 15 ml/kg
- Paediatrics 3–4 ml/kg to raise Hb by 10 g/l

#### Question 2

We prescribe 10-15 ml/kg.

#### Question 3

We prescribe 10-15 ml/kg.

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# **BLOOD TRANSFUSION SPECIAL REQUIREMENT REQUESTS**

# FILE FORM AT FRONT OF PATIENT NOTES

A COPY OF THIS FORM  $\underline{\textit{MUST}}$  BE FAXED TO BLOOD TRANSFUSION ITEMS WITH \* ARE MANDATORY

# First form/updated form\* (delete where applicable)

| -  |                          | •   |                         | •                   |                      |             |                  |
|--|--------------------------|---|-------------------------|---------------------|----------------------|-------------|------------------|
| *Patient surname                                   |                          |   |                         | *Patient            | forename             |             |                  |
| *Hospital number                                   |                          |   |                         | *Date of            | Birth                |             |                  |
| *Diagnosis   |                          |   |                         | 1                   |                      |             |                  |
| Please include any antibodies                      | atypical                 |   |                         |                     |                      |             |                  |
| The chart overleaf g<br>equirements. Tick<br>late. |                          |   |                         |                     |                      |             |                  |
| *Requirement                                       |                          | Tick all that<br>(leave blan<br>longer requ | k if no                 | *Reason<br>See over | for requirem<br>leaf | ent/remova  | I *Review period |
| CMV negative                                       |                          |   | ,                       |                     |                      |             |                  |
| Irradiated compo                                   | onents                   |   |                         |                     |                      |             |                  |
| HLA matched components                             |                          |   |                         |                     |                      |             |                  |
| Donor Group  | Recipie                  |   | Red cel                 |                     | Renal (Ind           |             | Plasma           |
| RhD  | Recipient group Red cell |   |                         |                     |                      | RhD         |                  |
| laemoglobinopathi                                  | ies_                     | Referring ho                                | spital                  |                     | Last Trans           | fusion Date |                  |
|  |                          |   |                         |                     |                      |             | antibodies       |
| Authorised by                                      |                          |   |                         |                     |                      | Τ           |                  |
| *Name  |                          |   |                         | *Bleep ı            | number               |             |                  |
| *Position  |                          |   |                         | *Date               |                      |             |                  |
| *Form FAXED to<br>8288                             |                          |   |                         | Signed:             |                      |             |                  |
| Laboratory Use Only:                               |                          |   |                         |                     |                      |             |                  |
| Received in laboratory (Print name):               |                          | Signature: D                                |                         | Date                | / Time:              |             |                  |
| Entered on OmniLab (Print name)                    |                          |   | Signature: Date /       |                     | / Time:              |             |                  |
| #SBMS Check:(Print name):                          |                          |   | Signature: Date / Time: |                     |                      | / Time:     |                  |
| #Senior/Chief BMS to                               | check for a              | ccuracy and appr                            | opriatenes              | s. Refer to Ha      | em SpR if requ       | ired.       |                  |

# **Guidance on Special Requirements**

| CMV (C  | ytomegalovirus) negative req  | uirements   |  |  |
|---|---|---|--|--|
| CMV Negative<br>Components                    | <ul> <li>Neonates &lt;6 months</li> <li>Intrauterine transfusion</li> <li>Exchange transfusion</li> <li>CMV negative recipients of allogeneic SCT</li> <li>SCID</li> <li>Pregnancy for patients requiring transfusion (except in emergency).</li> </ul>   | Granulocytes – CMV negative if recipient negative Granulocytes must always be irradiated  |  |  |
|   | Irradiation requirements  |   |  |  |
| Stem Cell<br>Transplantation<br>(BMT or PBSC) | Allogeneic recipients – require irradiated co-<br>conditioning until Consultant states otherwise<br>All donors require irradiated components fro<br>the harvest.<br>Autologous recipients – require irradiated of<br>during harvest. Then from start of conditioning<br>transplant.   | m 7 days prior to and during  |  |  |
| Neonates                                      | Intrauterine Transfusion (IUT) – all compon<br>Neg and irradiated (use within 24 hours post<br>Neonates post IUT: all components including<br>Neg and irradiated up to 6 months post EDD.<br>Exchange Transfusion (ET) – Blood must b<br>within 24 hours post irradiation).   | irradiation).<br>ET are required to be CMV  |  |  |
| Miscellaneous/<br>Specific Drugs              | Purine Analogues – such as fludarabine, cladeoxycoformycin, clofarabine, nelarabine and Alemtuzumab (Anti-CD52) (Campath, Mabel Human Leucocyte Antigen (HLA) Matched from first or second degree relatives.  Known or suspected Congenital Immunod SCID, Wiskott Aldrich or Di George Syndro components are not required except for SCII Asplenia, together with any other immunode not qualify)  Aplastic Anaemia (on Anti-Thymocyte Glof Hodgkin's Lymphoma Granulocytes –need to consider if the pati | d bendamustine Campath) Components and components  deficiency – such as CID, ome (CMV negative blood D) ficiency. (Asplenia alone does bulin (ATG) treatment) |  |  |
| Other Requirements                            |   |   |  |  |
| Haemoglobinopathies                           | These patients are regularly transfused eigunder shared care  Phenotyped Blood – to help prevent the form patients are given Rh and K negative red cells are also given antigen negative red cells. The therefore order in advance (24 hours preferated Transfusion History – please inform the label last transfusion occurred.  | mation of allo-antibodies, all<br>s. If antibodies develop, they<br>ese may not be standard stock<br>bly)   |  |  |

**NB.** Remember to complete special requirement boxes on the pink blood prescription chart, if the requirement is new or has been amended please start a new chart.

Blood Transfusion Special Requirements Requests Page 2 of 2

Version 7 Issued August 2017 Review date August 2019

Anne Kinmonth, Mary Comande, Helen Savoia & Gemma Crighton

#### Australia

Subject 1. Hospital and transfusion service demographics

Question 1
Australia.

#### Ouestion 2

Yes, it is major specialist paediatric hospital, providing tertiary care for Victoria, Tasmania, southern New South Wales (as well as specialist treatment for patients from other states and overseas) and is a major teaching hospital.

#### Ouestion 3

Our hospital receives blood components from the Australian Red Cross Blood Service; requests for specialized blood products such as phenotype-matched or CMV-seronegative components are directed to the Blood Service. However, we have an onsite irradiator and irradiate our own components.

#### **Question 4a**

In our hospital, the definition of a neonate, specified by the neonatal department, is an infant that is less than 28 days of age.

#### Ouestion 4b

In our hospital, a paediatric patient is considered someone less than 18 years of age.

### Ouestion 5

357 inpatient beds.

#### Question 6

No, it only treats paediatric patients. On occasion, we will treat a young adult while they are awaiting transition to adult services.

Question 7a

Yes.

Question 7b

Yes.

**Question 7c** 

Yes.

Question 7d

Yes.

Question 7e

Yes.

Question 7f

Yes.

Question 7q

Yes.

**Question 7h** 

Yes.

**Question 7i** 

Yes.

**Question 7**i

Yes.

Question 7k

Yes.

Question 71

Yes.

Question 7m

Yes.

**Question** 7n

Yes.

Question 70

Yes.

Question 7p

Yes.

Question 8

Approximately 4700 RBC per year.

Question 9

Approximately 2350 per year.

Question 10

Approximately 3100 per year.

Subject 2. Transfusion indications for paediatric and neonatal patients

# Question 1

Yes, the National Blood Authority, Australia, has developed a series of six patient blood management (PBM) modules and Module 6 specifically focuses on evidence-

based neonatal and paediatric PBM. The Modules include recommendations based on systematic review, practice points which are based on consensus where systematic review found insufficient high-quality data to produce evidence-based recommendations. Finally, it includes expert opinion points based on consensus where relevant guidance is required. Module 6 includes general information about blood product modifications, foetal transfusion, blood conservation strategies and iron deficiency anaemia. The Appendices include information on transfusion volume calculations, a preoperative haemoglobin (Hb) assessment and optimization template, a critical bleeding template and dosing information for intravenous iron and tranexamic acid.

The Australian and New Zealand Society of Blood Transfusion have developed local guidelines, which provide guidance for paediatric, neonatal and foetal transfusions, and they include specific information on blood product administration, transfusion volume calculations and rates of infusion, in addition to blood product modifications.

The Transfusion Orientation Pack produced by the Australian Red Cross Blood Service provides specific paediatric guidance for red cell and platelet transfusion triggers and dosing.

Our hospital's blood product prescription guideline has been adapted for use in our state in Australia.

## **Question 2**

Yes, our hospital has developed its own neonatal and paediatric blood transfusion and blood management resources and guidelines for our clinicians. We do not have any specific adult transfusion guidelines on our website. Clinicians access our guidelines both internally from within our hospital, locally across the state and nationally across Australia.

Our guidelines differ from adult guidelines because they provide specific guidance for the different clinical scenarios that affect neonates and children and take into account some of the specific differences between these patient groups.

Our guidelines provide guidance for specific, transfusion indications, consent for transfusion, refusal of transfusion consent, blood group and antibody testing, calculating transfusion volumes, blood product modifications such as indications for CMV-seronegative blood products, fresh products and irradiated blood products.

Transfusion consent for example in an adult involves a discussion between the consenting adult and the clinician, whereas in children, it involves a discussion with the parents or guardians and may involve discussion with the child. Refusal of transfusion consent by a parent/guardian is a unique situation, and in this instance, parents' wishes may be over-ridden if a transfusion is considered life-

Most neonates are eligible for extended expiry for their blood group and antibody screening, because the development of antibodies to red cell antigens is very uncommon in the first 4 months of life and this reduces the requirement for repeated sampling.

There are a number of different considerations in neonates and children that determine whether a transfusion is indicated and these differ compared with adults. Neonates in particular have higher normal Hb values ranges compared with adults and children. Red cell transfusion thresholds in neonates depend on the gestational age, the postnatal age, the Hb concentration and the degree of respiratory support the neonate is receiving. There are a number of unique transfusion indications affecting only fetuses and neonates, such as foeto-maternal alloimmune thrombocytopaenia, intrauterine transfusion for haemolytic disease of the newborn and exchange transfusions and these have their own specific considerations such as human platelet antigen (HPA) or red cell alloantibody matching.

Transfusion volumes for neonates, infants and small children must be carefully calculated based on ml/kg and prescribed in ml (not units) with a specific appropriate transfusion rate.

There are some specific blood product modifications such as phenotype matched, irradiated. CMV-seronegative and washed blood products where indications differ between children and adults or the requests occur more frequently in children.

# **Question 3** Yes

| Paediatrics<br>Haemoglobin<br>threshold | Indication/clinical information   |
|---|---|
| Hb <70 g/l                              | Often indicated, however, lower thresholds may be acceptable in patients without symptoms (symptoms may include – tachycardia, flow murmur, lethargy, dizziness, shortness of breath and cardiac failure) and where specific therapy (e.g. iron) is available |
| Hb 70–90 g/l                            | RBC may be indicated, depending on the clinical setting, for example the presence of bleeding or haemolysis and clinical signs and symptoms of anaemia  Patients with disease or therapy-related bone marrow  |
| Hb >90 g/l                              | failure<br>RBC transfusion is often unnecessary and may be<br>inappropriate<br>Sickle cell disease  |

| Paediatrics<br>Haemoglobin<br>threshold | Indication/clinical information  |
|---|--|
|   | mulcation/clinical information   |
| Threshold not                           | Blood prime for circuit  |
| specified                               | Red cell exchange transfusion  |
|   | Haemolysis   |
|   | Clinically significant acute blood loss  |
| Transfusion may situations              | be indicated at higher thresholds for specific                                     |
| Preterm neonate                         | S  |
| Children with cy                        | anotic heart disease or on ECLS  |
|   | emoglobinopathies (thalassaemia or SCD) or congenital hronic transfusion programme |
| Neonates                                |  |
| Haemoglobin                             | Indication/clinical information  |

| Neonates<br>Haemoglobin<br>threshold | Indication/clinical information                                     |
|--------------------------------------|---|
| Hb <100-120 g/l                      | Preterm or term neonate with no respiratory support, week 1 of life |
| Hb <85-100 g/l                       | Preterm or term neonate with no respiratory support, week 2 of life |
| Hb <70-100 g/l                       | Preterm or term neonate with no respiratory support, week 3 of life |
| Hb <110-130 g/l                      | Preterm neonate with respiratory support, week 1 of life            |
| Hb <100-125 g/l                      | Preterm neonate with respiratory support, week 2 of life            |
| Hb <85-110 g/l                       | Preterm neonate with respiratory support, week 3 of life            |
|                                      | Clinically significant acute blood loss                             |

ECLS, extracorporeal life support; RBC, red blood cell; SCD, sickle cell disease.

## Question 4

Yes, it is mandatory.

Yes, the indication is entered electronically.

## Question 5

No, there is a space where the patient's clinical information or diagnosis can be entered, but it is not mandatory. As we have an electronic medical record, it is possible, to review in real time the patient's clinical diagnosis, medical notes and laboratory investigations to determine transfusion indication.

# Question 6

Yes, thresholds are included with some of the indications, please see table above.

# Question 7

Yes.

| Paediatrics |  |  |
|-------------|--|--|
|             |  |  |

| Platelet count            | Indication/clinical information   |
|---------------------------|---|
| <10 × 10 <sup>9</sup> /l  | Clinically stable paediatric patients receiving chemotherapy for leukaemia or post-HSCT Clinically stable patients with solid tumours (prophylactic) <sup>a</sup> <sup>a</sup> Transfusions at higher levels may be required for bladder, brain or necrotic tumours |
|                           | Critically ill patients with no bleeding  |
| <20 × 10 <sup>9</sup> /l  | Chemotherapy, HSCT and risk factors (e.g. fever, sepsis, minor bleeding, mucositis and DIC without bleeding)  |
|                           | Critically ill patients with no bleeding and risk<br>factors (e.g. sepsis, renal failure and medications)<br>Nasogastric tube insertion   |
|                           | Intramuscular injections, for example Erwinia asparaginase  |
|                           | Insertion of a nontunnelled central venous line   |
| <30 × 10 <sup>9</sup> /l  | LP and ongoing chemotherapy-induced thrombocytopaenia   |
|                           | CNS tumour and:   |
|                           | A VP shunt or Ommaya reservoir  |
|                           | Has a gross total resection and is receiving  |
|                           | chemotherapy and/or radiation   |
|                           | Has residual tumour and is receiving  |
| 9.6                       | chemotherapy and/or radiation   |
| $<50 \times 10^{9}/I$     | LP and new disease induced thrombocytopaenia  |
|                           | Patient undergoing invasive procedure (including  |
|                           | tunnelled central venous line insertion)  |
|                           | Moderate active bleeding (including bleeding associated with DIC)   |
|                           | CNS tumour and:   |
|                           | A past history of intracranial haemorrhage  |
|                           | Is receiving an anti-angiogenesis agent   |
| <75 × 10 <sup>9</sup> /l  | Major haemorrhage due to trauma or significant postoperative bleeding (e.g. postcardiac surgery)  |
| <100 × 10 <sup>9</sup> /l | Patient undergoing high-risk invasive procedure (e.g. neurosurgery/ophthalmology)   |
|                           | 3 / /   |
|                           | ECLS (lower platelets may be acceptable in stable patients)   |

# Platelet transfusion not indicated

Stable patients with chronic, stable, severe thrombocytopenia due to: Alloimmunization, ITP, TTP, aplastic anaemia or MDS

These patients should receive platelet transfusions with clinically significant bleeding only

BMA and trephine biopsy

Intravenous cannula insertion

| Platelet threshold       | Indication/clinical information  |
|--------------------------|--|
| <30 × 10 <sup>9</sup> /I | Stable term or preterm neonate with asymptomatic thrombocytopaenia and no bleeding |
| $30-50 \times 10^9/I$    |  |

| Paediatrics               |  |  |
|---------------------------|--|--|
| Platelet count            | Indication/clinical information  |  |
|                           | Preterm neonate with thrombocytopaenia being treated for sepsis or requiring respiratory support   |  |
| <50 × 10 <sup>9</sup> /l  | Term or preterm neonate with bleeding symptoms (mucocutaneous, gastrointestinal, petechiae/purpura), coaqulopathy or prior to surgery      |  |
| <100 × 10 <sup>9</sup> /l | Term or preterm neonate with major bleeding (drop in Hb requiring RBC transfusion) or those that require major surgery (e.g. neurosurgery) |  |

BMA, bone marrow aspirate; CNS, central nervous system; DIC, disseminated intravascular coagulopathy; ECLS, extracorporeal life support; Hb, haemoglobin; HSCT, haematopoietic stem cell transplantation; ITP, immune thrombocytopaenia; LP, lumbar puncture; MDS, myelodysplastic syndrome; RBC, red blood cell; TTP, thrombotic thrombocytopaenic purpure; VP, ventriculo-peritoneal.

# Question 8

Yes

| Doodiotuios | a al | <br>- fresh-frozen | laamaa |
|-------------|------|--------------------|--------|
|             |      |                    |        |

| Paediatrics and neonates – fresh-frozen plasma |   |  |  |
|--|---|--|--|
| Laboratory parameter                           | Indication/clinical information   |  |  |
| Significant                                    | Acute bleeding  |  |  |
| coagulopathy                                   | Liver disease, with clinically significant bleeding or in<br>the context of coagulopathy post liver<br>transplantation  |  |  |
|  | Acute DIC with bleeding   |  |  |
| Prolonged INR                                  | Warfarin reversal, in the presence of significant or life-threatening bleeding or prior to emergency surgical procedures  • Given in addition to vitamin K  NOTE: vitamin K-dependent clotting factor concentrates (e.g. Prothrombinex) may be given instead of FFP for bleeding secondary to warfarin or emergency warfarin reversal |  |  |
| Not specified                                  | During massive transfusion or cardiac bypass for the treatment of bleeding  |  |  |
| Not specified                                  | Plasma exchange for the treatment of TTP  |  |  |
| Not specified                                  | Specific factor deficiencies where a factor concentrate is not available  |  |  |

## FFP is not indicated for

The correction of minor coagulation abnormalities (minor prolongation of the INR/APTT) in the nonbleeding child

Liver disease when there are minor coagulation abnormalities and no bleeding

For reversal of a INR <2.0 in patients undergoing minor procedures

APTT, activated partial thromboplastin time; DIC, disseminated intravascular coagulopathy; FFP, fresh-frozen plasma; INR, international normalized ratio; TTP, thrombotic thrombocytopaenic purpura.

| Laboratory parameter | Indication/clinical information   |
|----------------------|---|
| Fibrinogen <1.5 g/l  | Active bleeding   |
|                      | During massive transfusion or cardiopulmonary bypass  |
| Fibrinogen <1·0 g/l  | Acquired fibrinogen deficiency of acute DIC for the treatment of bleeding   |
| Fibrinogen <1·0 g/l  | Prior to an invasive procedure and there is a risk of significant bleeding associated with the surgery or it is at a critical site (e.g. neurosurgery or eye surgery) |
| Hyperfibrinolysis    | During massive transfusion or cardiopulmonary bypass  |

#### Cryoprecipitate is not indicated for

Nonbleeding children with mildly reduced fibrinogen levels Liver disease when there are minor coagulation abnormalities and no bleeding

| Additional neonate indication – cryoprecipitate |                                 |  |  |  |
|---|---------------------------------|--|--|--|
| Laboratory parameters                           | Indication/clinical information |  |  |  |
| Fibrinogen <0·5 g/l                             | Active bleeding in the neonate  |  |  |  |

DIC, disseminated intravascular coagulopathy.

# Subject 3. Product manipulations for paediatric and neonatal patients

# Question 1

Yes, our hospital has an onsite irradiator and is able to provide irradiated red cells. Our hospital has a universal blood product irradiation for patients in the following units: paediatric intensive care unit (PICU), neonatal unit (NNU) and oncology patients, in addition infants under 3 months of age and patients treated with extracorporeal life support (ECLS) or left-ventricular assist devices receive irradiated components. Although not all patients in these units are at risk of transfusion-associated graft-versus-host disease, this policy ensures patients who require irradiated products are not missed. In addition, blood is irradiated for intrauterine and exchange transfusions, directed donations and immune-compromised patients, this includes immunology patients with conditions such as severe combined immune deficiency (SCID) and common variable immune deficiency (CVID), as well as patients with malignancy, transplantation recipients (bone marrow, cord blood and solid organ).

#### Ouestion 2

All red cells and platelets components (both apheresis and pooled) manufactured by the Blood Service are prestorage leucodepleted. The leucodepletion system used by the Blood Service enables leucodepletion of some but not all clinical plasma. As a result, there is a mixed inventory of leucodepleted and nonleucodepleted clinical plasma components.

#### **Ouestion 3**

Washed blood components are produced by the Blood Service on special requests. Requests for these components are usually discussed with the laboratory haematology team. Indications include patients with IgA deficiency and patients with severe allergic reactions to previous red cell or platelet transfusions.

#### Question 4

Yes, two units of AB plasma are available for emergency use and for life-threatening bleeding.

#### **Question** 5

No, at present, the Blood Service does not offer pathogen-inactivated blood components.

#### **Question** 6

At present, all patients at our institution receive Kellnegative blood, due to the risk of Kell alloimmunization, by not matching for Kell in young girls with child-bearing potential. Patients with haemoglobinopathies such as sickle cell disease and patients with congenital anaemia will have red cell genotyping assessment at diagnosis. Their transfusions will be matched for Rh (C, D and E) and Kell. Patients receiving exchange transfusions for SCD or RBC transfusion for warm autoimmune haemolytic anaemia will have RBC most suitably matched to negative antigens of Rh/Kell, Jka, Fy and Ss systems.

If a patient forms a red cell alloantibody, they will have extended red cell matching and antigen-negative blood will be provided for any alloantibodies they may have or at risk of making and consideration given for matching for Fya, Jk, M, N and Ss.

Fetuses receiving an intrauterine transfusion and neonates receiving a exchange transfusion for haemolytic disease of the foetus and newborn will be provided with red cells that are RhD-negative (or RhD identical with neonate if mother not Rh immunized), Kell-negative and antigen-negative for any maternal alloantibodies. If there is time, an extended maternal phenotype will be performed and consideration given for provided red cells matched for Fya, Jk, M, N and Ss.

#### **Ouestion** 7

CMV testing is performed on an ad hoc basis by the treating clinician and is not performed by the transfusion laboratory. We have a policy, which outlines in which patients' CMV-seronegative blood products are indicated. CMV-seronegative blood products are indicated for intrauterine transfusion, neonatal exchange transfusions, preterm and term infants up to 28 days postestimated due date, patients with SCID, who are CMV-negative

(including those undergoing haematopoietic stem cell transplantation), pregnant women and granulocyte infusions. All other indications – haematology/oncology patients, allogeneic and autologous stem cell transplantation, solid organ transplant patients and other immunodeficiency patients, leucocyte deplete blood products – are considered equivalent to CMV-seronegative products.

#### Ouestion 8

We attempt to reduce donor exposure in the multiple-transfused haemato-oncology patient and allocate apheresis platelets when possible. If we are aware a neonate is going to need multiple transfusion, we will attempt to minimize donor exposure by allocating pedipaks. Since implementation of an electronic medical record, blood components are now generally only issued at the time a request is made for a transfusion. This means they are not allocated prior to transfusion, and therefore, a neonate does not have blood components allocated prior to transfusion unless we are aware they are going to require multiple transfusions.

#### Question 9

We have no formal policy in place to minimize the volume and frequency of blood samples drawn; however, we do have a number of strategies in place, which aim at blood conservation in these patients. We have paediatric collection tubes, which are standard of care for our neonates and small paediatric patients. Patients in the NNU and PICU have discard volumes returned after sampling. We use of point of care testing for INRs for patients on warfarin, arterial and venous blood gas sampling and glucose analysis for diabetics. We have a policy for neonates for extended blood group and antibody testing in the first 4 months of life if the DAT is negative and there are no maternal alloantibodies.

# **Question 10**

Yes, in Australia, red cell units can be stored up until 42 days. The following table gives information regarding the age of red cells on the day of issue for particular patient groups.

| Patient group                 | Age of red<br>cells on<br>day of<br>issue | Consideration  |
|-------------------------------|---|--|
| Neonatal exchange transfusion | <5 days                                   | Kell, negative, CMV<br>negative, Rh and<br>phenotype matched |
| Intrauterine transfusion      | <5 days                                   | Kell, negative, CMV<br>negative, Rh and<br>phenotype matched |

| Patient group   | Age of red<br>cells on<br>day of<br>issue | Consideration  |
|---|---|--|
| Paediatric large volume transfusion <sup>a</sup> Cardiac surgery requiring cardiopulmonary bypass ECLS Craniofacial surgery   | <7 days                                   | In an emergency, the<br>blood bank scientist<br>will issue the most<br>appropriate unit<br>available at the time |
| Massive transfusion   | Aim<br><10 days                           | In an emergency, the<br>blood bank scientist<br>will issue the most<br>appropriate unit<br>available at the time |
| Paediatric-routine transfusion Postoperative surgical patient Oncology patient Solid organ transplantation  | Standard<br>issue                         |  |
| Chronically transfused patients Congenital anaemia who are transfusion dependent Haemoglobinopathies (SCD, thalassaemia) Aplastic anaemia haemoglobinopathies are matched with Rh and Kell Extended phenotype matching for patients with alloantibodies Extended phenotype matching may take precedence over age of red cells | Aim<br><14 days                           | Patients with  |
| Small-volume (top up) neonatal<br>transfusions<br>Children and adolescents not<br>included in any groups listed<br>above  | Standard<br>issue<br>Standard<br>issue    |  |

CMV, cytomegalovirus; ECLS, extracorporeal life support; SCD, sickle cell disease.

# Subject 4. Blood product dosing

#### Ouestion 1

The transfusion volume for a red cell transfusion is calculated for children <20 kg from the equation

$$\begin{array}{l} ml = weight \text{ (kg)} \times Hb \text{ (g/l) rise(desired Hb} - actual \text{ Hb)} \\ \times 0.5 \end{array}$$

For children >20 kg - 1 unit of red cells is prescribed.

#### Question 2

|   | <10 kg                         | 10–<br>19 kg                   | 20–<br>29 kg                   | 30–<br>39 kg  | >40 kg        |
|---|--------------------------------|--------------------------------|--------------------------------|---------------|---------------|
| Pooled platelets (adult unit)                     | 10 ml/kg                       | 10 ml/kg                       | 10 ml/kg<br>up to 1<br>unit    | 1 unit        | 1 unit        |
| Apheresis<br>platelets<br>(adult<br>unit)         | 5–10 ml/<br>kg                 | 5–10 ml/<br>kg                 | 5-10 ml/<br>kg up to<br>1 unit | 1 unit        | 1 unit        |
| Paediatric<br>apheresis<br>platelets<br>(pedipak) | 5–10 ml/<br>kg or 1<br>pedipak | 5–10 ml/<br>kg or 2<br>pedipak | 3–4<br>pedipaks                | 4<br>pedipaks | 4<br>pedipaks |

#### Question 3

A dose of 10-20 ml/kg of fresh-frozen plasma is prescribed for stable paediatric or neonatal patients with consideration given to the average fresh-frozen plasma unit size of about 284 ml. A dose of 5-10 ml/kg of cryoprecipitate is prescribed for stable paediatric or neonatal patients, with consideration given to the typical unit size of approximately 37 ml.

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<sup>&</sup>lt;sup>a</sup>Equivalent to a single circulating blood volume (~80 ml/kg).

Joanne Yacobovich & Vered Yahalom

Israel

Subject 1. Hospital and transfusion service demographics

Question 1
Israel.

Question 2 Yes.

Question 3

Prepared in a neighbouring facility.

Question 4a
Less than 30 days old.

Question 4b

Less than 18 years old.

Question 5a

300 inpatient beds, 100 are intensive care/emergency.

Question 6 No.

Question 7a Yes.

Question 7b

Yes.

Question 7c Yes.

Question 7d Yes.

Question 7e Yes.

Question 7f Yes.

Question 7g Yes.

Question 7h Yes.

Question 7i

Yes.

Question 7j Yes, both.

Question 7k

Yes, including dialysis.

Question 7l Yes.

Question 7m

Yes.

Question 7n Yes.

Question 70 Yes.

Question 7p Yes.

*Question 8a* >5000 in 2017.

Ouestion 9

>700 in 2017 (variable volume).

Question 10

>2150 in 2017 (variable volume).

Subject 2. Transfusion Indications for paediatric and neonatal patients

Ouestion 1b

Yes – Only for platelets, approved by Israeli paediatric haematology–oncology society.

(a) Higher platelet thresholds for preterm babies, sick newborns and babies on ECMO.

Platelet dosing per kg weight up to 1 SDP.

Question 2b

Yes - Lower threshold for RBC Tx (see 3b).

Question 3b

Yes.

RBC.

• HB <8.0 g/dl for anaesthesia, patients on oxygen, fever,

- HB <6.0 g/dl in general,
- HB 6-8 g/dl can consider RBC

## PLT.

- If <10 000/µl
- or <20 000 for febrile patients,
- <80 000 postneurosurgery

#### **Question 4b**

Yes - requested not mandatory.

(a) No.

Question 5

Yes.

**Ouestion** 6b

Yes – see Q.3.

Question 7b

Yes - see 0.3.

**Question 8a** 

No.

Subject 3. Product manipulations for paediatric and neonatal patients

**Question 1b** 

Yes - Haematooncology and oncology patients.

Question 2b

Yes - Haematooncology and oncology patients.

**Question 3b** 

Yes – Severe transfusion reactions.

Ouestion 4b

Yes -2 units of every ABO blood group.

**Question** 5a

No.

**Question** 6b

Yes - Haemoglobinopathies, AIHA patients.

**Question 7** 

CMV-negative components for selected patients - SCID CMV negative.

**Question 8** 

Only if required by the treating physician.

Ouestion 9

Yes, microtainers for CBC, smaller tubes for coagulation tests, finger prick monitoring for INR monitoring in inpatients.

Question 10b

Yes - 7 days exchange Tx, preterm. Ten days thalassaemia/SCD patients (up to 14 if multiple antibodies).

Subject 4. Blood product dosing

**Question** 1

10-15 ml/kg.

**Question 2** 

15 ml/kg up to 1 SDP unit.

**Question 3** 

10-15 ml/kg.

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Wendy Lau

# Canada

Subject 1. Hospital and transfusion service demographics

Question 1

Canada.

**Question 2** 

Yes.

**Question 3** Blood center.

Ouestion 4a

Neonate less than 4 months.

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Question 4b

Paediatric less than 18 years.

**Question** 5

Yes, 350 beds.

Question 7a

Yes.

**Question 7b** 

Yes.

**Question 7c** 

Yes.

**Question 7d** 

Yes.

Question 7e

Yes.

Question 7f

Yes.

Question 7g

Yes.

Question 7h

Yes.

Question 7i

Yes.

Question 7j

Yes.

Question 7k

Yes.

**Question 71** 

Yes.

Question 7m

Yes.

Question 7n

Yes.

Question 70

Yes.

Question 7p

Yes.

#### Question 8a

Just over 8000 units per year (range from 8000 to 8700 units in recent years).

#### Question 8b

It is not possible at this time to subgroup by age, maybe in the future (with help with EPIC/HCLL).

#### Question 9a

About 1500 units per year (range 1400–1600 in recent years).

#### **Question 9b**

It is not possible at this time to subgroup by age, maybe in the future (with help with EPIC/HCLL).

#### **Question 10a**

Just over 3500 units (total of pools and apheresis platelets) per year (range 3500–3800 in recent years).

#### Question 10b

It is not possible at this time to subgroup by age, maybe in the future (with help with EPIC/HCLL).

# Subject 2. Transfusion Indications for paediatric and neonatal patients

#### **Question 1b**

Yes, some recommendations (neonatal, platelet).

- (a) Recommendations for neonatal red cell transfusions differ from adults.
- (b) C17 recommendations for platelets threshold higher for bleeding, fever.

## Question 2b

Yes.

- (a) Red cells similar to adults.
- (b) Platelets more similar to C17 guidelines.

#### Question 3b

Yes.

(a) This is what we have in EPIC right now.

## Red cells

Acute or surgical blood loss

To maintain Hb. Level >70 g/l

To maintain Hb. Level >100 g/l

To maintain Hb. Level > \_\_\_\_\_specify

Chronic transfusion programme

Exchange transfusion

ЕСМО

Dialysis prime

#### Apheresis prime

#### Plasma

Bleeding with INR >1.5 or APTT >1.5 $\times$  top of age-related reference

Preinvasive procedure with INR >1.5 or APTT >1.5 $\times$  top of age-related reference range

Replacement therapy when specific factor concentrate not available Therapeutic plasma exchange

Massive haemorrhage/transfusion

Urgent reversal of warfarin if prothrombin complex not available **Platelets** 

PLT.ct <10 000/µl

PLT.ct <20 000/µl stable infant

PLT.ct <20 000/µl (sepsis, bleeding, other considerations)

PLT.ct <50 000/µl unstable infant

PLT.ct <50 000/µl actively bleeding

PLT.ct <50 000/µl pre invasive procedure

PLT.ct <50 000/µl pre major surgery

PLT. Ct <100 000/µl CNS bleeding

PLT. Ct <100 000/µl CNS surgery

## **Question 4b**

Yes, have to select in EPIC.

## Question 5

No.

#### Ouestion 6b

Yes.

- (a) Over 4 months, similar to adults (as per AABB paediatric transfusion handbook).
- (b) Neonates as per AABB paediatric transfusion handbook, except we use Hb 140 instead of 150 for congenital cyanotic heart disease.

## **Question 7b**

Yes.

Guidelines for platelet transfusion in neonates:

- 1 Stable infant, platelet count  $<20 \times 10^9/l$ .
- 2 Unstable infant, platelet count  $30-50 \times 10^9/l$ .
- 3 Infant with active bleeding, or invasive procedure, platelet count  $<50 \times 10^9/l$ .

Prophylactic platelet transfusion (for hem/onc patients over 4 months of age):

Platelet count  $10 \times 10^9$ /l:

• All patients.

Platelet count  $20 \times 10^9/l$ :

- Patients with fever, sepsis, DIC, bleeding, coagulopathy, splenomegaly, AML M3, brain tumour.
- Bone marrow aspirate and biopsy.

• Insertion of PICC line (peripherally inserted central

Platelet count  $50 \times 10^9/l$ :

- Lumbar puncture.
- · Major surgery.
- Hurler's syndrome post-BMT.
- For placement of central venous catheters other than PICC lines, consult IGT (image-guided therapy) in advance.

Platelet count  $100 \times 10^9$ /l:

• CNS bleeding, CNS surgery, invasive procedure in patients with D.I.C.

#### **Ouestion 8b**

Yes.

See the indications in Epic in 3.

# Subject 3. Product manipulations for paediatric and neonatal patients

## Question 1b

Yes.

Irradiated cellular blood products (red cells and platelets):

The following patient populations receive irradiated cellular blood components at Sick Kids:

- Infants up to 6 months of age
- Primary immune deficiencies (SCID, Wiscott-Aldrich syndrome, Di George's syndrome, 22q deletion etc).
- · Cardiac patients with a diagnosis of truncus arteriosus or interrupted aortic arch, until test result for 22q deletion is known.
- · Patients receiving or have received: purine analog (e.g. fludarabine, cladribine, pentostatin/deoxycoformicin), alemtuzumab (anti-CD 52), antithymocyte globulin (ATG).
- Leukaemia, lymphoma. severe aplastic anaemia.
- · Solid tumours if receiving intense chemotherapy such as those listed above.
- · Bone marrow transplant recipients (and donors during bone marrow harvest).
- · Heart transplant and lung transplant recipients (ATG for conditioning).
- · Components from blood relatives (directed dona-
- HLA-matched single-donor apheresis platelets.

# Ouestion 2b

Yes.

(a) All RBCs are prestorage leucoreduced by blood centre.

#### Question 3b

Yes, but rarely do.

(a) Patients with anti-IgA and those still reacting to plasma-reduced red cells.

#### Ouestion 4a

No.

## **Question** 5a

No.

#### **Question 6b**

Yes.

- (b) K neg for all female patients.
- (c) Rh and K for sickle cell and WAIHA.
- (d) Fya, Jka, Jkb, S for SCD who have already made clinically significant antibodies.

#### Ouestion 7

No.

#### Question 8a

Yes.

Dedicated units for neonates under 1000 g birthweight. Split units for patients >4 months who only gets part of unit at a time (e.g. to prevent alloimmunization prekidney transplant) (give half a unit now, half a unit at a subsequent date).

Our routine platelet product is pooled buffy coat (4 donors) for children. We try to give part of an apheresis platelet to neonates and to small infants who only need 100 ml or less of platelets.

#### **Question 9**

Yes, microtubes.

### Question 10b

Yes

- (a) Fresher units for cardiac surgery
- (b) <7 days for infants <3 months
- (c) <7 days if possible for infants 3–6 months old, otherwise <14 days
- (d) <14 days for 6 months to 4 years
- (e) <28 days for patients over 4 years
- (f) Neonatal top-up transfusions any age
- (g) Neonatal exchange transfusion <7 days
- (h) Haemoglobinopathy patients on chronic transfusion programme phenotyped units <21 days ordered for scheduled patients, but will use up to 35 days
- (i) Sickle cell red cell exchange <21 days

# Subject 4. Blood product dosing

## Question 1

Peds: 10–15 ml/kg. Neonates: 15–20 ml/kg.

## Question 2

Peds: 5-10 ml/kg.

Neonates: 10-15 ml/kg.

# Question 3

10-15 ml/kg.

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