

Neonatal transfusion practices

Cases on TML testing and Blood administration

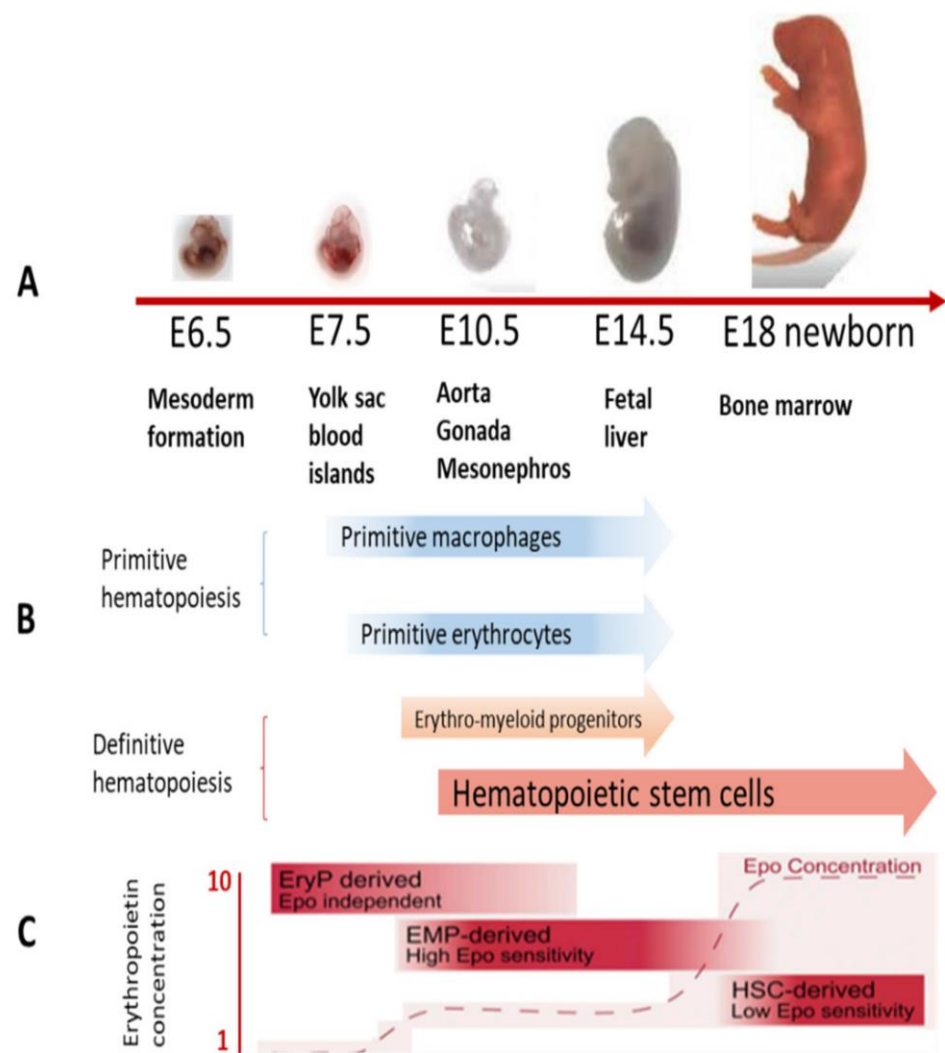
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Objectives

- Understand the developmental differences in hemoglobin and coagulation parameter values between neonates, children and adults
- Recognize the transfusion thresholds for red cells and platelets in neonates
- Understand the differences in pre-transfusion testing for neonates
- Recognize the specifics of blood product administration and modification for neonates

Developmental Hematology 101

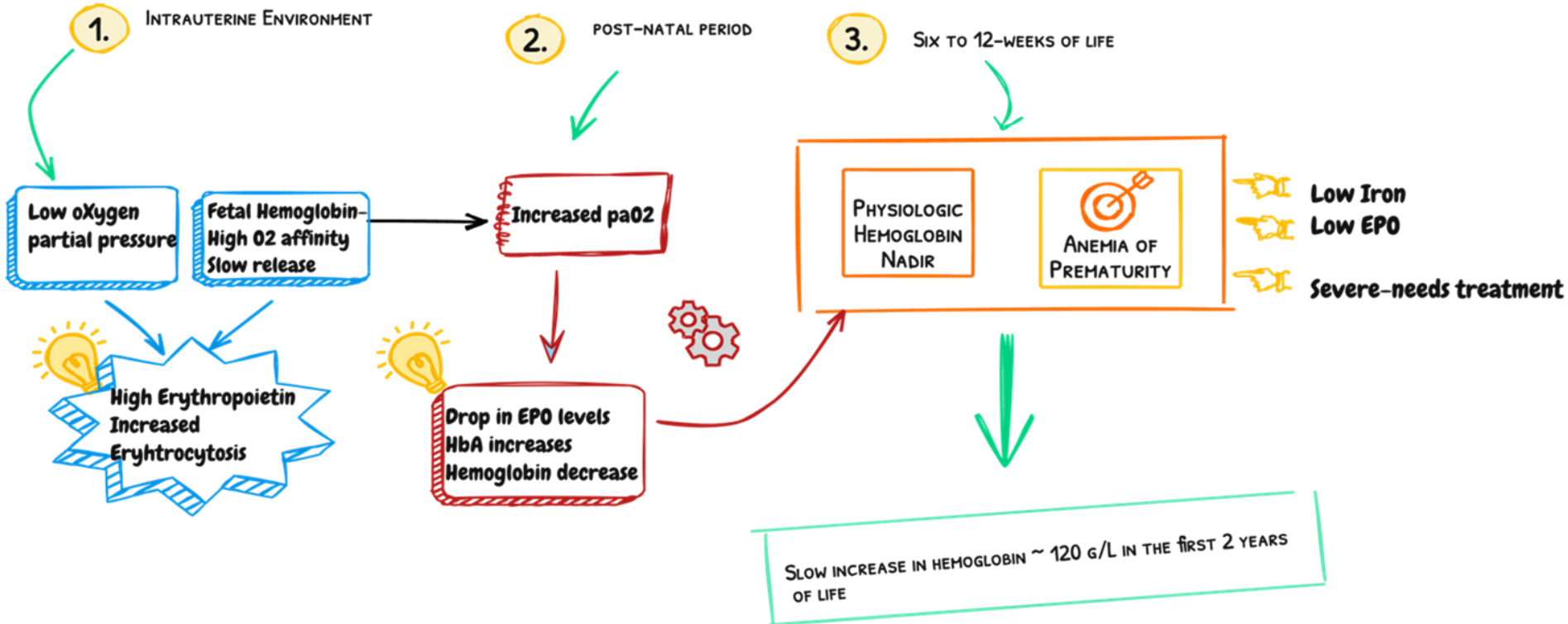


Case 1:

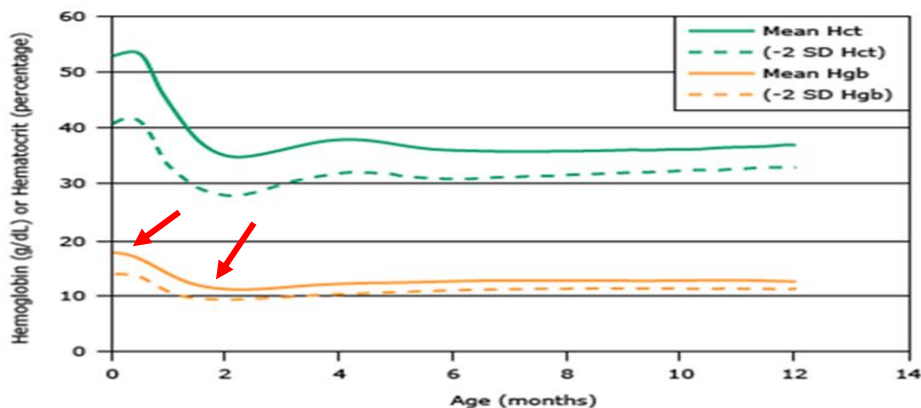
A 2-month old male born at 30-weeks gestation has a hemoglobin of 85 g/L today whereas it was 170 g/L at birth.

What is the most likely reason for the change in his hemoglobin level?

Developmental changes in Hemoglobin



Normal values for hematocrit and hemoglobin during the first year of life in healthy term infants



Hct: hematocrit; SD: standard deviation; Hgb: hemoglobin.

Data from:

1. Jopling J, Henry E, Wiedmeier SE, et al. Reference ranges for hematocrit and blood hemoglobin concentration during the neonatal period: data from a multihospital health care system. *Pediatrics* 2009; 123:e333.
2. Oski FA, Naiman JL. Hematologic problems in the newborn, 2nd ed, WB Saunders, Philadelphia 1972; p.13.
3. Saarinen UM, Siimes MA. Developmental changes in red blood cell counts and indices of infants after exclusion of iron deficiency by laboratory criteria and continuous iron supplementation. *J Pediatr* 1978; 92:412.

Immediately after birth

Low
vitamin K

Immature
Liver cells

Low vitamin
K-dependent
factors

Increased
Fibrinolysis

Low
procoagulant
factors

Factors 2, 7, 9, 10

High PAI-1,
plasminogen

Factors 5, 11, 12

Protein C, S

BALANCED COAGULOPATHY
HIGHER PTT 'NORMAL' VS.
ADULTS
SIMILAR TO ADULT LEVELS
IN 3-12 MONTHS OF LIFE
PREMATURITY INCREASES THE
DIFFERENCE

Normal looks different

Normal values for hemoglobin, INR and PTT are different for neonates and infants

Age is important

Differences in coagulation parameter compared to adults increase with decreasing GA.

There is a 'balance'

Neonates have a 'balanced' coagulopathy. How do we affect it by giving adult derived products?

Numbers don't tell the truth-entirely

Numbers alone do NOT predict bleeding risk. Even less so compared to children and adults

WHY DOES THIS MATTER?

They grow fast

The timeline during which the values become similar to adults is very dynamic and usually in the first year of life

So what do I do with this info?

Deciding to transfuse a neonate? Look at age-based reference ranges. Consider full clinical picture

Transfusion Thresholds

A stylized illustration of a medical form or checklist titled "Transfusion Thresholds". The form has a light blue background and a thick black border. At the top is a large empty rectangular box. Below it are four rows, each starting with a small square checkbox. The first three checkboxes are yellow with a black checkmark, and the fourth is red with a black "X". To the right of each checkbox is a horizontal line for text. A large orange circle with a black outline and a black "i" inside is positioned at the bottom right of the form.

Case 2:

You receive request for RBC transfusion in a 2-day old male in NICU. He was born at 28 weeks gestation. His hemoglobin was 140 at delivery and is 100 today. He is not bleeding.

Is this request appropriate?

How is that determined?

What does that mean for RBC transfusion in neonates



IMPORTANT CLINICAL TRIALS FOR ANEMIA & RBC TRANSFUSION

THE PINT TRIAL (2006)

RESTRICTIVE VERSUS LIBERAL
TRANSFUSION STRATEGY
(N = 451)



THE ETTNO TRIAL (2020)

EUROPEAN MULTICENTER
RANDOMIZED CLINICAL
TRIAL (N = 1083)
RBC TRANSFUSION
THRESHOLDS



THE TOP TRIAL (2020)

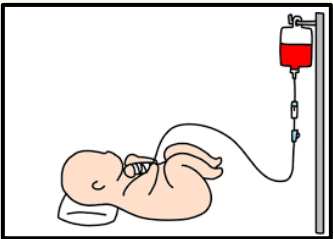
NORTH AMERICAN
MULTICENTER RANDOMIZED
CLINICAL TRIAL (N = 2561)
RBC TRANSFUSION
THRESHOLDS



THE PENUT TRIAL (2020)

MULTICENTER RANDOMIZED
TRIAL COMPARING EFFECT
OF ESA'S VERSUS PLACEBO
FOR NEURODEVELOPMENTAL
OUTCOMES (N = 941)

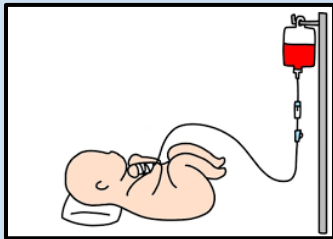
Red cell transfusion thresholds for non-bleeding neonates



Days 1-7

Cardiorespiratory Support.

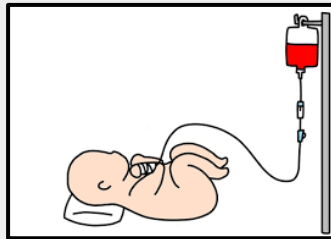
Yes	115 g/L	No	100 g/L
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Days 8-14

Cardiorespiratory Support.

Yes	100 g/L	No	85 g/L
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Day 15 onwards

Cardiorespiratory Support.

Yes	85 g/L	No	75 g/L
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Case 3:

A 5-day-old infant born at 30-weeks gestation had a head US done today. It showed a grade 1 hemorrhage in CNS ventricles which is stable compared to last imaging. His CBC is shown below.

Is platelet transfusion an appropriate next step for him?

WBC count	$10 \times 10^9/L$
Hemoglobin	120 g/L
Platelets	$68 \times 10^11/L$

Research favors a restrictive transfusion threshold for platelets at 25 in non-bleeding neonates and infants

Low grade prematurity related CNS bleeds alone should not prompt platelet transfusion unless count is < 25-50

Canadian Multi-center trial

N = 251

Intervention: Platelet transfusion for count below 150 vs. counts below 50.
Outcome variables: Risk or worsening of prematurity related ICH
Results: No significant difference

1993

The PLANet-2 trial

N = 660

Intervention: Restrictive (25) versus liberal (50) transfusion threshold in non-bleeding infants.

Outcome variables: Mortality, bleeding severity, LOS

Results: Higher odds of death or major hemorrhage in liberal threshold group.

2019

Post Hoc analysis of PLANet-2 trial for heterogeneity of effect

Identify preterm infants at higher bleeding risk in the trial cohort
Reduction in bleeding risk observed across all groups.

High Bleeding Risk: Previous major bleed, allocated to liberal threshold, low GA

2020

CLINICAL STUDIES ON THROMBOCYTOPENIA AND TRANSFUSION THRESHOLDS IN PREMATURE INFANTS

Post Hoc analysis of PENUT trial: Platelet transfusions effect on neuro-developmental outcomes (N = 891)

Infants exposed to one or more platelet transfusions during trial had higher likelihood of impaired neurodevelopment and death.

2024

Pre-transfusion testing conundrums

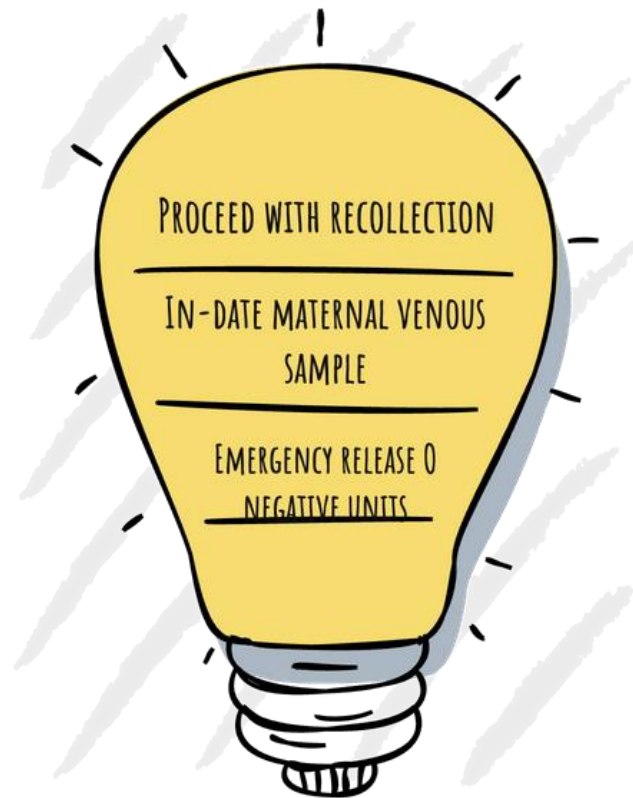
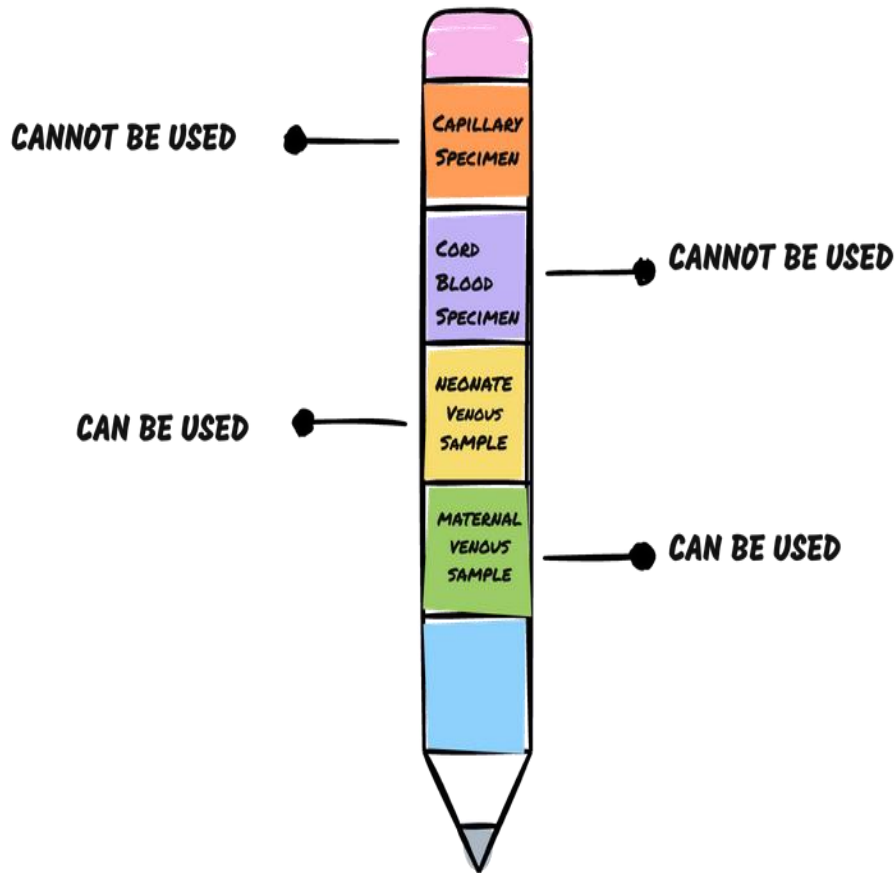


Case 4:

A red cell transfusion is ordered for an NICU infant at 5-days of life. His requisition for pre-transfusion type and screen is not signed by any clinical care providers. When you request a recollection, the nurse says 'well, he is an extremely hard draw and really needs blood, can I just come down and sign the requisition?'

How do you approach this scenario

WHAT CAN I USE FOR NEONATAL GROUP, SCREEN AND CROSSMATCH



POTENTIAL SOLUTIONS

Case 5:

A new technologist gets an order for blood group typing in a neonate. Following are the results

Anti-A	Anti-B	Anti-A,B	A1 cells	B cells
0	0	0	0	0

What is the blood type for the neonate?

ANTI-A AND ANTI-B ANTIBODIES IN NEONATES

1

**NATURALLY OCCURRING
ISOHEAMAGGLUTININS**

Require exposure to A and B
antigens in food and gut
flora

2

**ANTIBODIES IN
NEONATAL PLASMA ARE
OF MATERNAL
ORIGIN**

IgG anti-A/B, antibodies
against minor RBC antigens
or anti-D via recent
maternal RhIg

3

**INFANTS DE NOVO
ISOHEAMAGGLUTININS**

Develop by 3-6 months of
life

4

**PRE-TRANSFUSION
TESTING IN FIRST 4
MONTHS**

May only reveal forward
type and no reverse type

ABO specific RhD compatible

CAN be given to neonates.



Blood group O RhD compatible

Current common practice in
most North American
Institutions

Caveat with ABO specific RBCs

ABO compatibility with
mother. As maternal IgG anti-A/B
antibodies can be detected in
neonatal plasma

Blood Administration specifics



Case 6:

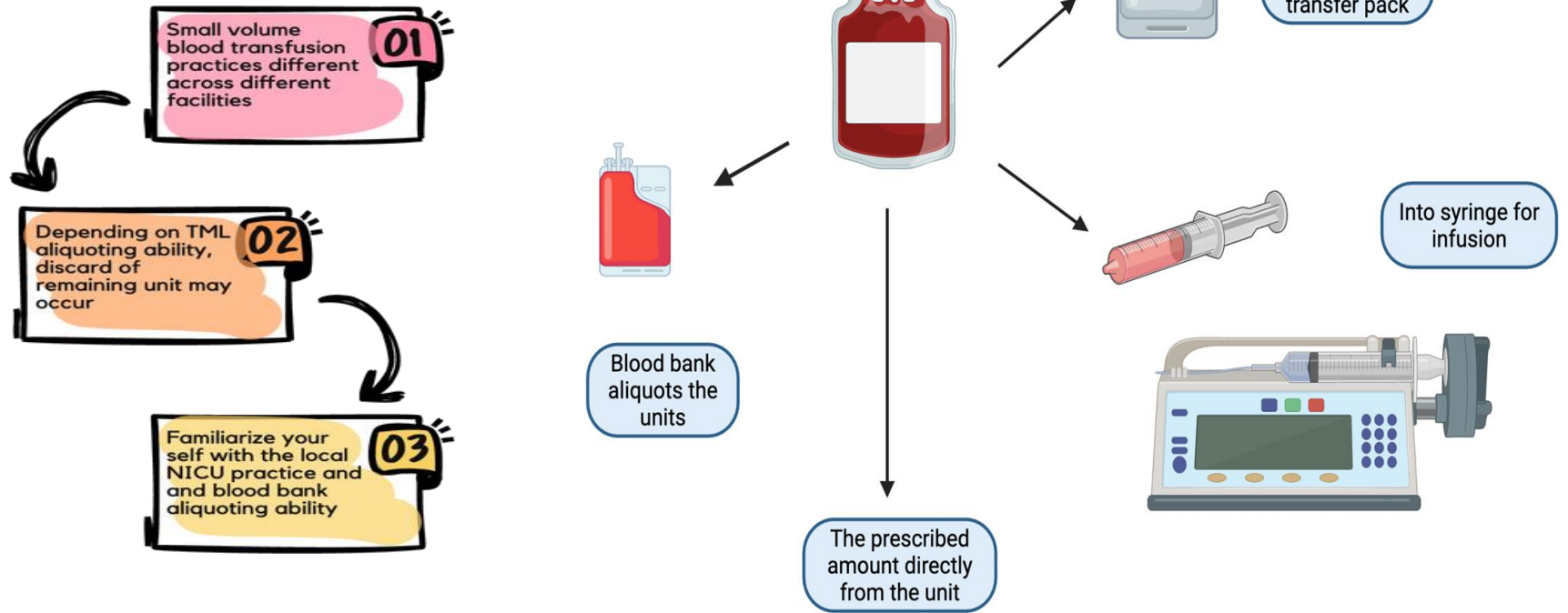
A nurse is preparing to infuse 22 ml of red cells to a 1500 gram neonate in NICU.

Physician order recommends to give over 4 hours.

The TML sends a 300 ml unit of red cells up to NICU

How to approach this scenario?

SMALL VOLUME BLOOD TRANSFUSIONS IN INFANTS



Blood Product Modification

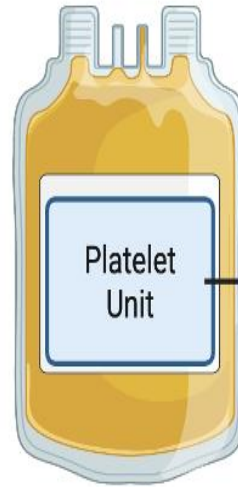
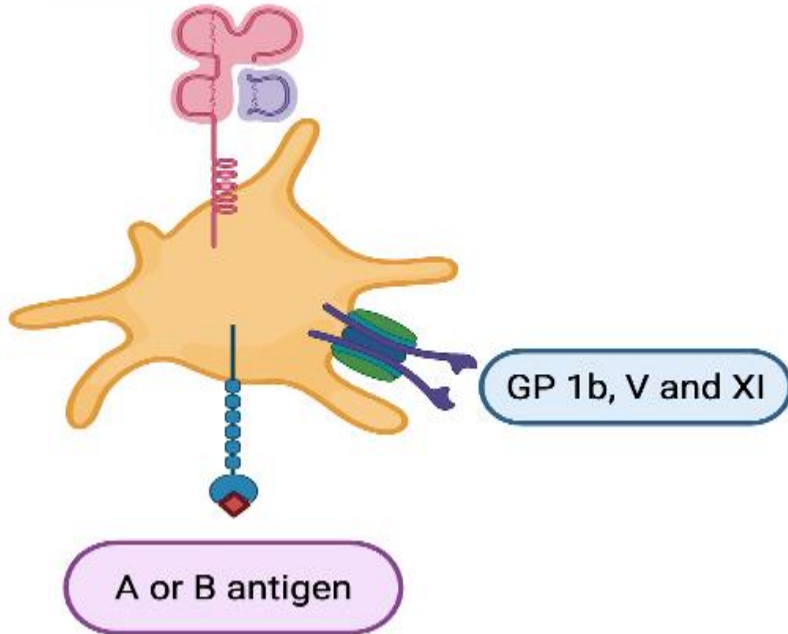
Case 7:

You receive a request for platelet transfusion for a neonate who is A positive. The blood bank only has O positive platelets at time.

Does this matter?

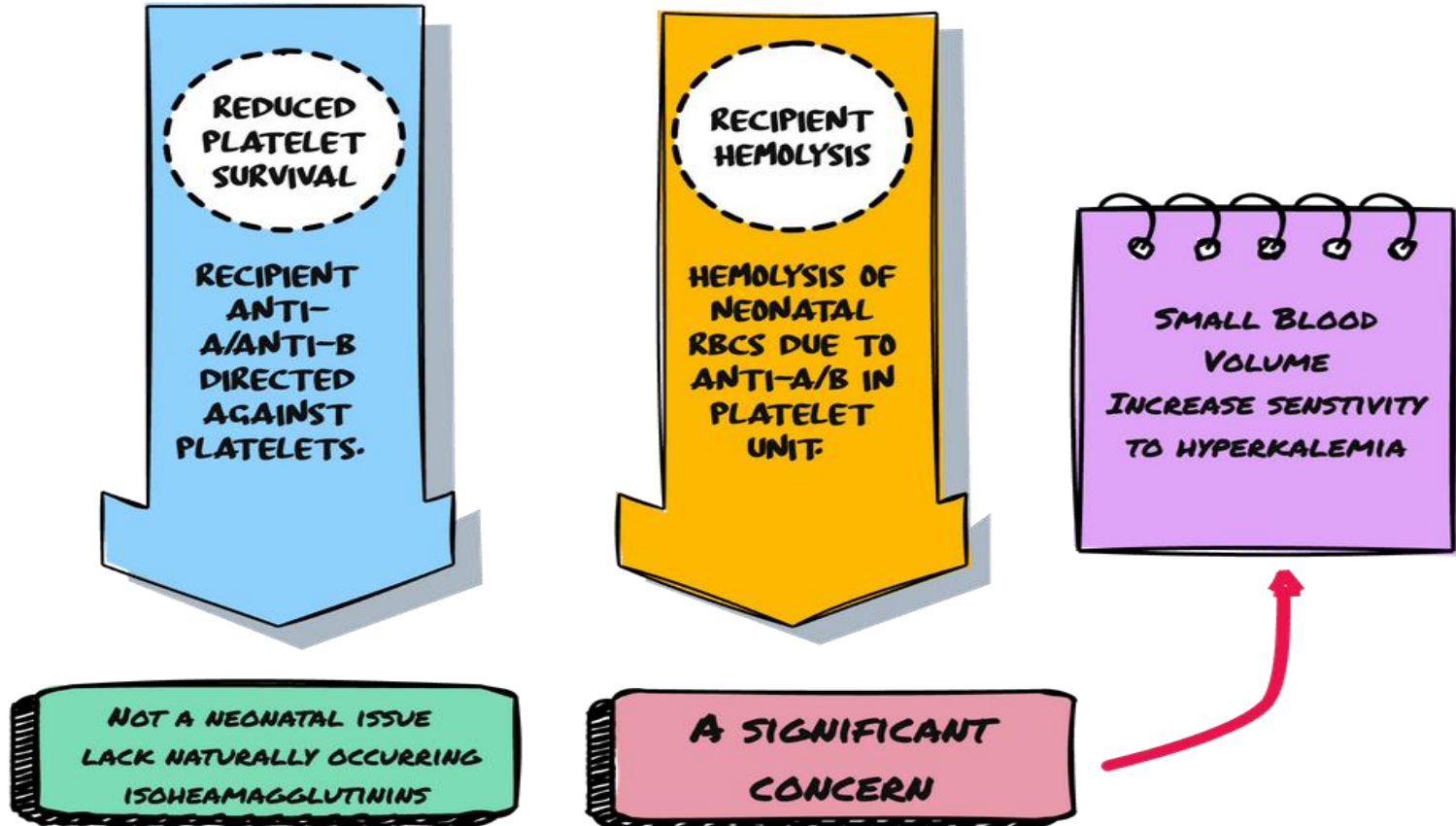
How do you approach this situation?

HLA Class 1

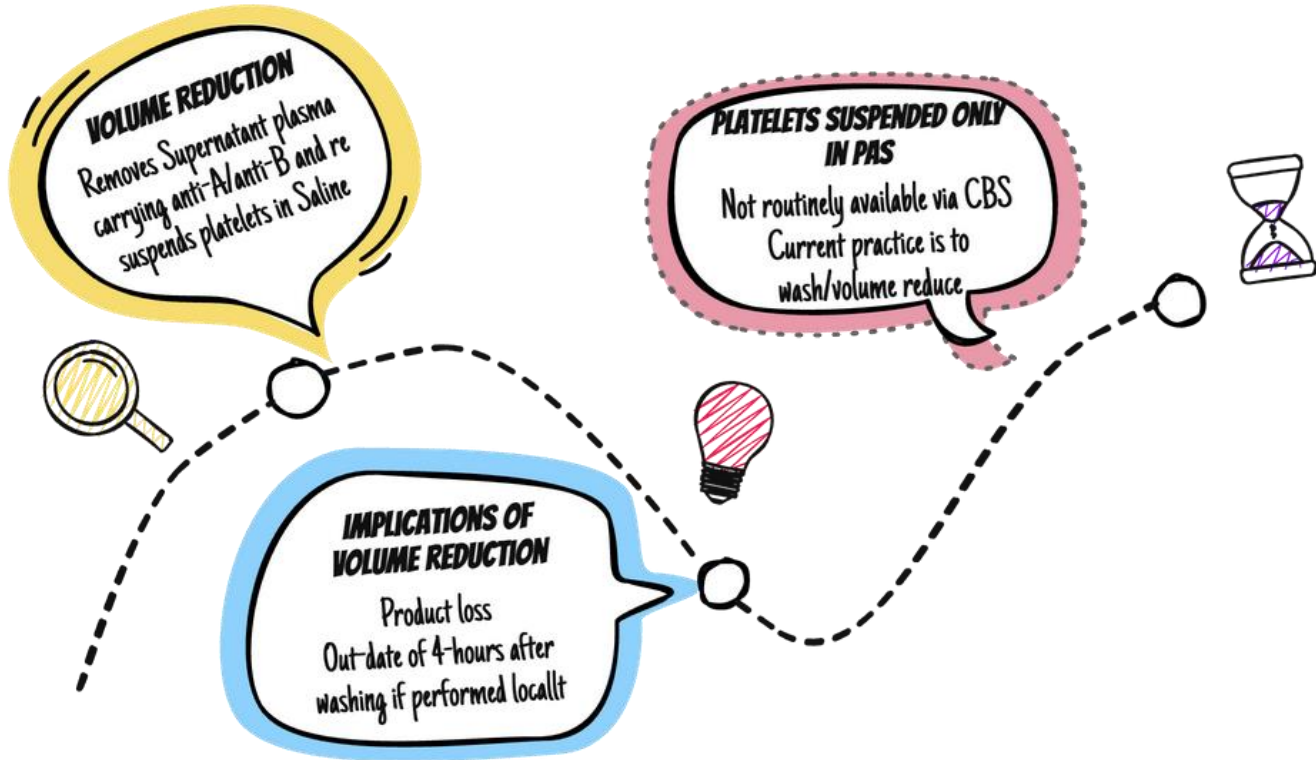


Donor plasma
Platelet additive solution
Donor anti-A/B antibodies
Donor RBC fragments

Implications of ABO incompatible platelets for neonates



! HOW TO ISSUE ABO INCOMPATIBLE PLATELETS TO NEONATES



Case 8:

You receive a call from RRPL in Saskatoon that Baby Yoda's newborn screen is positive for Severe Combined Immunodeficiency as she has low TRECS.



Why is this information important for the blood bank?

What are they?

Circular DNA fragments in plasma produced during T-cell maturation

Why does it matter in TML?

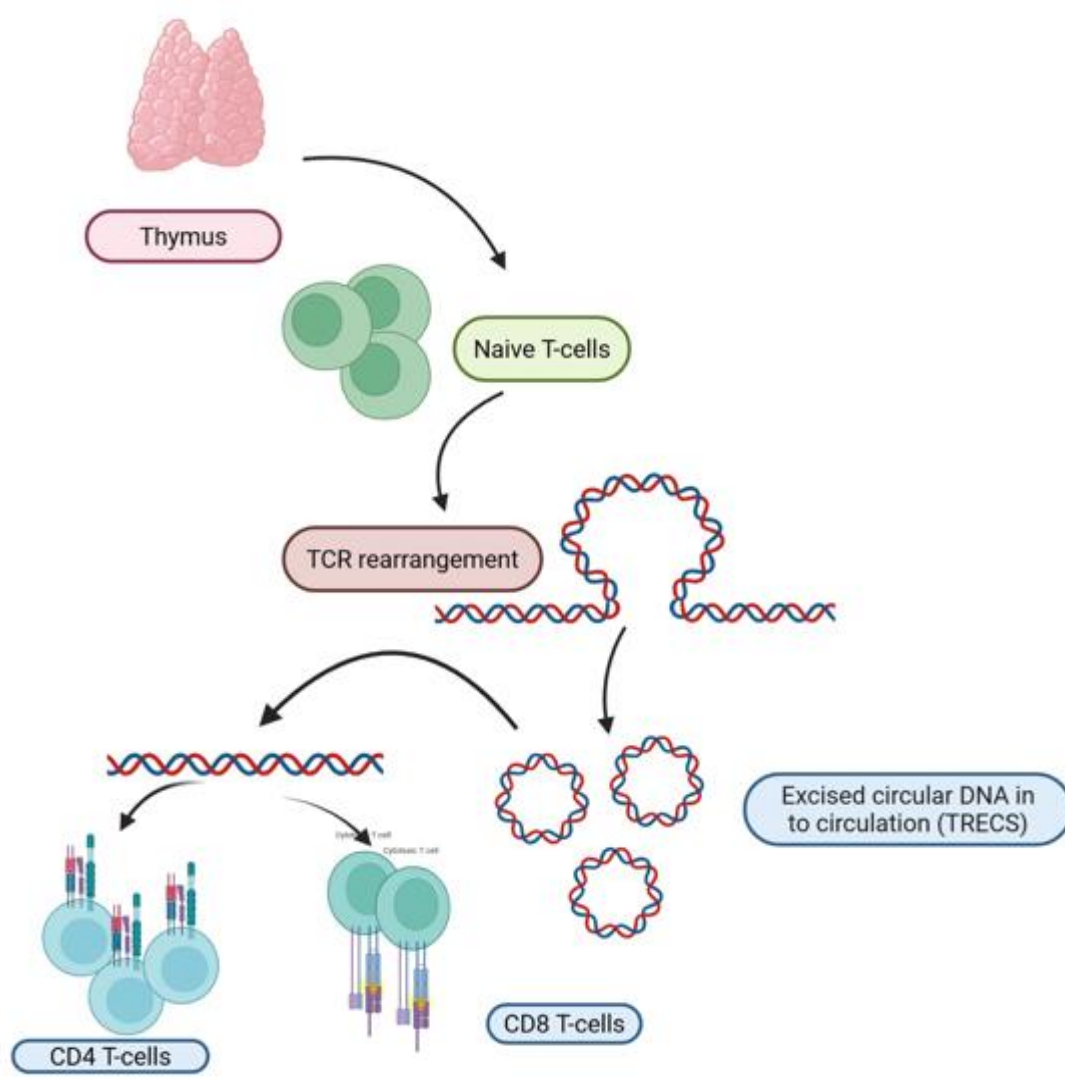
Neonates with SCID are at high risk of developing transfusion associated graft versus host disease

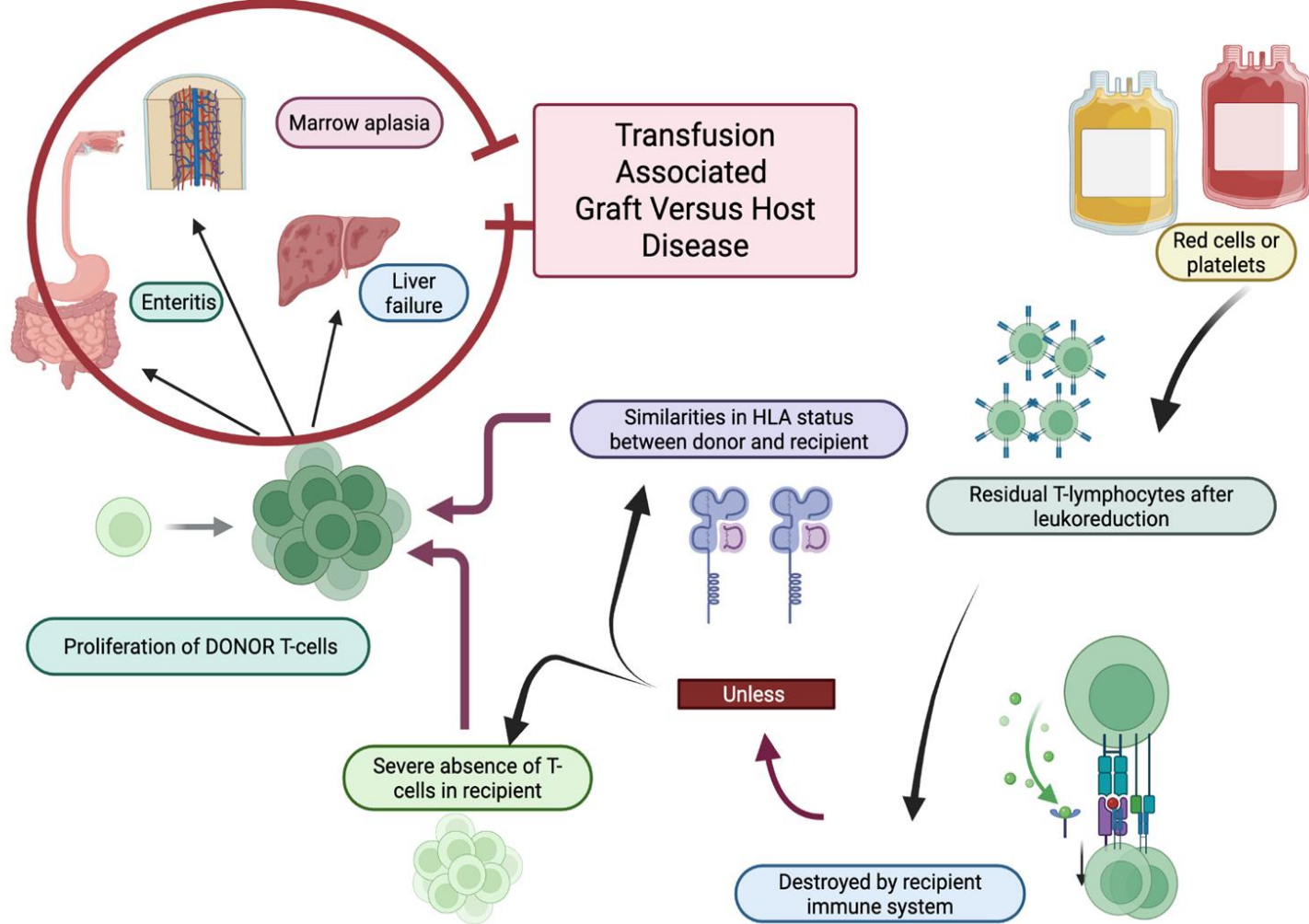
Why do they matter?

Highly sensitive screening tool for severe combined immunodeficiency in neonates

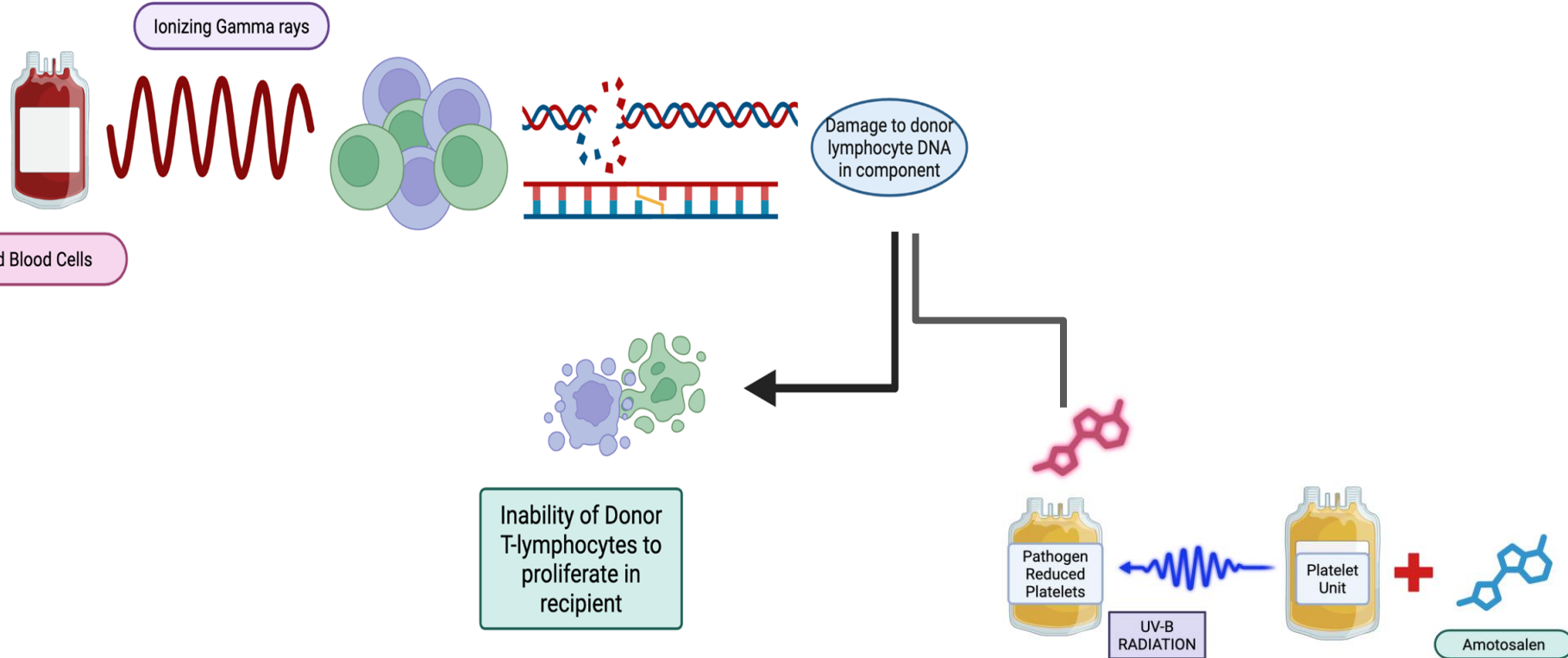
What is the prevention?

Irradiation of blood products

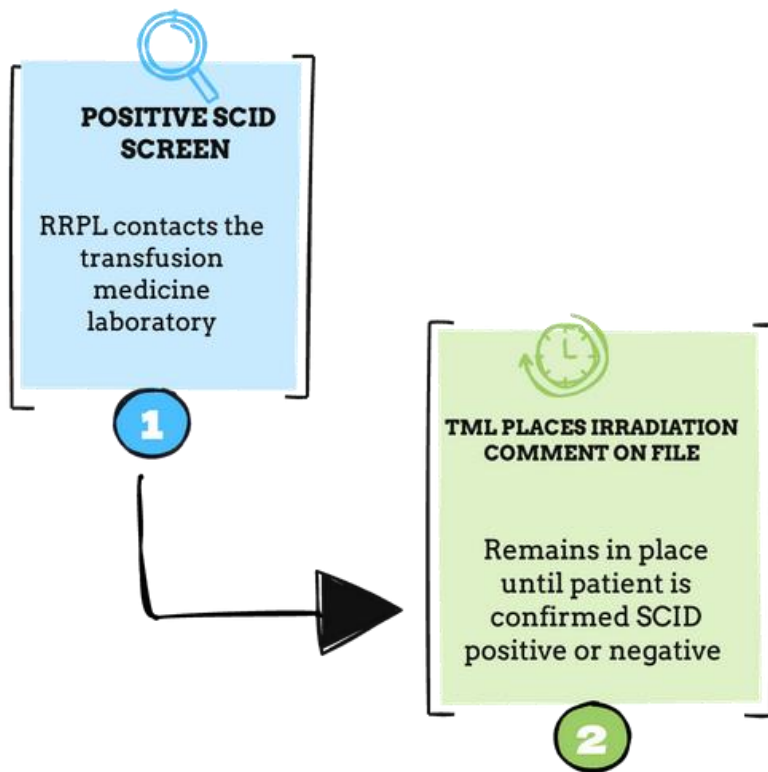




TA-GVHD Prevention strategies



Current Process for neonates screened positive for SCID in SK



Summary

Neonatal transfusion practice differs from adults

