Non-Infectious Transfusion Reactions

DR MOGHADDAM

Transfusion Reactions

ANY

unfavorable consequence is considered a transfusion reaction of blood TX

The risks of transfusion must be weighed against the benefits

Transfusion Reactions

- 1. Acute (<24 hours) Transfusion Reactions
- 2. Immunologic
 - Hemolytic; Febrile-non hemolytic; Allergic; Anaphylactic; Transfusion Reaction of Acute Lung injury(TRALI)
- 3. Nonimmunologic
 - Hemolytic (Physical or Chemical destruction of RBC); Circulatory overload; Air embolus; Hypocalcemia; Hypothermia
- 4. Delayed (>24 Hours) Transfusion Reaction
- 5. Immunologic
 - Hemolytic ; Graft vs. Host Disease; Posttransfusion Purpura
- 6. Nonimmunologic
 - Iron Overload
- 7. Infectious Complications of Blood Transfusion

Acute (<24 hours) Transfusion Reactions

- 1. Immunologic
 - Hemolytic
 - Febrile-non hemolytic
 - Allergic; Anaphylactic
 - Transfusion Reaction of Acute Lung injury(TRALI)
- 2. Nonimmunologic
 - Volume overload
 - Hemolytic (Physical or Chemical destruction of RBC)
 Air embolus
 - Hypocalcaemia; Hypothermia

Acute Hemolytic Transfusion Reaction

- Associated with Intravascular Hemolysis
- Etiology: Antibodies that activate complements in the vasculature: ABO antibodies are predominant / not the only ones.
- Prevention: Give ABO compatible blood.

- May also occur due to ABO incompatible plasma in platelet products
- Very rare; less than 20 case reports, all involving group O platelets
- Usually occurs in group A patients or those with anti-A titers greater than 1:1000
- Can prevent by removing plasma from platelets, or limiting number of incompatible group O platelets in a 24 hour period

Intravascular Hemolysis

Characteristics

- Within minutes
- IgM &/or IgG antibody
- Complement activation
- Release of histamine and serotonin

Signs may include:

- Pain along infusion site
- Shock
- Abnormal bleeding/DIC/ Hemoglobinemia/uria
- Release of cytokines: fever, hypotension
- Renal failure/ Oliguria, may progress to...anuria

Febrile non-hemolytic TX Reactions

- An INCREASE in temperature of 1°C during infusion of blood component
 - Usually "mild & benign" = not life threatening
 - Can have more severe symptoms, not usually
- Non-hemolytic
- Incidence of 0.1% of RBC transfusions
- 0.1-1.0% of platelet transfusions
- Cause: Recipient antibodies to donor WBCs & Cytokines in the transfused blood component.

Febrile Transfusion Reactions

Seen in...

- Multiply transfused patients
- Multiple pregnancies
- Previously transplanted

Must rule out...

- Hemolytic transfusion reaction
- Bacterial contamination of unit

Prevention

• Leukocyte reduction (pre-storage reduction may be more effective than post-storage reduction) or plasma removal is also helpful.

Acute Transfusion Reactions

Immunologic

Allergic (Urticarial-Hives)

- Etiology: Form of cutaneous hypersensitivity triggered by recipient antibodies directed against:
 - Donor plasma proteins or
 - Other allergens (food, medicines) in donor plasma
- Begins within minutes of infusion
- Characterized by rash and/or hives and itching .
- Common (1 per 2000 transfusions)
- Usually involves release of histamine.

Allergic (Urticarial) Reactions

- MUST be sure that the only reaction is the development of urticaria
- Must rule out more severe symptoms that could lead to anaphylaxis:
 - angioneurotic edema
 - laryngeal edema
 - bronchial asthma
- Prevention:
- Can pre-treat recipient with anti-histamines before transfusion.

Anaphylaxis

- Life threatening!!
- Etiology:
 - Recipient is IgA deficient & has anti-IgA in serum
 - Recipient anti-IgA can react to even small amounts of donor IgA in the plasma in any blood component
 - Idiopathic & Haptoglobin deficiency
- Reaction may occur within minutes : <u>Onset of symptoms is</u> <u>SUDDEN</u>
- **Prevention:** Wash cellular components or blood products from IgA deficients

Anaphylaxis

Symptoms

- Burning sensation at infusion site
- Coughing, difficulty in breathing, and bronchospasms can lead to cyanosis
- Nausea, vomiting, severe abdominal cramps, diarrhea
- Hypotension which can lead to shock, loss of consciousness, & death

• MUST STOP TX IMMEDIATELY

Acute Transfusion Reactions

Immunologic

TX Reaction of Acute Lung Iinjury Etiology:

- Acute onset of hypoxemia and pulmonary edema on CX-RAY within 6 hrs of TX without evidence of cardiac failure.
- Mechanism's
- Primary Suspect: Donor antibodies to recipient WBCs
- Another cause: Biologically active lipids in the lungs causing edema

Transfusion Reaction of Acute Lung Injury(TRALI)

Symptoms

- Chills, fever, cough, cyanosis, hypotension, increased difficulty breathing
- Prevention:
- For recipients : give male products
- For donors: watch/defer.

Acute Transfusion Reactions

NONimmunologic

Circulatory Overload

• Etiology: Rapid increases in blood volume to patient .Risk factors: compromised cardiovascular function, current volume overload, small intravascular volume (elderly, young children), severe chronic anemia.

Signs and Symptoms

- Dyspnea, cyanosis, severe headaches, hypertension or CHF (congestive heart failure). Chest Xray: pulmonary edema, distended pulmonary artery, cardiomegaly
- Laboratory: elevated B-natriuretic peptide (BNP) is 81% sensitive and 89% specific
- Prevention: Slow Tx.
- Treatment: Stop infusion and place patient in sitting position.

Physically or Chemically Induced Red Cell Destruction

Etiology:

• Destruction of red blood cells in the collection bag and infusion of free hemoglobin, etc.

Improper temperatures: High or Low

 Microwave blood bag, malfunctioning blood warmer or water bath, inadvertent freezing of blood. Physically or Chemically Induced Red Cell Destruction

Osmotic Hemolysis

• Addition of drugs or hypotonic solutions (5% dextrose, deionized water, etc.) to transfusion.

Mechanical Hemolysis

- Caused by rollers in blood pump
- Pressure infusion pumps
- Small bore needles
- Prevention: Adherence to procedures for all aspects of procuring, processing, issuing and administering red blood cell transfusions.

Hypocalcemia

- Excess citrate: When infused at rate >100 mL/minute or individuals with impaired liver function:
 - Citrate is broken down by liver.
- Seen more in pediatric and elderly patients
- Signs and Symptoms: Facial tingling, nausea, vomiting.
- Prevention: Slowing or discontinuing infusion.

Hypothermia

- Etiology: Drop in core body temperature due to rapid infusion of large volumes of cold blood.
- Symptoms: Decreased body temperature and ventricular arrhythmias.
- Seen in small infants or massive transfusion
- **Prevention:** Reduce rate of infusion or use blood warmers.

Air Embolism

- Etiology: If blood in an open system is infused under pressure or if air enters the system while container or blood administration sets are being changed.
- Treatment: Place patient on left side with head down to displace air bubble from pulmonic valve.

Delayed (>24 Hours) Transfusion Reaction -

- 1. Immunologic
 - Hemolytic
 - Graft vs Host Disease
 - Posttransfusion Purpura
- 2. Nonimmunologic
 - Iron Overload

Delayed Transfusion Reactions

Immunologic

Delayed Hemolytic Transfusion Reaction (Red blood cell alloimmunization)

- Onset within days (>24 hours)
- Associated with Extravascular Hemolysis
- Etiology: Antibodies that usually do NOT activate Complements : Rh, Kell, etc.
- Prevention: Give antigen negative blood.

Extravascular Hemolysis

Characteristics

- Reaction within days
- Antibody attaches to RBC: RBC destroyed in spleen or liver, etc.
- Commonly IgG
- May or may not activate Complement

Signs may include:

- No release of free Hgb, or enzymes into circulation
- May be immediate (hours) or delayed (days)
- Bilirubinemia or bilirubinuria

Extravascular Hemolysis

Signs & Symptoms continued...

- 1. Fever or fever & chills
- 2. Jaundice
- 3. Unexpected anemia
 - Some may present as an ABSENCE of an anticipated increase in Hemoglobin and hematocrit.

Delayed Transfusion Reaction

Immunologic

Graft vs Host Disease (GVHD)

- Etiology:
- Donor CD8+ T-Lymphocytes attack recipient (host) tissues. Very rare in blood stored 4+ days due to WBC *inactivation*
- Groups at risk:
 - Immunocompromised patients (Cancer, fetus, neonatal, bone marrow transplant).
- Signs:
- Fever, dermatitis, or erythroderma, hepatitis, diarrhea, pancytopenia, etc.
- Prevention:
- Irradiation of blood products.

Delayed Transfusion Reaction Immunolgic

Post-transfusion Purpura

- Etiology:
- Antibodies to platelet antigens (HP1a) causes abrupt onset of severe thrombocytopenia (platelet count <10,000/µl) 5-10 days following transfusion. Usually affects multiparous women.
- Signs:
- Purpura, bleeding, fall in platelet count
- treatment:
- IVIG, plasmapheresis or corticosteroids; platelet transfusions usually NOT recommended

Delayed Transfusion Reaction

NONimmunolgic

Iron Overload

- Etiology:
- Excess iron resulting from chronically transfused patients such as hemoglobinopathies, chronic renal failure, etc.
- Signs:
- Muscle weakness, fatigue, weight loss, mild jaundice, anemia, etc.
- Treatment:
- Infusion of deferoxamine an iron chelating agent has been useful.

1. Acute (<24 hours) Transfusion Reactions - Immunologic

- Hemolytic; Febrile-non hemolytic; Allergic; Anaphylactic; Transfusion Reaction of Acute Lung injury(TRALI)
- 2. Acute Transfusion Reactions Nonimmunologic
 - Volume overload; Hemolytic (Physical or Chemical destruction of RBC); Air embolus; Hypocalcemia; Hypothermia
- 3. Delayed (>24 Hours) Transfusion Reaction Immunologic
 - Hemolytic ; Graft vs. Host Disease; Posttransfusion Purpura
- 4. Delayed Transfusion Reactions Nonimmunologic
 - Iron Overload
- 5. Infectious Complications of Blood Transfusion

Infectious Complications of

Blood Transfusion (Viral is rare)

Infectious Complication of Blood Transfusion

Bacterial Contamination

- Etiology:
- At time of collection: either from the donor or the venipuncture site.
 - During component preparation, etc.
- Usually involves endotoxins
 - Staph, Pseudomonas, E.coli, Yersinia

Bacterial Contamination

• Components:

- Most often from platelet components (room temp). Red cell units will look dark.
- Symptoms:
- Rapid onset
 - Fever, hypotension, shaking chills, muscle pain
 - Vomiting, abdominal cramps, bloody diarrhea, hemoglobinuria, shock, renal failure, & DIC.

Bacterial Contamination

- **Transfusion must be stopped immediately**
- Gram stain & blood cultures should be done on the unit, patient and all infusion sets .
- Broad-spectrum antibiotics should be given immediately intravenously
- Prevention: Maintain standards of donor selection, blood collection and proper maintenance of collected blood components.

Transfusion Reactions

1. Acute (<24 hours) Transfusion Reactions - Immunologic

- Hemolytic; Febrile-non hemolytic; Allergic; Anaphylactic;
 Transfusion Reaction of Acute Lung injury(TRALI)
- 2. Acute Transfusion Reactions Nonimmunologic
 - Volume overload; Hemolytic (Physical or Chemical destruction of RBC); Air embolus; Hypocalcemia; Hypothermia
- 3. Delayed (>24 Hours) Transfusion Reaction Immunologic
 - Hemolytic ; Graft vs. Host Disease; Posttransfusion Purpura
- 4. Delayed Transfusion Reactions Nonimmunologic
 - Iron Overload
- 5. Infectious Complications of Blood Transfusion

Transfusion Reaction Follow-up

Clinical Information Needed:

- Recipient diagnosis
- Medical history of pregnancy &/or transfusion
- Current medications
- Signs & symptoms during transfusion reaction
- How many mL's of RBC's or plasma were transfused?

Clinical Information Needed

- Were rbc's cold or warm when transfused?
- Were red cells infused under pressure?
- What was the size of the needle used?
- Were other solutions given through the IV line at the same time? If so what?
- Were any other drugs given at the time of transfusion? If so, what?
- What were pre- & post- transfusion vital signs?

Transfusion Reaction Follow-up

Post Transfusion Reaction blood samples to be collected from the recipient:

Clotted specimen

 Repeat ABO, Rh, IAT and Crossmatch. Visual check for hemolysis and compare with pre transfusion sample.

• EDTA specimen

• DAT (Direct Antiglobulin Test)

- Clotted specimen
- 1st voided urine specimen post-tx'n

- Collect 5-7 hours post transfusion to check for bilirubin
- Free hemoglobin determination

Transfusion Reaction Workup

CLERICAL CHECKS

- Correct identification of patient, specimen, and transfused unit.
- 2. Agreement of records and history with current results
- 3. Correct labeling of transfused unit

SPECIMEN CHECKS

 Visual inspection of post-transfusion specimen

 Visual inspection of blood bag and lines

Post Transfusion Lab Testing

Direct Antiglobulin Test (DAT)

- Recipient post-tx'n spec.
- Positive: Perform eluate and identify antibody if the pre-TX spec negative.

ABO Grouping and Rh Typing

- Recipient pretransfusion and posttransfusion specimen
- Donor bag.

Post Transfusion Lab Testing

Indirect Antiglobulin Test (IAT)

- Recipient Pre- & post-transfusion reaction specimens
- Pre neg and post pos:
- Identify antibody and compare results of serum panel with eluate panel.

In Summery:

- 1-Allergic: Minor VS Serious
- 2-Febrile: Minor VS Serious

3-Onset<15 Mints, Temp>1C with other symptoms or Temp>39C, BP , Shock, SOA, Rigors, Back/Chest Pain, Bleeding from IV site, Tachy/Arrhythmia, Nausea/Vomiting and Generalized Flushing

In Summery:

1- Stop TX immediately and keep an IV open with 0.9 Saline 2-Contact the clinician 3-Check vital signs every 15 minutes 4-Check labels, forms, and lds 5-Send bags & patient's blood to BB 6-Minor(allergic-febrile non-hemolytic) VS Serious (hemolytic &febrile)

Thank You