

Transfusion Reactions Maher A. Sughayer M.D Blood Bank Director Chairman, Pathology and Lab. Medicine

# **TRANSFUSION REACTIONS**



> Either acute or delayed.

Common: in up to 10% of recipients therefore benefit of Tx should outweigh the risk

> Informed consent





During or shortly after (within 24 hrs) of Tx

The most severe occur early in Tx, so careful monitoring during Tx.

# Hemolytic Reactions



Immune mediated lysis of Tx rbc's: acute or delayed, intra- or extra vascular.

#### **AHTR**



INCOMPATIBLE rbc's given to a pt. with clinically significant preformed ab's to an rbc Ag.

- Most common cause is ABO incompatibility due to clerical error.
- > Within minutes of starting Tx



> Usually by complement-fixing IgM or IgG: intravascular lysis. > By anti-A or anti-B; bind mac(C5-9): intravascular lysis: Hbemia and Hburia. Shock, hypotension, bronchospasm: due to C3a, C5a, and other inflammatory mediators. > Renal ischemia: ATN



Activation of cytokines (IL-1, IL-6, IL-8, TNF-@): FEVER, hypotension, activation of WBC's, coag. : DIC
 Severity is: DOSE- & RATE-RELATED
 MEDICAL EMERGENCY.

# **MANGEMENT OF AIHTR**



When suspected:
Stop Tx immediately
maintain IV by crystalloid or colloid.
maintain BP, pulse
ventilation



# > give diuretic or fluid diuresis > obtain bl. and urine for Tx-reaction investigation

# Perform work-up of Tx-reaction

- > paper-work: Id's, labels: right component to right pt?
- Check plasma for Hbemia, urine for Hburia
- > Perform DAT.
- > Repeat x-match
- Repeat typing ABO-Rh as needed.

# If AIHTR is confirmed



> monitor renal status (BUN,....
 > DIC workup: PT, PTT, fibrinogen, plt.
 > monitor for hemolysis (bili., LDH, hapto, Hbemia)
 > If sepsis is suspected C&S and Rx....



> most AHTR are intravasc. but if Ab doesn't fix complement or only C3: extravascular.

> AHTR may be caused by Ab's in ABOincompatible plt's or IVIG which may be severe.

# NONIMMUNE HEMOLYSIS (NIH)



- If AHTR; but no ABO incompatibility: look for NIH.
- Mechanical: mech. heart valves, extracorporeal cir.
- Somotic lysis: hypotonic saline, drugs, H2O, infused in same line.
- Heating (bl. warmer) or freezing (refrigerator malfunction): may hemolyze rbc's prior to Tx



There may be Hbemia and or Hburia but no serious symptoms, however should be investigated promptly to exclude immune-mediated



# **FEBRILE REACTIONS**



FEVER: may be due to any of: AHTR, bacterial contamination, or most commonly due to the pt's underlying dis. OR FNHTR.

FNHTR: a rise of more than 1 C, may be with chills, not attributable to any cause, occurring during or up to 2 hrs post Tx.



- Less than 1% of all Tx.; while a 2nd FNHTR in same pt. is 1 in 8.
- > Up to 10% in chronically transfused pts.
- Cause: Ab's to transfused WBC's or plt.
- > As a result of Ab-Ag rxn: comp. fragments and IL-1, IL-6, TNF@.

## **FNHTR**



#### Dx of exclusion.

 May be prevented by use of LR blood products, but WBC's during storage in plt. units may produce cytokines.
 not life-threatening but fearful and annoying and responds to antipyretics



# **ALLERGIC REACTIONS**

- <1% OF Tx.
- > caused by Ab's to donor plasma proteins.
- vary from mild (urticaria, itching, mild fever) to anaphylactic Rxn.

if mild rxn develops: no progression to severe rxn with continuation of infusion ( not dose-related)



- > most mild rxn do not recur and respond to antihistamines.
- So with mild localized rxn give antihistamines and if symptoms subside continue the Tx.
- > the risk of Tx transmitted dis. with new exposure is higher than that of restarting Tx.
- > not to be applied to rxn's with fever, chills, or vasomotor instability.
- > LR will not prevent it.

# **IF SEVERE:**



> Stop Tx, IV access, fluids, epinephrine/steroids. > for further Tx: washed rbc's. > Tx of plasma containing products? difficult: risk-benefit balance, premedicate with high dose steroids, antihistamines.



> Make epinephrine available.

Anaphylaxis may be seen in IgAdeficient pt's who were immunized previously, so use products from IgAdeficient donors (difficult).

# HYPERVOLEMIA



> IF too large a volume given too quickly.

- S &S: headache, plethora, CHF, dyspnea and systolic rise over 50 mm Hg.
- > Will subside if: Tx stopped, sitting position, O2, diuretics...



- if not phlebotomy.
- > prevention: not more than 2-4 ml/Kg/hr.
- > pts at risk (children and the elderly): slower or smaller aliquots.

## TRALI



- > Acute lung inj.: noncardiogenic acute pulmonary edema.
- > due to donor high titer anti-HLA or WBC's Ab's (leukoagglutinines).
- > accumulations of activated recipient's WBC's in pulmonary capillaries leading to microvascular occlusion and leakage.



> most common cause is: WBC Ab's in plasma-containing products from multiparous female donors, but may be due to recipient high titer anti-wbc reacting with donor's wbc's infused with RBC's, plt. or granulocyte components.



> clinically: severe resp. distress, hypoxia, hypotension, fever, pulmonary edema, occurring during or within 4 hrs of transfusion. Treatment is supportive: O2, ETT, ICU...

> further Tx: LR



- > when suspected: call bl. supplier to quarantine all products derived from the same donor.
- Dx: test donor and recipient for anti-HLA or antileukocyte Ab's and if positive HLA typing.

# **BACTERIAL CONTAMINATION**



A rare but severe complication; can be fatal

- bacteria enter bl. bag at collection, during handling or from occult bactermia in the donor.
- > 0.3% of blood units.

Sm-ve and Gm+ve: in rbc's and plt. (at 4 C or at room temp.)



> during or several hrs after Tx: chills, rigor, severe fever, shock.
> usually no Hbemia or Hburia.
> Aggressive therapy and antibiotics.
> Culture the suspected unit + Gm stain

# HYPOTHERMIA



> Rapid Tx of refrigerated blood.

- > may induce cardiac arrest due to rapid cooling of sino-atrial node.
- > also overwarming >42 C: shock.
- So bl. must be warmed by electric warmer to <42 and not by microwave or under hot water.

# **METABOLIC COMPLICATIONS**



- > Hypocalcemia: from citrateted bl. infused rapidly: circumoral and finger parasthesia.
- Slow the Tx or if large volumes as in aphaeresis: oral ca.
- > with normal liver function: citrate to bicarb.



V Calcium: NOT recommended and adding ca to bl. bag is prohibited (it reverses the citrate effect resulting in bl. clots)



## Hyperkalemia



# rare, due to massive Tx of old bl. (in neonates)

# Hypokalemia



#### > from alkalosis due to bicarb. resulting from metab. of citrate in massive Tx.

#### **Delayed Transfusion Reactions**



Adverse effects of Tx can be seen from several days to yrs after Tx.
Risk vs. benefit difficult to judge.
Think twice before Tx.



# Delayed hemolytic Tx rxn:



> transfused rbc's induce Ab response days (anamnestic) or weeks (primary response) after Tx.

> with each unit Tx an estimated risk of 1-1.6% of sensitization to Ag's other than D.

> not life-threatening: because Ab's do not fix Comp. beyond C3



- > extravascular hemolysis: mild symptoms, mild anemia despite Tx, mild hyperbili., +ve DAT, no Hbemia.
- If pt. shows severe symptoms: Rx as AIHTR.
- > Ab's involved are: Rh and Kell.

**Transfusion-Associated Graft**vs-Host Disease (TA-GVHD) > Immunocompetent donor lymphocytes to immunsupp. pt's. > also when donors are homozygous for **HLA and immunocompetent recipients** are heterozygous for same HLA haplotypes. (usually in related but may be in unrelated donors).



#### > 10-12 days after Tx

- > fever, skin rash, diarrhea, hepatitis and marrow aplasia.
- > fatal in most cases.
- Prevention: irradiate cellular components 25 Gy for susceptible pt's and all cellular components from relatives regardless of immune status.

# HEMOSIDEROSIS



> 1 mL of rbc's: 1 mg iron; in Tx dependent pt's damage to heart and liver.

> deferoxamine.



## **Transfusion Transmitted Diseases**

Despite testing: HIV, HBV, HCV and syphilis may occur.

- > we do minimum testing (no HBc antibody, p24, HTLV-I/II)
- > most common is hepatitis (window + low level Ag).

 in north America: HBV 1 in 31,000 units, HIV 1 in 3 million, HCV 1 in 4 millions.
 (Kleinman; Tran med rev; 17:2; 2003).



# Thank you