Oh "G," What Role Does the Blood Bank and Transfusion Service Play in Perinatal Care?

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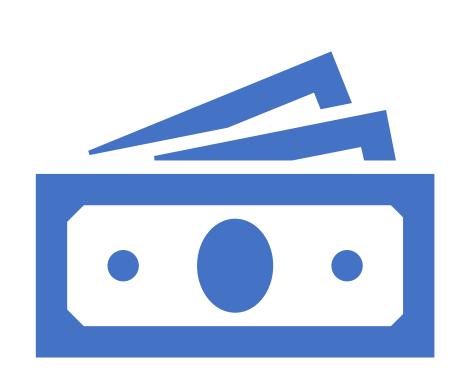
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## Background

Blood Bank System Coordinator at Billings Clinic

Adjunct Instructor of Immunohematology at Montana State University Bozeman

Board Certified as a Medical Laboratory Scientist and Specialist in Blood Banking



# Conflicts of Interest and Financial Disclosures

None

## Objectives

1

Recognize the importance of prenatal testing and consistent monitoring of antibodies.

2

Describe the clinical use of available blood products and compare their storage and transport requirements.

3

Describe the importance of Kleihauer-Betke/Fetal Hgb Flow testing and how to appropriately calculate RhIG dosing.

## Hemolytic Disease of the Fetus and Newborn (HDFN)

- Occurs when fetal cells enter maternal circulation
- Fetal red blood cell antigens that the mother lacks stimulate her to produce antibodies
- Maternal IgG antibodies cross the placenta and bind to fetal antigens, causing red blood cell destruction

## Types of HDFN

#### **ABO**

- Most Common
- Occurs in 1:125 births
- Mild HDFN

#### Rh

- Affects Rh(D)NegativeWomen
- Moderate to Severe HDFN

# Alloantibodies other than anti-D

- Mild to Severe:
   E, c, C, k, Kp<sup>a</sup>,
   Kp<sup>b</sup>, Js<sup>a</sup>, Js<sup>b</sup>, Jk<sup>a</sup>,
   Jk<sup>b</sup>, Fy<sup>a</sup>, Fy<sup>b</sup>, S, s,
   and U
- Severe: K, Jr<sup>a</sup>, Ge

# Recommended Prenatal Testing to Identify/Predict HDFN

#### **Initial Prenatal Visit**

- ABO/Rh
- Antibody Screen
  - If positive, Antibody Identification and Titration of IgG antibodies

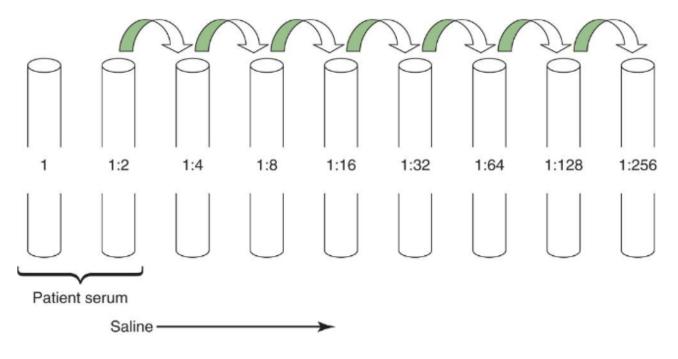
#### Follow-up Visits (If IgG antibody is identified)

- Antibody Screen with Identification
- Antibody Titration—performed in parallel with initial sample at 4-6-week intervals

#### 26-28 Weeks Gestation

- ABO/Rh to confirm D typing
- Antibody Screen (Rh negative patients)

Antibody Titration



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## Additional Methods to Monitor HDFN



### Middle Cerebral Artery Doppler

Typically done every 2 weeks to track the degree of fetal anemia

>1.5 multiples of the mean (MoM) predicts significant fetal anemia



#### **Cordocentesis**

Can be done to collect a sample of fetal blood to determine hematocrit

## Opportunities for Fetomateral Hemorrhage

# Events of Pregnancy

- Delivery
- Antepartum hemorrhage
- Spontaneous or therapeutic abortion
- Abdominal trauma

# Complications of Pregnancy

- Ectopic pregnancy
- Stillbirth
- Fetal Demise
- Placental Abruption

# Medical Procedures

- Amniocentesis
- Chorionic villus sampling
- Cordocentesis
- External cephalic version
- Manual removal of placenta

Gestational Age	Fetal Hgb Flow	RhIG dosing
<20 weeks	Not indicated	1 vial
20-28 weeks	Yes	Calculated from Hgb Flow results
28 week (Prophylaxis)	NA	1 Vial
28-40 weeks	Yes	Calculated from Hgb Flow results

Antenatal
Care for
suspected
FMH in Rh(D)
Negative
Patients

<sup>\*</sup>Type and Screen should be performed prior to any RhIG administrations

## Upon Admission for Delivery



Hold for Blood Bank

- -Low risk for HDFN
- -No previous uterine incision
- -Singleton pregnancy
- -≤ 4 previous vaginal births
- -No known bleeding disorder
- -No history of PPH



Type and Screen

- -Prior cesarean birth or uterine surgery
- -Multiple gestation
- -> 4 previous vaginal births
- -Chorioamnionitis
- -History of previous PPH
- -Large uterine fibroids



Crossmatch

- -Positive Antibody Screen (previous or current)
- -Placenta previa, low lying placenta
- -Suspected Placenta accreta or percreta
- -Hematocrit <30 AND other risk factors
- -Platelets <100,000
- -Active bleeding on admit
- -Known coagulopathy

## Available Blood Products

Product	Information	Storage Temp	Transport Temp
Red Blood Cells	Packed Red Blood Cells: Increase recipient's oxygen carrying capacity by increasing the mass of circulating red cells.  Used in treatment of symptomatic or critical deficit of oxygen-carrying capacity.  Whole Blood  Contains RBC's, plasma, and platelets.  Increases recipient oxygen carrying capacity by increasing the mass of circulating red blood cells.  Used in life-threatening hemorrhage where oxygen carrying capacity, coagulation factors, platelets and volume expansion is needed.	1-6°C	1-10°C
FFP/Plasma	<ul> <li>Serves as a source of plasma proteins (coagulation factors) for patients who are deficient in or have defective plasma proteins.</li> <li>Used in management of preoperative or bleeding patients who require replacement of multiple coagulation factors (e.g. liver disease, DIC), MTP's, and exchanges in patients with TTP.</li> <li>EXPIRATION: FFP: 24-Hours from thaw date/time PLASMA: 5 days from thaw date/time Liquid Plasma: 26 days</li> </ul>	Frozen: ≤-18°C Thawed: 1-6°C	Frozen: ≤-18°C Thawed: 1-10°C

## Available Blood Products (cont'd)

Product	Information	Storage Temp	Transport Temp
Platelets	Goal of platelet transfusion is to provide adequate numbers of normally functioning platelets for the prevention or cessation of bleeding.  Platelets can be off the rocker for a maximum of 24-hours during transport	20-24°C (Room Temp)	20-24°C (Room Temp)
Cryoprecipitate	<ul> <li>Serves as a source of fibrinogen, vWF, Factor VIII, and Factor XIII.</li> <li>Used to control bleeding associated with fibrinogen deficiency, and when recombinant and/or virally inactivated preparations of fibrinogen, Factor VIII, Factor XIII, and vWF are not readily available</li> </ul>	Frozen: ≤-18°C Thawed: 20-24°C (Room Temp)	Frozen: ≤-18°C Thawed: 20-24°C (Room Temp)
	EXPIRATION: 6-Hours from thaw date/time		

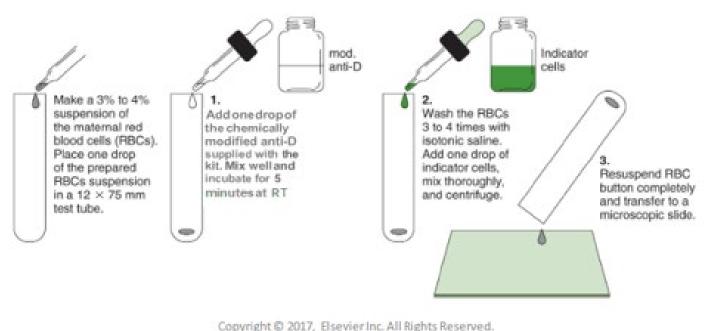
## Neonate Blood Bank Testing

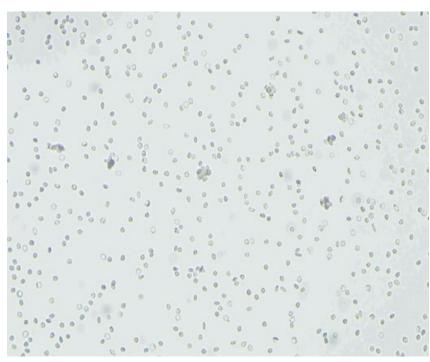
#### Cord Blood Screen

- Includes
  - Neonatal ABO/Rh
  - DAT (IgG Only)
  - Cord Blood Interpretation
  - Maternal Rhogam Interpretation
- REQUIRED on all babies born to Rh(D) Negative mothers
  - Rh of baby is used to determine mother's eligibility for post partum RhIG
- Recommended on all babies born to type O mothers
  - ABO and DAT are used to determine if ABO HDFN is present

# Additional Testing: Rh(D) Negative Mothers Who Give Birth to Rh(D) Positive Babies

### Fetomaternal Hemorrhage Screen (FMH Screen)

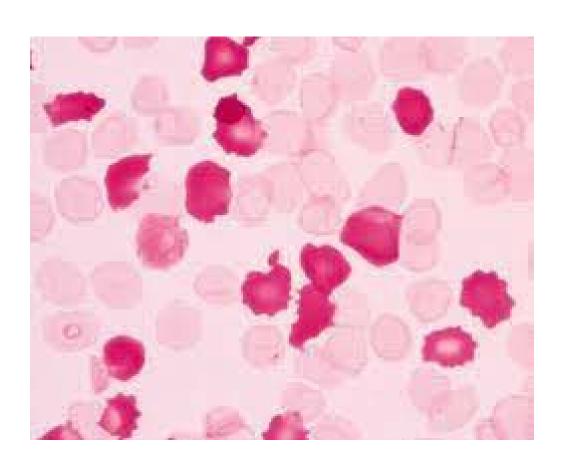




# Additional Testing: Rh(D) Negative Mothers Who Give Birth to Rh(D) Positive Babies

#### Kleihauer-Betke

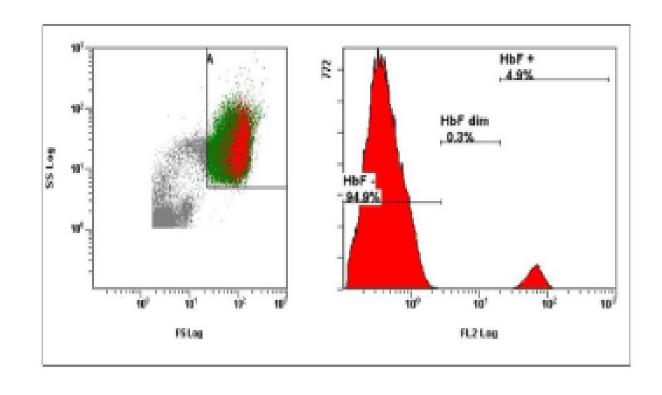
- 1. Prepare thin blood smears by diluting blood with equal volume saline.
- 2. Fix and stain smears using erythrosine B followed by Harris hematoxylin.
- 3. Examine dry smears under 40x magnification, count a total of 2,000 cells and record the number of fetal cells observed.
  - Fetal cells stain dark pink
- 4. Calculate the percentage of fetal red cells in the total counted.



# Additional Testing: Rh(D) Negative Mothers Who Give Birth to Rh(D) Positive Babies

### **Fetal Hgb Flow**

- Fluorescent tags are added to well mixed maternal sample collected after delivery and incubated, then washed
  - Fetal cells-antibody to HbF
  - Maternal cells-antibody to Carbonic Anhydrase (CA)
- 2. Sample is placed into flow cytometer and at least 50,000 events are read.
- 3. # of HbF and CA are counted and percent of HbF is determined.



## Calculating RhIG Doses

### Current Height and Weight Provided

First, calculate Body Surface Area
(BSA):

BSA (m<sup>2</sup>) = 
$$\sqrt{\frac{(h\times w)}{3600}}$$

h=height in centimeters w=weight in kilograms \*round to 3 decimal places

Next, calculate Total Blood Volume (TBV):

TBV (mL)= BSA ( $m^2$ ) x 2370 ( $mL/m^2$ )

Whole Blood FMH (mL) = 
$$\left(\frac{\% \text{ fetal cells}}{100}\right) x \text{ TBV}$$

# Vials of RhIg = 
$$\frac{Whole Blood FMH (mL)}{30 \left(\frac{mL Whole Blood}{vial}\right)}$$

If # after decimal point is	Then	And
<5	Round down	Add 1 vial
≥5	Round up	Add 1 vial

## RhIG Calculation Example

### Current Height and Weight Provided

First, calculate Body Surface Area
(BSA):

BSA (m<sup>2</sup>) = 
$$\sqrt{\frac{(h \times w)}{3600}}$$

h=height in centimeters w=weight in kilograms \*round to 3 decimal places

Next, calculate Total Blood Volume (TBV):

TBV (mL)= BSA ( $m^2$ ) x 2370 ( $mL/m^2$ )

Patient weighs 84 kg and is 172 cm 1.5% fetal cells were detected

• BSA=2.003m<sup>2</sup>

• TBV=4747 mL

## RhIG Calculation Example (cont'd)

Whole Blood FMH (mL) = 
$$\left(\frac{\% fetal cells}{100}\right) x TBV$$

# Vials of RhIg = 
$$\frac{Whole Blood FMH (mL)}{30 \left(\frac{mL Whole Blood}{vial}\right)}$$

If # after decimal point is	Then	And
<5	Round down	Add 1 vial
≥5	Round up	Add 1 vial

Whole Blood FMH (mL)= 71.2 mL

• # Vials=2.4

 # after the decimal is <5, so we round down and add 1

Total # of vials given= 3

Questions?

## Case Study

29-year-old G2P1 patient presented at an outside facility at 10 weeks and 4 days with planned pregnancy. At this visit, the prenatal panel was ordered, which included an ABO/Rh and Antibody Screen.

O Negative

Positive Antibody Screen



At this time, it was also noted that with her first pregnancy, she was induced and had an operative vaginal delivery (vacuum extraction). She had a positive antibody screen (passive anti-D), which led to post partum dose of RhIG getting missed even though baby was Rh(D) positive.

Patient was referred to MFM at our facility, where an ABO/Rh and Antibody Screen were repeated.

- O Negative
- Positive Antibody Screen—Anti-D and anti-C were identified

#### Questions

- Does this combination of antibodies warrant any concern?
- What should we do next?

### Anti-G



G antigen is found on red blood cells that possess the C or D antigens



Antibodies to G appear as an anti-D and anti-C that cannot be separated



Identification is done by using adsorption and elution techniques



Only clinically significant in OB cases to determine if RhIG is necessary

Anti-G WITHOUT anti-D—Can receive RhIG

Anti-G WITH anti-D—Not a candidate for RhIG

Through Adsorption and Elution procedures, it was found that the patient did have an anti-D, anti-C and anti-G

NOT a candidate for RhIG and at risk for severe HDFN.

Antibody titers were done

- Anti-D 1:64
- Anti-C 1:128

Titers were >1:16 MFM determined that the risk for fetal anemia is severe and performed MCA Doppler's from 18 weeks until delivery.



Blood Bank was actively involved in this patients care with the MFM.

This patient has a very rare Rh phenotype D-C-c+E+e- (0.1%)

Recommended patient undergo autologous red blood cell donation



At 29 weeks, MCA Doppler was >1.5 and patient was referred out for intrauterine transfusions

- Received 2 intrauterine transfusions (IUT) at an Academic Medical Center and returned for delivery.
  - D-, C-, c+, E-, e+ red cells were administered during IUT
  - In addition to previously identified anti-D, anti-C, and anti-G, mom developed anti-e
  - Due to the rarity of D-C-e- red blood cells, patient had to deliver at the Academic Medical Center where she received IUT
  - 2 weeks following delivery, baby presented to EMR requiring transfusions due to severe anemia due to HDFN

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