



OBSTETRICS & GYNECOLOGY

(25) Rh Isoimmunization

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Objectives:

Not given!

Rh Isoimmunization

Rh Isoimmunization:

- Is an immunological disorder occurs in a pregnant Rh-ve mother carrying Rh+ve fetus.
- It affects 1 in 250 live births in Europe and North America, it is much less frequent in other parts of the world such as Asia, where the Rh-negative blood group is uncommon.
- Pathophysiology: The Rh antigen is limited to the red cell surface (Rh complex, C,D,E,c,e). *Rh negativity is referring to the big D.

Fetomaternal Haemorrhage (FMH):

Rh isoimmunization can only take place if fetal red cells **cross the placental barrier** into the maternal circulation.

When does it happen?

- The placenta is subjected to maximal **trauma** during delivery.
- After **abortion** with a gestational age above 14 weeks.
- APH/trauma.
- External version. *A delivery procedure for the breech position.
- Amniocentesis.
- Complicated and difficult deliveries.
- Caesarean section.

The Natural History Of Rh Isoimmunization:

Rhesus antibodies are humoral antibodies or free antibody:

- IgM large, unable to cross the placenta.
- IgG small, able to **cross** the placeta and attach itself to Rh positive red cells leading to haemolytic anaemia.

Immune Responses Are:

- A. Primary first response to an antigen appears after several weeks and is **IgM**.
- B. Secondary when exposed for the 2nd time a primed, antibody will appear within a few days and its IgG.

Generally, the quantity of antigen required to produce a secondary immune response is **very much smaller** than that required to initiate the primary immune response.

Pathophysiology:

- The first pregnancy is usually unaffected by RHD because FMH's
 of sufficient magnitude to induce primary immunization do not
 usually take place until delivery.
- Only about 5% of all Rh-negative mothers form antibodies.
- Vast majority of FMH's after delivery are small but about 0.2% of mothers have larger bleeds of 30 ml or more. The risk of Rh immunization is proportional to the size of the FMH.

ABO Incompatibility:

When the mother and the baby are ABO incompatible such as an O mother and an A baby any fetal red cell (Group A) entering the maternal circulation (Group O) is destroyed, in an exactly similar way to that occurring in an ABO incompatible blood transfusion.

Pregnancy:

FMH does occur during pregnancy but is much less common than following delivery. Most of the bleeds occur in the <u>last trimester</u> when the placenta is degenerating and the barrier may become a little more pervious.

The Prevention of RHD:

D-positive FMH's can be neutralized by passively administered anti-D antibody (Rh immunoglobulin).

When to give it? (During every pregnancy):

- At 28 weeks.
- Post delivery.
- After abortion.
- APH/trauma.
- External version.
- Amniocentesis.

Failure Rate:

About 1% of Rh-ve women become immunized after D-positive pregnancies despite treatment with Rh immunoglobulin.

- Those **already primed**, even though overt antibody is undetectable by present techniques.
- Large FMH's before delivery e.g. epileptic or eclamptic patients.
- Extreme sensitivity to the D-antigen: thus small bleeds will produce primary response.
- Large FMHs after delivery more than the amount that can be taken care of by standard dose of immunoglobulin.
- Failure to give the immunoglobulin patients who slip through the net.

Management of Isoimmunization:

Pregnancies complicated by clinically relevant isoimmunization are managed in centers with <u>fetal medicine</u> units and regional blood transfusion.

- a) Maternal blood group and antibody quantification.
- b) Paternal blood group genotyping.
- c) Fetal blood group genotyping.

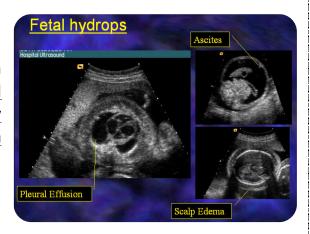
- d) Ultrasound assessment.
- e) Amniotic fluid spectrophotometry.
- f) Fetal blood sampling.
- g) Fetal blood transfusion.

MGT:

- Maternal blood group and antibody quantification at booking and 28 weeks if initial is –ve.
- Paternal blood group genotyping.
- Anti-D titer is not serious **if below 1:16** and should be repeated every 2-4 weeks.
- If titer is **above1:16** invasive testing:
 - Ultrasound assessment.
 - Amniotic fluid spectrophotometry.
 - Fetal blood sampling.
 - Fetal blood transfusion.

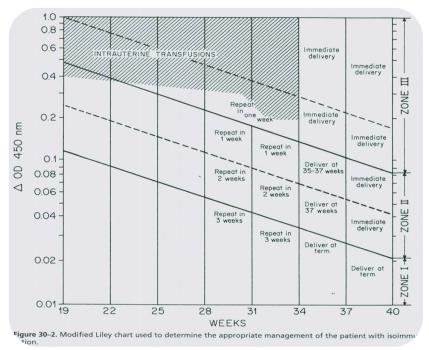
Ultrasound Assessment:

- The severely anemic fetus on scan will have: <u>skin edema</u>, <u>ascites</u>, <u>pleural</u> or <u>pericardial effusions</u>, <u>cardiomegaly</u> and an <u>edematous</u> <u>placenta</u> (<u>Hydrops</u>).
- Middle cerebral artery blood flow is increased.



Amniocentesis and Amniotic Fluid Analysis:

- When fetal heamolysis occurs the amniotic fluid becomes bright yellow from the **bilirubin**.
- Amniotic fluid bilirubin concentration can be quantified by spectrophotometry by assessing the change in optical density at 450nm (\triangle OD 450).
- <u>Amniocentesis is started after 24 weeks</u> under ultrasound guidance.



Complications Following Intrauterine Transfusion:

- 1. Premature labour.
- 2. Pre-labour ruptured membrane.
- 3. Fetal haemorrhage.
- 4. Fetal bradycardia.
- 5. Failure to obtain a sample.
- 6. Increase in maternal Isoimmunization by inducing feto-maternal haemorrhage.

CTG:

Fetal heart rate changes have been noted with severe anemia. A sinusoidal pattern with the loss of normal baseline variability of the CTG is highly suggestive of severe anaemia.

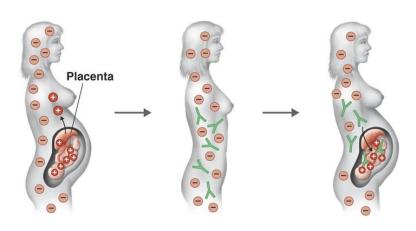
Irregular Antibodies:

- 2% *Other than Rh D.
- Kell, Duffy, Kidd....etc.

Summery

- ✓ Rh Isoimmunization is an immunological disorder occurs in a pregnant Rh-ve mother carrying Rh+ve fetus.
- ✓ Rh isoimmunization can only take place if fetal red cells cross the placental barrier into the maternal circulation.
- ✓ Rhesus antibodies are humoral antibodies or free antibody:
 - IgM large, unable to cross the placenta.
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- ✓ Immune Responses Are:
 - Primary first response to an antigen appears after several weeks and is IgM.
 - Secondary when exposed for the 2nd time a primed, antibody will appear within a few days and its IgG.
- ✓ D-positive FMH's can be neutralized by passively administered anti-D antibody (Rh immunoglobulin). When to give it?
 - At 28 weeks.
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 - ✓ Management Of Isoimmunization:
 - Maternal blood group and antibody quantification.
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 - ✓ Ultrasound Assessment:
 - The severely anemic fetus on scan will have: skin edema, ascites, pleural or pericardial effusions, cardiomegaly and an edematous placenta (Hydrops).
 - Middle cerebral artery blood flow is increased
 - ✓ CTG: Fetal heart rate changes have been noted with severe anemia. A sinusoidal pattern with the loss of normal baseline variability of the CTG is highly suggestive of severe anaemia.





- 1 Rh+ father.
- Rh⁻ mother carrying her first Rh⁺ fetus. Rh antigens from the developing fetus can enter the mother's blood during delivery.
- In response to the fetal Rh antigens, the mother will produce anti-Rh antibodies.
- If the woman becomes pregnant with another Rh+ fetus, her anti-Rh antibodies will cross the placenta and damage fetal red blood cells.

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MCQ's:

- 1. The blood group of a 28-weeks pregnant lady is ARH negative. Her husband's group is O Rh negative. What is your next step regarding her blood group?
 - a. Check Rh-antibody every 4 weeks.
 - b. Give anti-D prophylaxis at 34 weeks.
 - c. Give anti-D within 72 hours after delivery.
 - d. Reassure her that there is no risk of Rh immunization.

2. Anti-D prophylaxis:

- a. Should be given to all sensitized Rhesus negative women after delivery.
- b. Should be given to all Rhesus negative women after amniocentesis.
- c. Should be given to all Rhesus positive women who give birth to Rhesus.
- d. Is contra-indicated during pregnancy if the women is Rhesus negative.

Ans: 1:d. 2:b.