



Transfusion Reactions

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BMS Education Day

28th January and 4th February 2013

This presentation provides realistic case studies which have been designed with audience participation in mind and should encourage discussion within the group. The cases are NOT designed for individual learning and if used in this way caution is advised. Each case unfolds as time goes on reflecting a true clinical situation and, as a result, there are no clear-cut correct answers until the case is concluded. Discussion of these case studies should use problem-solving skills, be based on effective communication between the laboratory and the clinicians and take into consideration national and local policies on the management and investigation of transfusion reactions.

Note: *It will be beneficial for staff to read the BCSH guidelines on the investigation and management of acute transfusion reactions and to review the local transfusion reaction policy.*

Disclaimer: *All of these cases reflect a similar case in clinical practice but they have been modified so that the hospital, the staff and the patient are not identifiable.*

The information given reflects current clinical and laboratory practice but local variations may be in place.

Slides 1-7 Introduction to the subject

This presentation is for Biomedical Scientists and assumes a basic knowledge of the adverse effects of transfusion. This brief introduction provides an overview of other support and learning materials.

Category of Reaction	Infectious	Non-infectious
	Bacterial contamination	<u>Immune</u> Febrile Non-Haemolytic Allergic Acute Haemolytic Reaction -ABO Anaphylactic -anti-IgA Transfusion Related Acute Lung Injury
		<u>Non-immune</u> TACO/TAD
	Viral – HIV, Hepatitis, Parvovirus Protozoal – Malaria, Syphilis Prion - vCJD	<u>Immune</u> Delayed haemolytic reaction Transfusion Associated-GvHD Post Transfusion Purpura
		<u>Non-immune</u> Iron overload

As you participate in this session please;

- Understand your role in when a patient has an adverse reaction to blood components



- At the same time try and understand everyone else's role too

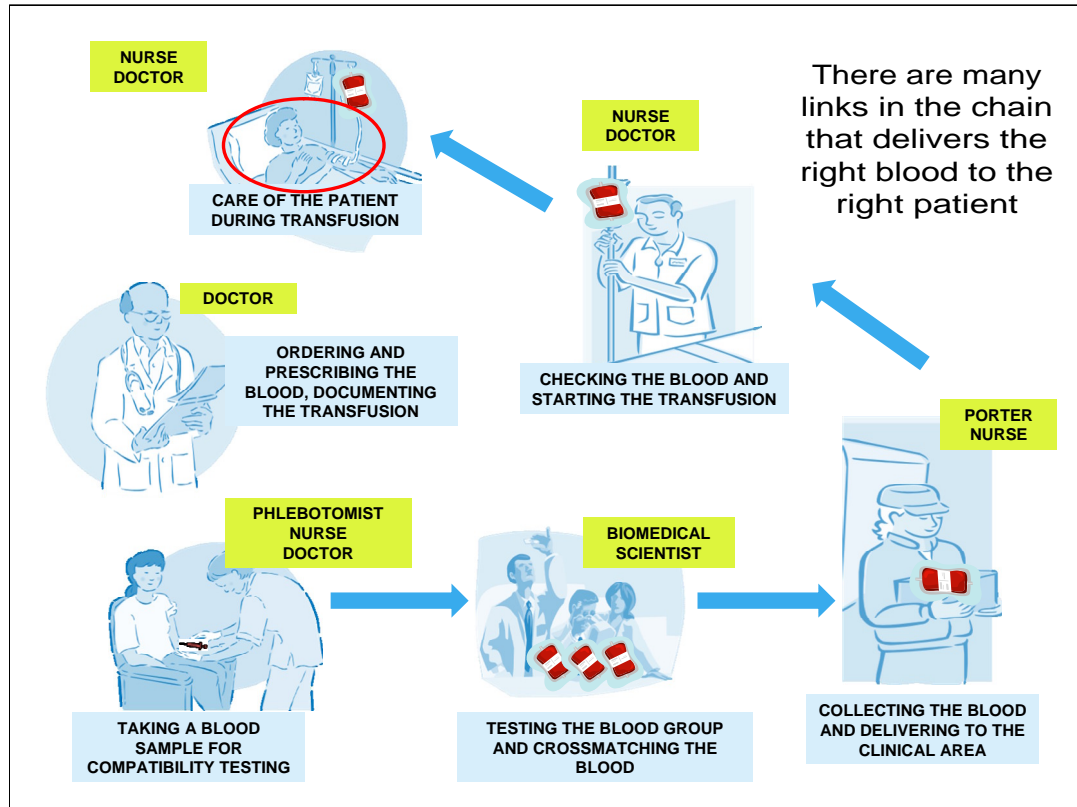
DOCTOR, NURSE,

BIOMEDICAL SCIENTIST, PORTER



- Never forget the patient who is having the reaction. That's why we are all here.....





How do we know, what we know?

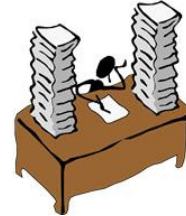
FIRSTLY: What can go wrong?

- We have been reporting **Serious Hazards Of Transfusion** to SHOT since 1996
- MHRA have been collecting **Serious Adverse Blood Reactions** and **Events** via the SABRE website since 2005

Look at the SHOT definitions and the annual report on the website www.shotuk.org

SECONDLY: What should we do about it?

- Transfusion Medicine Handbook (for clinicians)
 - Useful flow chart
- BCSH guidelines (for laboratory and clinicians)
 - Compatibility guidelines
 - Acute transfusion reaction guidelines



Look at websites www.bcshguidelines.com and www.transfusionguidelines.org.uk

Hospitals & Science

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NHS Blood and Transplant

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adverse events & reactions

Requests for Investigation of Adverse Events and Reactions

All files are stored in a PDF format and require Adobe Acrobat Reader.

Download Adobe Acrobat Reader.

Component supply:

Cell Immuno-haematology

in D Neg Women and inadvertent transfusion of D

Transfusions within NHS Blood and Transplant (NHSBT)

Supply of Allogeneic Blood and Supply of Blood for Transfusion

Products for Recipients of ABO/Rh Mismatched Stem Cell

Distinguishing Passive from Immune anti-D in Pregnancy (PDF)

Provision of Red Cell Transfusion Support for Transfusion Dependent Patients (PDF)

Investigation and Clinical Management of Suspected Reactions to IgA (PDF)

Investigation and Clinical Management of Patients with a positive DAT with and without Haemolysis (PDF)

Diagnosis and Management of T Antigen Activation (PDF)

Platelet and White Cell Transfusion, Histocompatibility and Immunogenetics

Transfusion Associated Graft Versus Host Disease (PDF)

Guidelines for The Management of Platelet Transfusion Refractoriness (PDF)

Transfusion Related Acute Lung Injury (TRALI) (PDF)

Post-Transfusion Purpura (PDF)

Clinical Guidelines for the use of Granulocyte Transfusions (PDF)

Neonatal Alloimmune Thrombocytopenia (PDF)

Thrombasthenia (PDF)

hospital.blood.co.uk





Interactive Session



Clinical information

The orange boxes represent the clinical picture.

There are many things going on and transfusion may only be a small part of the picture.

Remember, they know more about the patient than you do

Laboratory information

The blue boxes represent the laboratory situation.

This is often the only information you have access to when deciding what is going on.

Remember, you know more about transfusion practice than they do

Learning Points

To take back and use in everyday practice

Slide 8 – Introduction to the cases

This part of the education session was interactive and examples were used to stimulate conversation and group learning. No answer was deemed incorrect but justification was needed for your answer. Notes have been added after the presentation to help with some guidance when delivering this back in the work place and used for learning. Discussion with your lab manager and/or training manager will aid learning.

There are learning points throughout the presentation (highlighted in black) containing knowledge that can be applied in daily practice.

Monday 9.30 am Transfusion Dept.

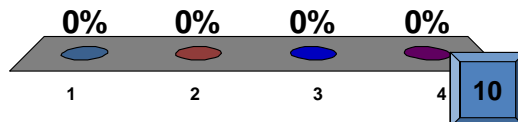
- 30 year old woman with suspected ectopic pregnancy came into the ED at 8.45
- Described as 'in severe pain but stable'
- FBC, coag. screen, G&S already processed by out of hours BMS.
- Second G&S sample requested, received and tested
- All 4 core identifiers present on samples; not previously known to your hospital
- Hb 90g/L, coag. screen normal, A RhD positive
- Going to theatre in 10 minutes, request for 4 units of blood immediately

Monday 9.30 am Emergency Dept.

- 30 year old woman with severe left sided abdominal pain arrived 45 mins ago.
- Positive pregnancy test
- Scan suggests ectopic with free-fluid +++
- Following morphine is pain-free but very drowsy & clammy, hypotensive (80/? mmHg) and fast pulse (120/min)
- Decided to operate
- Urgent transfusion needed because of internal bleeding

Q1 : What blood would you issue?

1. Emergency O RhD negative, no XM
2. Group specific, uncrossmatched
3. Group specific, electronic issue
4. None of these, something else



Slide 11 – Case study 1, question 1

Option 1: (Emergency O RhD negative) Whilst O RhD negative is a safe option, it is appropriate to conserve O RhD negative blood for those patients where an alternative cannot be used. A group and screen on 2 samples was already available and, providing the samples were labelled according to the sample acceptance policy and had been taken from the right patient, group specific blood could have been given.

Options 2 and 3: The patient had a second sample taken which was received and tested so the patient should be eligible for electronic issue if this is the laboratory policy and all the other criteria have been met. As the case is an emergency it would not be inappropriate to release group specific uncrossmatched units as 2 samples have been tested.

Discussion point:

What processes do you have in the laboratory to issue blood in an emergency?

What crossmatching techniques are available in your laboratory?

What are the criteria for electronic issue?

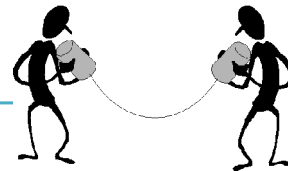
Option 4: If this option was selected there should be discussion as to what the 'something else' is. Be aware that this is a female of childbearing potential so avoiding sensitisation to the D and K antigens should be considered.

In an emergency situation the provision of blood should be a priority.

It may be beneficial to find out from the clinical team if other blood components are required.

Monday 9.45 am Transfusion Dept.

- You ask “do you want to activate the major haemorrhage protocol?”
- Gynae SpR says “no, but please issue 4 units of blood as soon as you can and send it straight to theatres”
- You issue 4 units of A RhD positive red cells by electronic issue by 9.55am



Monday 10 am Gynae. Theatre

- 2 units of blood are given over 10 minutes
- Surgery starts at 10.05am
- The ruptured ectopic in the left fallopian tube is removed and bleeding stops
- 1 litre of blood is drained and surgery is finished by 10.30am
- In recovery, patient is cold (35°C) and hypotensive. She is on oxygen and warmed iv fluids. She has produced no urine.

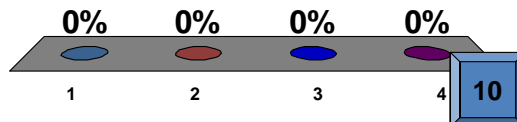
Monday 10.45 am Haematology Dept.

- Repeat FBC and coagulation screen arrive
- After 30 minutes you phone the results
Hb 50g/L, platelets $45 \times 10^9/L$,
PT 20 seconds, APTT 60 seconds, fibrinogen 1.0 g/L
- She has moved to intensive care and they request 4 more units of blood, 4 FFP and 1 ATD platelets
- You issue group A blood, platelets and FFP

Q2 : What is the most likely explanation for these blood results?

1

1. She has DIC because of the ectopic
2. She has dilutional coagulopathy because of all the blood and iv fluids
3. The surgery has not succeeded in stopping the bleeding
4. She has DIC because of an ABO incompatible red cell transfusion



Slide 15 – Case study 1, question 2

There may be not enough information at this stage to give a full and complete answer and there should be a discussion with regards to which answers were chosen and why.

Option 1: Ectopic pregnancy can cause DIC, this option would not explain the drop in Hb.

Option 2: We do not have information on what other IV fluids, or the volume, the patient has received. A review of the patient's treatment may help support or exclude this diagnosis.

Option 3: This option may explain the drop in Hb in this case. This is a possibility.

Option 4: DIC and drop in Hb with no urine production are symptoms of acute haemolytic transfusion reactions. If a patient has had a blood transfusion this may be an option due to sampling/administration/checking errors. However, in this case there were 2 samples grouped for this patient and a two sample policy reduces the risk of WBIT.

Discussion points:

How confident are staff with the laboratory testing for ABO? What are the signs and symptoms of an ABO transfusion reaction?

What additional samples would you request and what tests would you perform?

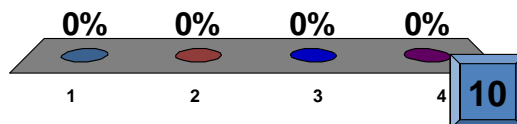
Monday 11.15 Intensive Care Unit

- The lab phone to say the blood components are ready
- The lab is concerned that the blood is haemolysed
- You have noticed very poor urine output and very dark urine
- The patient is very sick and deteriorating rapidly
- You discuss with the haematology SpR and both of you are worried about an acute haemolytic reaction

Q3 : The patient is retested and is group O instead of group A. What is the most likely explanation?

1

1. The transfusion laboratory tested the wrong patient's sample
2. The clinicians bled the wrong patient
3. The transfusion laboratory made a mistake when testing the sample
4. There were probably two patients with a similar name admitted at the same time



Slide 17 – Case study 1, question 3

The options were given to encourage discussion and learning between the group of delegates.

Options 1 & 3: This is a possibility, but this would have meant that the same errors occurred twice on 2 different runs, as the laboratory received and tested 2 separate samples for this patient.

Discussion points:

How confident are you with regards to your laboratory ABO typing?

Could there be a possibility a mistake could be made on 2 separate samples?

Option 2: Wrong blood in tube (WBIT) has been reported in all Trusts. The new BSCH Guidelines state that 2 samples, taken at 2 separate times, should be tested prior to a transfusion. There has been reports of 2 samples been taken at the same time and one kept and sent later in order to save time and not bleed the patient twice – this practice does not reduce the risk of a wrong blood in tube incident. The Department of Health have listed an ABO incompatible transfusion as a 'never event'.

Discussion points:

Why might the person taking the blood take 2 samples at the same time?

Can you think of any safe guards to prevent an ABO incompatible transfusion?

Option 4: Failures in the final administration stage have occurred and have been documented in past SHOT reports. It may be that the wrong patient was bled at the time of admission although 2 samples were received in the laboratory. This is a likely source of error but the patient may have had a dedicated team involved in transfusion and communication with the lab as she was bleeding and in urgent need of treatment.

ABO-incompatible transfusions

- The 'two-sample rule' can prevent ABO incompatible transfusions **IF** the samples are taken at separate times (but you don't always get this in a massive haemorrhage situation)
- Shift hand-over times are error-prone; incomplete handovers, hurrying and cutting corners
- In a complex clinical situation ABO incompatibility (or any transfusion reaction) can be missed because the symptoms are attributed to something else
- If you have any suspicions or clues in the lab there is a WBIT or that wrong blood has been given, always speak out immediately

Thursday 9pm, Care of the Elderly Ward

- Mrs. P is 75 and has had myelodysplasia for the last 18 months, requiring red cell transfusions every 3 weeks and occasionally requires platelets. She is refractory to random platelets as a result of HLA-antibodies
- She was admitted 2 days ago with a high fever and is now on iv antibiotics for pneumonia
- The nurse checks today's blood results and contacts the on-call doctor
Hb 75 g/L, platelets $23 \times 10^9/L$, WBC $27 \times 10^9/L$
- The doctor reviews the patient, who is quite breathless, and arranges 3 units of blood

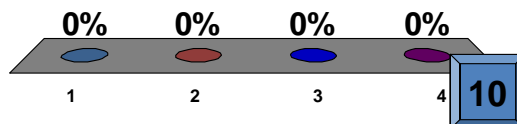
Thursday 10pm, Haematology Dept.

- The on-call BMS takes a verbal request for 3 units of red cells
- The patient is O RhD positive and has anti-Fy^a
- The last transfusion was 10 days ago
- 2 units of O+ Fya- blood have already been crossmatched for tomorrow
- There is also one unit of HLA-matched platelets for her, which expires at midnight tonight

Q4 : What is the most important thing to communicate to the doctor caring for Mrs. P tonight?

2

1. That blood and platelets are available, but give the platelets before midnight
2. That the patient has RBC and HLA antibodies
3. To discuss the case with the haematology team
4. That it is hospital policy to avoid overnight transfusion, unless the patient is unstable



Slide 21 – Case study 2, question 4

All the options contain information that would be useful for the Dr. Get the group to vote for the option and have a discussion regarding the choices made.

Option 1: This is useful information although the patient may have already been reviewed by the haematologist managing the patients care, hence the 2 units already crossmatched for tomorrow. The platelets have not been requested and by telling the Dr that they are expiring at midnight may mean that they decide to give these. Making sure all requests are appropriate should be the key concern.

Option 2: The fact that the patient has antibodies may not be something the Dr knows about his patient and is useful information.

Option 3: As the patient has MDS and has been treated for it for the last 18 months they will be under the care of the haematology team. Referral to this team to check the treatment plan would be important as there may be a good reason to wait and transfuse the patient in the morning as originally planned.

Option 4: Due to reduced number of staffing and patient safety it is not recommended that patients are routinely transfused at night. If a transfusion is not urgent it should be given during routine hours. In this scenario the patient is breathless and therefore may be deemed symptomatic.

Discussion point:

If unsure of the appropriateness of a request who do you refer to?

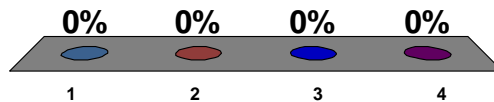
Friday 2.30am, Care of the Elderly Ward

- The doctor prescribes the platelets over 30 minutes and each RBC unit over 2 hours, covered with oral furosemide (diuretic)
- The baseline pre transfusion obs. at 11pm were
 - BP 130/85, pulse 102, temp 37.5°C, resp. rate 28
- The platelets were given uneventfully
- Within 15 minutes of the first RBC unit the patient becomes more breathless
 - BP 140/90, pulse 110, temp 39.2°C resp. rate 32

Q5 : You receive a phone call from the nurse on the ward to say Mrs. P is having a transfusion reaction and they have stopped the blood. What would you say?

2

1. Don't call me, there's nothing I can do
2. Get the doctor looking after the patient to assess the reaction and speak to the on-call haematologist for advice
3. Tell the clinical team to document the reaction and send the implicated RBC unit back to the transfusion department
4. None of these, something else



Slide 23 – Case study 2, question 5

Patient has an increase in temperature, blood pressure and respiratory rate which are signs of a transfusion reaction.

Option 1: The transfusion labs are the experts in the transfusion medicine field and are the point of contact and support for the clinical area. If it is felt that dealing with this is not within your knowledge base you should be able to advise them who to contact.

Options 2 and 3: These are both considered correct actions. Reference to local policy should be made. Suspected transfusion reactions should always be referred to the haematologist.

Discussion points:

If you received the implicated unit in the laboratory what tests would you perform?

Would a post transfusion sample be requested?

Option 4: Discussion should be held as to what other actions would have been done at this point if this was selected by anyone in the group. It may be that this option has been chosen because both option 2 and 3 would have been selected.

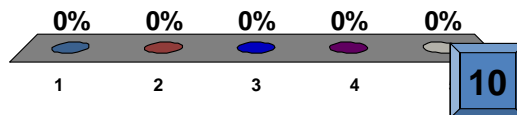
Friday 9am, Haematology Dept.

- The haematology SpR reports that Mrs. P is now ventilated on the Intensive Care Unit
- She has had blood cultures and a change of iv antibiotics but is still pyrexial
- She had poor renal function before but this is now much worse
- At 8am her Hb was 60g/L, platelets $55 \times 10^9/L$, WBC $12 \times 10^9/L$

Q6 : What do you think the likely cause of this transfusion reaction?

2

1. Transfusion Associated Circulatory Overload
2. Transfusion Related Acute Lung Injury
3. Haemolytic Transfusion Reaction
4. Anaphylaxis
5. Bacterial contamination the red cells



Slide 25 – Case Study 2, question 6

Option 1: The patient has had significant changes in observation without a significant fever and is breathless. These are signs of TACO. The weight of the patient has not been given and it is unclear what other fluids the patient has been given. The patient also has poor renal function.

Option 2: The patient has had significant changes in observation without a significant fever and is breathless. These are signs of TRALI. With TACO, the central venous pressure (CVP) will be raised and with TRALI the CVP will be normal.

Option 3: The patient has anti-Fya but the units transfused were Fya negative and crossmatch compatible. Haemolytic transfusion reactions related to non-ABO blood group systems do not tend to be immediate.

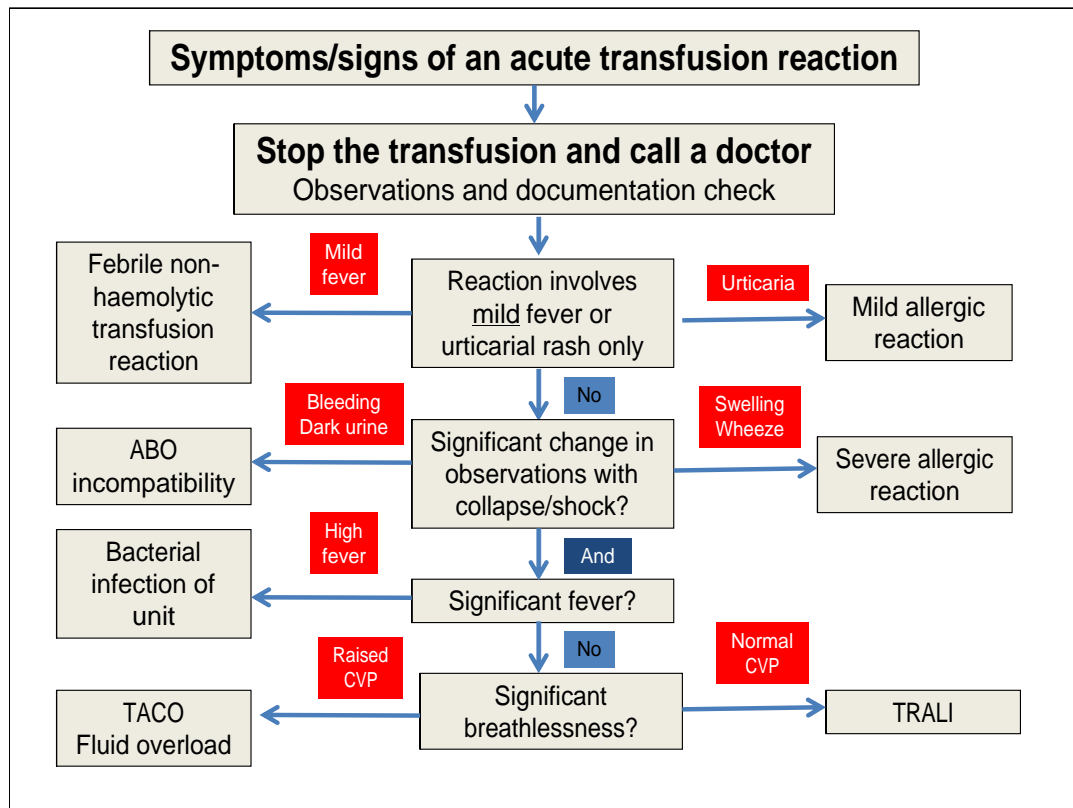
Option 4: Anaphylaxis causes shock/severe hypotension with a wheeze or stridor. This can be caused by transfusion in patients who are IgA deficient. The patient should be treated with adrenaline.

Option 5: The temperature rise was less than 2°C but the temperature is above 39°C. Bacterial contamination presents with shock/severe hypotension without clinical sign of anaphylaxis or fluid overload.

Discussion points:

What would be the most likely diagnosis based on the results and clinical picture presented so far?

Could more than one diagnosis be possible?



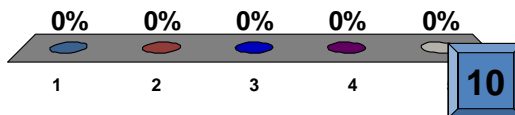
Friday 11.15am, Intensive Care Unit

- Blood cultures taken the night before grow pseudomonas, the same organism that was detected in the sputum on admission
- Chest X-ray suggests worsening pneumonia with bilateral infiltrates and a large heart
- She has clinical signs to suggest heart failure and is in positive fluid balance
- The haematology SpR reviews the patient and suggests a plan of investigation

Q7 : Which set of investigations would be the most useful in this case?

2

1. Repeat blood group, crossmatch and antibody screen. Check bilirubin and look at a blood film
2. Send the remains of the RBC unit to NHSBT for culture and ask them to recall any associated units
3. Discuss the case with the TRALI panel
4. Check an IgA level and anti-IgA antibodies
5. None of these, something else



Slide 28 – Case Study 2, question 7

Discuss all options bearing in mind the following notes, BCSH guidelines and local policy.

Option 1: These tests are useful when looking for a serological transfusion reaction. In this case these are not the most useful set of investigations.

Option 2: When bacterial contamination is suspected NHSBT should be informed to quarantine other products made from this donor. In this case the patient's blood cultures have shown no additional bacterial growth and the pseudomonas detected is the same as the organism detected on admission. Therefore it is very unlikely that the reason for the patient's deterioration is due to bacterial contamination of the transfusion.

Option 3: If a TRALI is suspected the case should be discussed with the TRALI panel at NHSBT. Samples will be requested and tested for neutrophil antibodies. If a TRALI is suspected treatment should not be delayed until the laboratory results are returned.

Option 4: IgA deficient patients may have anaphylactic reactions following a transfusion. This patient however has been transfused many times and therefore would have had a reaction previously.

Option 5: TACO. This patient has a positive fluid balance and has clinical signs to support cardiac failure. These are symptoms of TACO.

The patient also has impaired renal function

Respiratory Complications of Transfusion

2

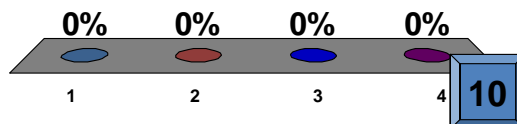
- Breathlessness is a feature of most severe acute reactions
- After resuscitating the patient, there needs to be a clinical review to guide further investigations
- It is important to focus investigations on the case – don't do every test on every case
- The pattern of symptoms and the time of onset after the start of transfusion is usually more helpful than laboratory investigations
- Not all breathlessness is transfusion-related!

Friday 4pm, Haematology Day Ward

- 21 year old male with very severe aplastic anaemia diagnosed 2 months ago
- He requires platelets twice a week and RBC transfusion every 2 weeks
- He continues to attend his university course and intends to finish his degree. The final exams are in 3 weeks
- He has not had any recent infections
- There have been no previous transfusion reactions
- He is about to go home when he starts to feel itchy and shows you a rash over his arms and body

Q8 : What should the day-ward staff do?

1. Arrange for him to have washed platelets next time
2. Take a set of observations and advise him to wait to be reviewed by the doctor
3. Give him antihistamines and then let him drive home if the rash settles
4. Alert the blood service and send the unit of platelet for culture to exclude bacterial contamination



Slide 31 – Case Study 3, question 8

Option 1: it is possible he could have developed an allergy to the plasma in platelets although this is the first time the patient has had a reaction. Washed platelets should be considered if severe reactions occur on more than one occasion.

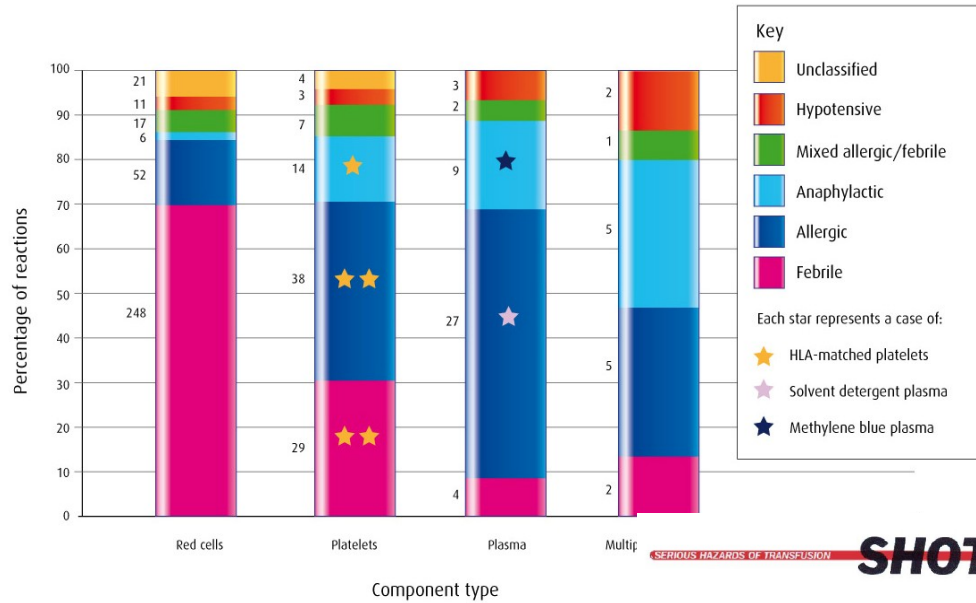
Option 2: The patient should wait and be reviewed by the doctor before being allowed to go home.

Option 3: Antihistamines can be given to deal with the allergy but this causes drowsiness so he should not be allowed to drive home

Option 4: Unlikely, but we do not have all the clinical details on the patient. If there had been a significant rise in temperature there may be reason to suggest bacterial contamination.

Category	Mild	Moderate	Severe
Febrile reaction	Temperature rise up to 2°C No other symptoms/signs	Temperature rise of 2°C or more Rigors, chills other inflammatory symptoms	Temperature rise of 2°C or more plus symptoms that require urgent medical review or prolong hospital stay
Allergic reaction	Transient flushing, urticaria or rash	Wheeze or angiooedema but no significant breathlessness or hypotension	Bronchospasm, stridor, angiooedema, circulatory problems. Anaphylaxis
Reaction with both allergic and febrile features	Features of mild febrile and mild allergic reactions	Features of both allergic and febrile reactions One feature in moderate category	Features of both allergic and febrile reactions One feature in severe category
Hypotensive reaction		Isolated fall in BP of 30mm or more No allergic or inflammatory symptoms	Shock without allergic or inflammatory symptoms

Reaction by Component Type



Allergic Reactions

- Are quite common
- Sometimes related to a factor in the donor
- Occur in patients with other allergies
- Sometimes mild reactions can get worse and patients should be observed for a period of time

Learning Points

1. **A multi-professional team approach to transfusion reactions produces the best results**
2. **Patients present with symptoms of their underlying disease as well as those of transfusion reactions**
It is **GOOD** when clinical staff consider the possibility of a reaction, so be encouraging!
3. **Symptoms and timing of onset are clues to the cause of the reaction**
Flowcharts are a useful complement knowledge of the type of reactions
4. **Mild reactions should be discussed and documented in the notes**
But do not require extensive investigations