

CLINICAL GUIDELINE

Special Requirements of Transfusion

A guideline is intended to assist healthcare professionals in the choice of disease-specific treatments.

Clinical judgement should be exercised on the applicability of any guideline, influenced by individual patient characteristics. Clinicians should be mindful of the potential for harmful polypharmacy and increased susceptibility to adverse drug reactions in patients with multiple morbidities or frailty.

If, after discussion with the patient or carer, there are good reasons for not following a guideline, it is good practice to record these and communicate them to others involved in the care of the patient.

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Approval Group:	Overarching Blood Transfusion Committee

Important Note:

The Intranet version of this document is the only version that is maintained.

Any printed copies should therefore be viewed as 'Uncontrolled' and as such, may not necessarily contain the latest updates and amendments.

Greater Glasgow and Clyde Blood Transfusion

Special Requirements Guideline

Issue Number: 3 Version: v3 Date: June 2022 Review Date: June 2024 Ownership: GG&C Overarching Transfusion Committee

Introduction

This guideline is for all staff involved in authorising, supplying and administering blood and blood components throughout Greater Glasgow and Clyde Health Board (GG&C). It should be used in conjunction with the GG&C Clinical Transfusion Policy. The aim of the Clinical Transfusion policy is to ensure that the **right** blood is given to the **right** patient at the **right** time, **every** time.

This policy focuses on those patients who have a special requirement for transfusion and it aims:

To provide healthcare professionals with clear guidelines on when Cytomegalovirus (CMV) seronegative components, irradiated components or other special requirements for transfusion are indicated

- To outline the process of notification to the hospital Blood Bank, of patients with special requirements
- To outline how this information will be disseminated across Greater Glasgow and Clyde
- To outline how errors should be reported if they arise.

Advice on blood transfusion may be obtained from Biomedical Scientist (BMS) staff in Blood Bank, from Haematology Medical Staff and from hospital Transfusion Practitioners.

Hospital Site	Contact details for Blood Bank within core working hours	Contact details for Blood Bank out with core working hours Monday – Friday 5pm-8am Friday 5pm – Monday 8am 0141 242 9606/9609		
Glasgow Royal Infirmary	Monday – Friday 8am-5pm 0141 242 9606/9609			
Stobhill ACH	See above No Blood Bank on site – Contact GRI	See above No Blood Bank on site – Contact GRI		
Gartnavel General	Monday – Friday 9am-8pm 0141 301 7728/7729	Monday – Friday 8pm-9am Friday 8pm -Monday 9am GRI Blood Bank -0141 242 9606/9609		
Queen Elizabeth University Hospital	Monday - Sunday 24hrs 0141 354 9104 / 9105 (ext 89104 / 89105)	Monday - Sunday 24hrs 0141 354 9104 / 9105 (ext 89104 / 89105) Page 17602		
Victoria ACH	See above No Blood Bank on site – Contact QEUH	See above No Blood Bank on site – Contact QEUH		
Royal Alexandra	Monday – Friday 8.30am – 5pm 0141 314 6159 (ext 06159)	Monday – Friday 5pm – 8.30am Friday 5pm – Monday 08.30am Page 56359		
Inverclyde Royal	Monday – Friday 8.30am – 5pm 01475 504323 (ext 04323)	Monday – Friday 5pm – 8.30am Friday 5pm – Monday 08.30am Page 51005		
Vale of Leven Monday – Friday 8.30am – 5pm 01389 817502 (ext 87502)		Contact RAH Laboratory via page 56359		

This policy has been prepared for the GG&C Overarching Transfusion Committee and represents current standards. It will be reviewed every two years to accommodate future changes in the provision of health care in GG&C and in the provision of blood and blood components.

Reason For Special Requirements And Identification Of Patients

- Special requirements cover a range of additional specifications or processing of standard blood components
- In case of emergencies, if appropriate blood components meeting the patient's specific transfusion requirements are not available, then standard blood components must be issued to avoid unnecessary transfusion delays

The most commonly requested special requirements in practice are CMV seronegative components and irradiated components.

CMV Seronegative Components

Cytomegalovirus (CMV) belongs to the herpes family and in the healthy population causes a self-limiting illness. It is cell associated and therefore can be transmitted in cellular blood components (red cells, platelets and granulocytes). Typically it causes a subclinical illness similar to infectious mononucleosis (glandular fever). In immunocompromised patients it can result in potentially fatal pneumonitis or disseminated CMV infection. For this reason CMV seronegative products should be used for a selected group of patients, listed below. The requirement for CMV seronegative blood components has been reviewed by the Advisory Committee on the Safety of Blood, Tissue and Organs (SaBTO) ¹. Universal leucodepletion, introduced by the UK blood services in 1999 primarily as a vCJD risk reduction measure, results in a three log depletion of white cells. This is generally accepted to render these components "CMV safe"

Monitoring selected patient groups for CMV infection with pre-emptive therapy where required has been shown to be a successful strategy in certain groups (solid organ transplants, haematopoietic stem cell transplants). CMV seronegative components should continue to be used where leucodepletion is not possible (granulocyte transfusions) or where the potential severity of the consequences of CMV infection, and/or the difficulty in monitoring for infection are considered too great (IUT and transfusions in neonates up to 28 days post EDD, elective transfusions in pregnancy).

Pregnant women, where possible, should receive CMV seronegative components.

There is no requirement for this for emergency transfusions in pregnancy.

As CMV is a common virus and the prevalence of anti-CMV in the UK adult population is 50-60%, it not possible to provide CMV seronegative products for all recipients. Components that have been leucocyte depleted to less than 5×10^{6} /unit have a significant reduction in the risk of CMV transmission.

IgA Deficiency

Primary IgA deficiency, defined as selective IgA < 0.07g/l in adults, with normal levels of IgG and IgM, not secondary to other causes is relatively common (1:500 caucasians), 20-40% of these individuals have anti-IgA-antibodies although these are a poor predictor of possible transfusion reactions. Patients with IgA deficiency may have allergic or anaphylactic reactions to blood components, however severe transfusion reactions in this group of patients are extremely rare (1: 20,000 – 1: 47,000 transfusions).²

Most IgA deficient patients with or without anti-IgA-antibodies will not experience severe reactions with standard components. Plasma reduced or IgA deficient components can be supplied, however this may lead to a delay in transfusion and the consequences of the delay should be balanced against the risk of a reaction.

Patients with IgA deficiency receiving a transfusion should be closely monitored in a clinical area where severe allergic and anaphylactic reactions can be managed.

In cases where there is a history of severe allergic/anaphylactic transfusion reactions, the use of components from an IgA deficient donor should be considered if it will not cause significant delays. Transfusion requirements for patients with IgA deficiency should be discussed with the local Haematologist.

Irradiated Components

Irradiation is the only known method of preventing Transfusion Associated Graft versus Host Disease (TaGvHD). This is a rare but fatal complication of blood transfusion. It results from engraftment and clonal expansion of HLA compatible donor lymphocytes. Clinically TaGvHD presents with fever, rash, diarrhoea, hepatitis and pancytopenia 4-30 days after transfusion. This complication of transfusion is eliminated by inactivation of donor lymphocytes by irradiation. The risk associated with individual transfusions depends on the number and viability of contaminating lymphocytes, susceptibility of the host's immune system to their engraftment and the degree of immunological disparity between donor and recipient.^{3,4,5}

As new treatments are developed there may be additional therapeutic agents for which irradiated components are required. If there is any concern about the need for irradiated components, prescribers should first check with pharmacist or haematologist.

Hepatitis E Seronegative Components

All blood and blood components supplied by SNBTS are hepatitis E negative and no longer require to be requested as a special requirement.

The table below identifies the patients who have requirements for CMV seronegative and irradiated components and the length of time these are required.

Condition	Cytomegalovirus (CMV) Seronegative	Irradiated	Duration for special requirements 28 days post EDD	
Neonates up to 28 days post EDD	\checkmark			
Intrauterine Transfusion (IUT)	✓	√ **	For 6 months after 40 weeks gestation	
Neonatal Exchange Transfusion (ET)	✓	✓		
All donations from first or second degree relative		\checkmark		
Severe T lymphocyte immunodeficiency syndromes including Di George and Severe Combined Immunodeficiency	à	à		
Recipients of CAR-T Therapy		\checkmark	For 2 weeks prior to harvest until 3 months following CAR-T cell infusion	
Recipients of allogeneic haemopoietic stem cell transplantation (HSCT)		~	From three months pre transplant to 6 months after transplant or while patient is immunosuppressed	
Recipients of autologous haemopoietic stem cell transplantation (HSCT)		\checkmark	From transplant until 6 months post-transplant	
Stem cell harvesting		√	From mobilisation or 2 weeks before the harvest (whichever is earlier) until harvest is completed	
All haematological recipients of alemtuzumab (Campath, anti-CD-52)		✓ ***	Lifelong	
All patients with Hodgkin's lymphoma		~	Lifelong	
All patients treated with purine analogues, e.g. fludarabine, cladribine, deoxycoformycin, clofarabine, bendamustine		\checkmark	Lifelong	
All patients with aplastic anaemia treated with immunosuppressive therapy (until lymphocyte count >1.0 x10 ⁹ /L)		~	Lifelong	
Patients with aplastic anaemia (potential stem cell transplantation)		~	From diagnosis where there is a high likelihood of proceeding to allogeneic HSCT	
Pregnant women (In an emergency standard leucocyte-depleted products should be given to avoid delay)	v		Until delivery	

Leucodepleted CMV seropositive components (CMV safe) may be used for the above patient groups in the situation of life-threatening haemorrhage CMV seronegative components should only be used for CMV negative patients or those in whom CMV status is unknown.

*Top up transfusions in infants aged <6 months do not require irradiation unless there has been a previous IUT or if the donation is from a first or second-degree relative

**The requirement for irradiation after IUT remains until 6 months after 40 weeks gestation

***The use of irradiated blood components is not indicated following treatments with alemtuzumab using the schedule currently recommended for MS or vasculitis or indicated for patients undergoing solid organ transplantation

† Whilst a diagnosis of severe T lymphocyte immunodeficiency is considered all components transfused should be irradiated and CMV seronegative whilst further tests are being undertaken. Adults and children > 2 years without a significant history of infection do not need irradiated cellular blood components unless there is a significant history consistent with a severe T lymphocyte associated immunodeficiency

Plasma Reduced Components

Plasma-reduction (previously known as washing) of cellular components (red cells and platelets) reduces levels of potentially harmful substances, e.g. serum proteins (IgA), antibodies, additive solutions, electrolytes (K+) etc. The process results in some cell losses, and takes several hours to complete. Plasma reduced components have a reduced shelf life.

The indications for plasma reduced components include:

- recurrent and / or severe allergic reactions or febrile reactions to transfusion which are not abolished by the use of leucodepleted components
- IgA deficient patients, as long as provision of plasma-reduced components does not cause unacceptable delays. Of note, in cases with a history of severe allergic/anaphylactic transfusion reactions, transfusion with components from an IgA deficient donor should be considered if this would not introduce an unacceptable delay
- neonates with T-activation and haemolysis following transfusion of standard blood components, should receive platelets in additive solution if they require platelet transfusions
- patients with unexplained hypotensive transfusion reactions
- maternal platelets for transfusion to neonates with neonatal alloimmune thrombocytopenia (NAIT)

Life-long transfusion requirements (including patients with haemoglobinopathies)

For a small number of patients with high life-long transfusion requirements, e.g., thalassaemia or sickle cell, the use of phenotyped blood should be considered to reduce the risk of the patient developing antibodies. These patients may require extended crossmatched blood or sickle negative (HbS negative) blood. The blood bank should be informed of patients with these diagnoses, so that appropriate components can be supplied.

If there is any doubt as to whether or not a patient has a requirement for special blood components please discuss with the on call Haematologist at your local site.

Notification To The Laboratory Of Patients With Special Requirements

Once a patient has been identified as having a requirement for special blood components, the Special Requirements of Blood Transfusion Laboratory Request Form, used throughout GG&C (see appendix 1), should be printed from the intranet at the webpage below:

http://www.staffnet.ggc.scot.nhs.uk/Acute/Diagnostics/BloodTransfusion/Documents/Special%20Requirement% 20form%20for%20Blood%20Products%20v070213%201.doc

The fully completed form should be sent to the local Blood Bank. The information held on this form will be shared with all Blood Banks throughout GG&C and the Golden Jubilee National Hospital.

The Consultant responsible for the patient should ensure that the information on special requirements of transfusion is marked in the patient's electronic record (Clinical Portal and Trakcare as appropriate).

The form should be signed by the Consultant in charge of the patient's care. It should be scanned and uploaded onto Clinical Portal with the second copy forwarded to the Blood Bank. If the patient has been identified as having special blood component requirements out with core working hours and a Consultant is not available, the requirement can be requested on a regular transfusion form and the special requirements form forwarded at the earliest possible opportunity.

The information will not be disseminated throughout the Health Board Blood Banks until a Consultant signed copy of the special requirements form is received by the hospital Blood Bank.

If the need for the special blood components changes, an updated form should be forwarded to the hospital Blood Bank so their records can be amended.

Laboratory Dissemination of Information

As patients regularly transfer between hospital sites it is important that their transfusion requirement information is available throughout the Health Board Blood Banks. Once a Consultant signed form is received at a local Blood Bank, the form will be scanned and emailed to the Blood Bank managers in the South, North, Clyde and the Golden Jubilee Hospital so that their computer systems can be updated accordingly (as per the Blood Bank special requirements SOP).

The form will be returned to the requesting Consultant to ensure they are aware the laboratory tasks have been completed.

Patient Counselling And Consent

It is not a legal requirement in the United Kingdom that written consent is obtained before a transfusion of blood components; however, it is considered a required standard of good transfusion practice to ensure that the patient/parent/carer receives adequate information regarding the transfusion⁶. This discussion should include information about the risks and benefits of transfusion as well as information relating to available alternatives, for example, iron supplementation and must be documented in the case records. Nursing and medical staff therefore, have a professional duty to ensure that they have adequate knowledge of transfusion related issues or that they can access the information and support required by patients undergoing a transfusion.

Booklets are available specifically for patients who require irradiated components and all patients receiving irradiated components should be given these before their first transfusion. They should also be counselled and told to inform nursing and medical staff of this requirement before any future transfusions. Patients should be given a card to carry documenting their special transfusion requirement. An adhesive sticker, taken from the information leaflet, should be placed on the front of case records where these exist, detailing the requirement. It is the responsibility of the authorising practitioner to ensure these steps are undertaken.

Incident Reporting

There have been 13 reported cases of TaGVHD since 1996 but no cases after 2001. 2 cases have been reported in leucodepleted components (1998-99 and 2000-01) indicating that while leucodepletion significantly reduces, it does not eradicate the risk of this condition. It is important therefore that appropriate products are provided for patients at risk.

Errors do occur and the Serious Hazards of Transfusion (SHOT) ⁷ report 2019 showed 259 cases where special requirements were not met, an increase from 194 in 2018. Of the 259 cases, 102 occurred in the clinical setting and 157 within the laboratory setting. 83.3% of the incidents within the clinical setting involved patients who required irradiated components but did not receive them. Of the remaining cases, specific phenotype of red cells, use of a blood warmer and the requirement for CMV seronegative components were missed. It is important that any errors are documented appropriately to ensure that steps can be taken to avoid them in the future. If it becomes apparent that a patient has failed to receive appropriate components the hospital Blood Bank **must** be informed immediately. The incident will be fully investigated and reported on Datix and to SHOT.

Duty of Candour

Duty of Candour was formally introduced in Scotland during 2018 8.

Duty of Candour requires professionals and organisations to be open and honest with patients where there have been failings in their care. This legal requirement means when unintended or unexpected events happen, that result in death or harm as defined in the Act^{7,8}, that the people affected understand what has happened, receive an apology, and that organisations learn how to improve for the future. A Duty of Candour event is when an individual has received treatment or care that has resulted in an unintended or unexpected event, which, in the reasonable opinion of a registered health professional, results in death or harm as defined in the Act^{8,9}.

Some serious transfusion incidents may fall into this category. Please refer to GG&C Duty of Candour Policy for further support and guidance

http://www.staffnet.ggc.scot.nhs.uk/Corporate%20Services/Clinical%20Governance/Clinical%20Risk/Duty%20of%20Candour/DoC%20Policy%20and%20Guidance%20GGC%20Final%20v1%20(2018).pdf

References

1. Advisory Committee on the Safety of Blood, Tissues and Organs (SaBTO) <u>Microbiological safety guidelines (publishing.service.gov.uk)</u>

2. The transfusion management of patients found to be IgA deficient Scottish National Blood Transfusion Service Policy Record NATP Clin 034 02 (2018) <u>https://www.nss.nhs.scot/media/1023/transfusion-management-of-patients-found-to-be-iga-deficient.pdf</u>

3. Clinical Blood Transfusion. Chapter 16 Postgraduate Haematology Sixth Edition. P268-300. Hoffbrand AV, Catovsky D, Tuddenham EDD, Green AR.

4. Guidelines on the use of irradiated blood components prepared by the British Committee for Standards in Haematology blood transfusion task force British Journal of Haematology 2020, 191, 704-724 Guidelines on the use of irradiated blood components | British Society for Haematology (b-s-h.org.uk)

5. Guideline on Transfusion for Fetuses, Neonates and Older Children. British Journal of Haematology 2016, 175, 784-828 https://b-s-h.org.uk/guidelines/guidelines/transfusion-for-fetuses-neonates-and-older-children/

6. Guidelines from the expert advisory committee on the Safety of Blood, Tissues and Organs (SaBTO) on patient consent for blood transfusion. https://www.gov.uk/government/publications/blood-transfusion-patient-consent

7. Serious Hazards of Transfusion http://www.shotuk.org/

8. 2018 No. 57 - Scottish Statutory Instruments - National Health Service Social Care Social Work - The Duty of Candour Procedure (Scotland) Regulations 2018 http://www.legislation.gov.uk/ssi/2018/57/made/data.pdf

9. Scottish Government – Healthcare Standards – Duty of Candour http://www.gov.scot/policies/healthcarestandards/duty-of-candour/

Transfusion Transmitted Infection. Chapter 13. Practical Transfusion Medicine. Third Edition. P122-145. Murphy MF, Pamphilon DH.

Special Requirements of Blood Transfusion Laboratory Request Form

This form should be completed for **all** patients who have special requirements for blood components. A copy should be sent to Blood Bank and a copy filed at the front of the patients clinical notes. It is the responsibility of clinicians to update Blood Bank on any changes to special requirements. A minimum annual review is required. Additional forms are available from the transfusion page on Staffnet.

Patient Details (Addressograph label if available)	Referring Consultant				
Surname:	Consultant:				
First Name(s):	Hospital:				
DOB:	Location:				
CHI Number/ TJ Number:	Contact Number				
Sex:					
Blood Product Requirements (Please tick (✓) in white column as appropriate, see below table for Haemoglobinopathy Patients and plasma reduced components)					
	CMV Seronegative	Irradiated			
Neonates up to 28 days post EDD					
Intrauterine Transfusion (for 6 months after 40 weeks gestation)					
Neonatal Exchange Transfusion (ET)					
All donations from first or second degree relative					
Severe T lymphocyte immunodeficiency syndromes including Di George and Severe Combined Immunodeficiency					
Recipients of CAR-T Therapy					
Recipients of allogeneic haemopoietic stem cell transplantation (HSCT)					
Recipients of autologous haemopoietic stem cell transplantation (HSCT)					
Stem cell harvesting					
All recipients of alemtuzumab (Campath, anti-CD-52)					
All patients with Hodgkin's lymphoma					
All patients treated with purine analogues, e.g., fludarabine, cladribine, deoxycoformycin, clofarabine, bendamustine					
Patients with aplastic anaemia treated with immunosuppressive					
therapy (until lymphocyte count >1.0 x10 ⁹ /L)					
Patients with aplastic anaemia (potential stem cell					
transplantation)					
Pregnant woman					
Haemoglobinopathy Patient (Full Rhesus and Kell matched, HbS negative products	Document indication here:				
required)	Document indication here:				
Plasma Reduced Components (Washed cells) Please see policy for indications	Document indication here.				
Alert recorded on: Portal Trakcare (Responsibility of	the Consultant named below)				
Consultant Signature:Print nar					
Review of Blood Product Requirements (Tick as appropriate)					
CMV Seronegative Irradiated Plasma Reduced (Washed) Other Change:Effective from (date):					
Consultant Signature: Print Name:					
Date: Review Date:					
For Laboratory Use Only (Tick as appropriate)					
Information transcribed into LIMS System					
Scanned copy of form sent via generic email address to the SNBTS and GJNH Blood Banks \Box					
Copy of completed form sent to referring Consultant \Box					
BMS Signature: Date:					
Print Name:					