

ANZSBT ABSTRACTS

Abstracts of selected papers presented at the 39th Annual Scientific Meeting of the Australian and New Zealand Society of Blood Transfusion, Sydney Convention and Exhibition Centre, Darling Harbour, Sydney, Australia, 16–19 October 2005

ABSTRACT NO.: 1

Molecular Basis and Investigation of Blood Group Genes

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There are 284 blood group antigens recognized by the ISBT Committee on Terminology for Red Cell Surface Antigens. Of these, 244 antigens are included in one of 29 blood groups systems. The rest are included in collections of similar antigens, or in a series of either low incidence or high incidence antigens. The biochemical work of the middle of the last century revealed these antigens to be carried on different glycoproteins and glycolipids on the surface of the red blood cell (RBC). Their usefulness as protein markers recognized by highly specific immune antibodies permitted functional studies of many of these membrane components. In the last twenty years, the genes encoding all but one of the 29 blood group systems have been sequenced, and the molecular basis identified. These studies have shown that the majority of blood group antigens are encoded by simple single nucleotide polymorphisms within the coding sequence of the gene. Where gene families exist such as at the Rh and MNS loci, high nucleotide identity between related genes has resulted in the formation of hybrid genes that encode novel antigens and create unusual phenotypes. Mutations and other alterations in the non-coding regions of genes are also responsible for creating blood group antigen diversity. The molecular identity of blood groups has many different applications. From a clinical perspective, molecular assays can be used to determine likely RBC phenotype in situations where it is difficult or undesirable to obtain RBCs. It also provides a model for the study of diversity within humans and enabled scientists to look at ancestral genes. These studies not only tell more about who we are but also map the influences of different external factors such as disease, on our evolution.

ABSTRACT NO.: 2KODETMCAE: Creating the World's First ABO Analytical Control System

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AIM To engineer controllable expression ABO glycolipids onto the surface of group O red blood cells to create weak expression ABO cells for blood grouping sensitivity controls.

METHODS AND RESULTS The natural phenomenon by which glycolipids can insert into red blood cells underpinned the development of this technology. Blood group A and B glycolipids were constructed with a novel linker molecule that confers aqueous solubility, and a carbohydrate glycocone design of generic specificity. Extensive research was carried out to determine the conditions

which facilitate controlled engineering of the red blood cell membrane to express specific levels of A and/or B blood group antigen. Cells expressing low levels of antigen were able to be constructed with KODETM technology, providing for the first time a reliable supply of cells, consistently expressing antigens suitable as analytical sensitivity controls. Comprehensive internal comparative testing and a large external field trial were performed to analyse KODETM cell stability and performance against a range of monoclonal antibodies and technologies. These trials clearly demonstrated that the KODETM constructed cells are stable and performed in the same manner as naturally occurring ABO weak cells. Furthermore they demonstrated a wide range of blood grouping result errors and discrepancies with blood grouping in Australasia, thus highlighting the need for an ABO quality control system in Immunohaematology laboratories.

CONCLUSION KODETM technology enables production of the first ABO blood grouping sensitivity control cells that can be precisely manufactured in large volumes. They have been incorporated into a process control system that consistently tests and challenges blood grouping performance.

ABSTRACT NO.: 3

A Pilot Study Evaluation of a Tool for Assessing Clinical Competency in the Administration of Blood Transfusion: The Nurses' Experience

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The change in nurse education from apprenticeship training to the higher education setting in the United Kingdom (UK) has raised concerns about the lack of practical skills newly qualified nurses have on registration. A skill, which every practitioner must be able to carry out is the safe administration of blood components however, the UK Serious Hazards of Transfusion (SHOT) scheme have consistently demonstrated that "wrong blood" incidents are the major cause of morbidity and mortality. As a result the SHOT working group have recommended that all practitioners should have their clinical competency formally assessed.

To address the above concerns and provide a means for formal assessment of clinical competency, an assessment tool was developed and evaluated using a triangulated approach of phenomenology and survey. The tool was piloted in two different clinical settings by four registered nurses who each assessed two nurses. Individual semi-structured interviews were conducted to collate the nurses' and the assessors' experience of the process.

Only two National Health Service hospitals in Scotland were identified as undertaking some form of competency assessment of blood transfusion practice. The pilot study demonstrated that the participants were of the opinion that formal assessment using a

criterion-reference tool gave practitioners the opportunity to relate theory to practice, promote best practice and encourage adherence to hospital transfusion policies. Formal assessment of clinical competency therefore has the potential to ensure that nursing practitioners are fit for purpose. Although national implementation will be challenging, the nursing profession should lead the way in defining performance standards and ensuring the safety of patients is paramount.

ABSTRACT NO.: 4

The Evolution of Haematology Nursing

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Can you imagine nursing a haematology patient before there were antiemetics, cytokines, antivirals and antifungals? When bone marrow transplant was considered 'not for today and not for tomorrow' (George Mathe, 1970), a time before the great migration from the Mediterranean and Asia, when the few patients with Thalassaemia major in Australia reached adulthood, when blood for transfusion was collected into glass bottles and was mainly reserved for trauma cases, and when haemophilia patients were treated with bed rest and pain relief? This was not at the beginning of the last century; rather it was in the late 1950's when international studies began to expose effective medical treatment for many haematological conditions.

How nursing evolved from the general nurse in the 1950s with the haematology patient nursed in open 'Nightingale' style medical wards to the highly specialised nurse of today working in specifically purpose built haematology units will be explained as we take a journey with one who has 'lived the changes'.

We will examine the material and technical changes that have had an impact and we will explore the many different ways that nurses equipped themselves to meet the changing needs of patients as they managed the introduction of a wide variety of different treatments and drugs and developed protocols, care plans and innovative programs to support their patients and their carers, and their colleagues.

Haematology nurses today are found in a wide variety of settings. They liaise; provide counselling, leadership and consultancy along with expert primary care of the patient. Their practice is evidence based and they involve themselves with research and with quality improvement.

Haematology nurses have a unique opportunity to influence future direction of care and to maximise the contribution of haematology nursing as our healthcare providers undergo generational change.

ABSTRACT NO.: 5

Development of the Nurse Practitioner Role in Haematology Stem Transplantation

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A new nursing role is rapidly emerging and gaining acceptance in Australian health care. The Nurse Practitioner role is expanding the boundaries of nursing practice and changing the face of health care in Australia.

The Nurse Practitioner is a nurse who has advanced their knowledge and skills through education and clinical experience, and who is working in a role that allows for increased clinical discretion, responsibility and autonomy. The Nurse Practitioner is authorised to perform tasks outside the practice of a regular registered nurse eg ordering of investigations and procedures, 'prescribing' medications according to unit specific clinical guidelines, and referring to other health professionals.

In February 2003, Queensland Health commenced a trial to explore and define the role of the nurse practitioner. The trial was conducted in four sites, one of these sites being in Stem Cell Transplantation within Oncology Haematology Services at the Princess Alexandra Hospital.

The Nurse Practitioner in Stem Cell Transplantation provides advanced nursing care, utilising a case management model to coordinate and manage the care of patients at various stages on the Haematology Stem Cell Program.

This paper will discuss the Nurse Practitioner role, its implementation and its contribution to quality outcomes of patients on the Haematology Stem Cell Transplant Program at the Princess Alexandra Hospital.

ABSTRACT NO.: 6

Emerging Infections: WNV and Beyond

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AIM To discuss emerging infections that impact blood safety and to outline approaches to their identification and interventions to reduce their impact.

Emerging infections are defined as those that have increased in frequency over the past 20 years. They include novel agents, such as HIV and vCJD, those that have been imported into areas where they were not previously endemic, like *T. cruzi* and West Nile virus (WNV) and those whose geographic range is expanding, like *Babesia* spp and malaria. In addition, aggressive new therapies lead to a population of patients that is much more susceptible to serious outcomes from infection with normally benign agents such as CMV and the B19 parvovirus (erythrovirus). Each emerging agent must be evaluated for its potential impact upon blood safety, including issues of public perception. Where appropriate, interventions must be designed and implemented and such interventions should ideally be continuously evaluated for efficacy. Key examples that will be discussed are the explosive outbreak of WNV in the US, where nucleic acid testing was rapidly developed and implemented, and vCJD, where preventative measures were implemented even before it became apparent that the disease was transmissible by transfusion. Other infections that will be discussed include *T. cruzi*, (the agent of Chagas' disease), babesiosis, malaria, SARS and dengue. A variety of actual or potential interventions will be discussed.

ABSTRACT NO.: 7

Lessons Learned From SARS Epidemic

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In the period of March – July 2003, Hong Kong had experienced an epidemic of Severe Acute Respiratory Syndrome (SARS), which started out as an outbreak of community acquired pneumonias of unknown aetiology. It seeded and spread rapidly throughout hospital patients and healthcare workers, with high morbidity and mortality. The outbreak has led to a boom in the art and science of infection control and health protection. Precautionary measures, such as frequent cleansing of utilities, personal hygiene and protective gears have become important. The provision of public healthcare services was re-prioritized. The pattern of blood usage changed. The demand for blood (↓12.8%), as well as the availability of donations (↓16.9%), dropped during the period. With the concerted effort of medical and scientific experts, the culprit, which was later identified as a novel pathogen, SARS-associated coronavirus (SARS-CoV) transmitted through droplets person-to-person, was isolated. Brief viraemia in patients was documented in anecdotal reports. Still, not much was known about the biology of the virus, and its impacts in relation to transfusion.

Despite the advancement in testing blood borne viral infections, there are still limitations in detecting window period donations and "new" pathogens. Blood centres have to rely on the use of health history enquiry in donor screening to alleviate the risk of transfusion transmitted infections caused by novel pathogens, of which very limited knowledge are known. SARS-CoV is a typical example of one of these emerging pathogens. During the SARS period, the HKRCBTS implemented a donor deferral policy to ensure blood

safety. The application of healthcare informatics to capture SARS or suspected SARS patient database for screening donors/donations was useful.

No one knows when, or where, it will re-emerge. Therefore, it is important to carry on further research for the understanding of its pathogenesis, laboratory diagnosis and its role in transfusion, and continuous stringent global surveillance.

ABSTRACT NO.: 8

Pre-emptive Management of Challenging Surgical and Critical Bleeding Scenarios

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Three clinical case scenarios will be presented to illustrate management of challenging critical bleeding:

I. Case 1: A Jehovah's Witness with an unanticipated blood loss of 8500 ml at reoperative hip surgery. Teaching and discussion points will include:

- (1) How low can you go?
- (2) Lifesaving "matters of conscience" blood products and peri-surgical interventions useful in the management of Jehovah's Witness patients.
- (3) Use of erythropoietin and darbepoietin in the perisurgical setting.
- (4) The Oregon Health & Science University "Transfusion Blood Refusal" form as a way to guide clinical discussion and management with Jehovah's Witness patients; how we handle transfusion of minors.

II. Case 2: A 6½ year old boy with exsanguinating pulmonary hemorrhage post bone marrow transplantation despite correction of platelet count, PT INR, PTT and fibrinogen. Teaching and discussion points will include:

- (1) Off-label use of recombinant factor VIIa in control of refractory critical bleeding.
- (2) Limitations of current standard coagulation testing and potential value of the thromboelastogram (TEG) in monitoring coagulopathy and response to rVIIa.

III. Case 3: The CABG patient from hell and his massive transfusions × 2: the first massive occurred in the context of 4 vessel post-operative graft occlusion and cardiogenic shock suspected due to heparin-induced thrombocytopenia (HIT). The patient had an intraaortic balloon pump placed preoperatively and was undergoing his third cardiopulmonary bypass (CPB) procedure in as many days for BiVAD placement as a bridge to cardiac transplantation. Anticoagulation with the direct thrombin inhibitor (DTI) argatroban was used and post-operative bleeding was hard to control. The second massive transfusion occurred in the context of his heart transplant 12 weeks later. Teaching and discussion points will include:

- (1) Coagulopathy of cardiopulmonary bypass.
- (2) Monitoring the degree of anticoagulation on DTIs and treating bleeding due to DTIs.
- (3) Treating bleeding on antiplatelet agents.

ABSTRACT NO.: 9

Attempting to Apply Logic to Chaos in Perioperative Bleeding: An Anaesthetist's Perspective

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The idealized view of surgery and anaesthesia working in concert is sometimes stressed when dealing with unstable coagulopathic patients. The clinical trigger to consider the presence of a coagulopathy is the onset of non-surgically controllable bleeding. You see it in the surgical field, and may remain the best predictor of the need to administer haemostatic agents. While earlier consensus

documents considered this event predictable by modeling blood loss, and factor level decline¹, we now recognize confounding variables such as shock, hypothermia, metabolic disturbances and fibrinolysis confuse the picture to such an extent they are often irrelevant². The monitoring of coagulopathy by traditional coagulation screens also lacks validation in non surgical bleeding, and is usually so delayed in response time to become irrelevant to appropriate management.

Attaining euvolaemia is possible with large volume dedicated infusion systems but in a patient with massive mediator release and comorbidities may not achieve adequate tissue oxygen delivery, and inotropic support may be needed. Evidence based transfusion triggers are still lacking, but the degree of insult means while adequate haemoglobin replacement is possible, correction of the coagulopathy in the presence of ongoing loss, hypothermia and acidosis is not. Active warming methods are more advanced and effective, with forced air warming, infra red radiant heaters, and extracorporeal circuits available. Management of coagulopathy is by component replacement. Delays occur because of slow coagulation test turnaround times, and the place of newer point of care monitors and thrombelastography needs evaluation, and shows promise³. Delay in delivery of blood components due to processing time may worsen the coagulopathy. Factor VIIa probably has a unique place in the management of coagulopathy in these patients, but studies are needed to define dosage and time of dose⁴. The management of acidosis and hyperlactaemia are fundamentally improved by better perfusion, but short term correction may be indicated to reinforce coagulation factor function⁵.

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ABSTRACT NO.: 10

Your Body, Your Choice

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Many authorities have identified the need for change in transfusion practice. The precautionary principle, unsustainable increasing (and currently underestimated) direct and indirect costs of blood, chronic blood shortages, donor deferrals, loss of altruism, wide variations in transfusion practice and growing knowledge of transfusion limitations and possible adverse outcomes necessitates a paradigm shift in transfusion practice and the management of anaemia and critical bleeding. Historically, changing medical practice in a sustained manner has been challenging. A recent editorial suggested changes in transfusion practice would require "a cultural shift among clinicians, managers, and policy makers."

Informed and empowered patients can be important drivers for change in transfusion practice. Surveys in Australia, United States, Canada and Europe reveal consumers, including lay public and

health professionals, would prefer alternatives to donor blood transfusions¹.

A force for change in recent years has been the international evolution of comprehensive, patient-centred, blood conservation/management programs. These programs are effecting significant reductions in blood usage (42–95%) along with positive patient and fiscal outcomes. Strategies include managing the patient's own blood by a multidisciplinary, peri-event approach, utilising individualised patient assessment and work-up, strategies to minimise blood loss, optimise red cell mass and a greater understanding of anaemia and its appropriate management. Transfusion decisions are based on individualised patient-specific clinical and physiological factors. Decisions also need to consider patient values and choices.

Bloodless surgery, initially developed as a response to patient request, has metamorphosed into the global approach of comprehensive patient blood management – a cultural shift addressing “bloody problems” that can take transfusion medicine into the 21st century.² The AABC is working to establish such programs in Australia.

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ABSTRACT NO.: 11

Cost Effectiveness of Transfusion Therapy vs. Blood Conservation Therapies

Axel Hofmann, ME

To date governments, hospital administrators and clinicians have grossly underestimated the cost of blood transfusions and transfusion related processes. This may be attributable to a lack of comprehensive and precise cost analyses in this clinical field. In Europe blood components and blood products already account for the biggest proportion of therapeutics purchased by hospitals, but they represent only a *fraction* of the overall transfusion related cost. The total cost might be in the vicinity of 20 billion USD for the European Union alone. Many different activities and services within a hospital are directly related to transfusions such as receiving, controlling and appropriate storage of RBCs, managing and administering the internal blood supply, transfusion preparation, including extensive laboratory work with numerous regular and irregular tests, non-productive labor time/stand-by time, administration and monitoring of transfusion, treatment of immediate adverse effects, reporting and documenting, cleaning, hemovigilance and long-term outcomes-tracking. In addition, several of these cost elements have a constant tendency to increase disproportionately high, compared to public health expenditures.

From this perspective, blood conservation strategies (“bloodless medicine”) and patient blood management are becoming increasingly important. If accurate cost data can prove a more favorable cost effectiveness ratio of blood conservation therapies compared to transfusion therapies, then it is more likely that the paradigm will shift towards a new gold standard for maintaining patient's tissue oxygenation.

The challenge to measure the total cost of these competing therapies can be best met by using ‘Process Cost Analysis’ or ‘Activity Based Costing’. This methodology enables the capturing of actual resource consumption in terms of labor, materials, third party services and capital for each single process step of a therapy. Such costing models, to compare blood conservation strategies and transfusion therapies, are currently being developed.

ABSTRACT NO.: 12

Removal of Infective Prions by the Chromatographic Processes of Plasma-derived Albumin and Immunoglobulin

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AIM To demonstrate, using a validated scaled-down laboratory model, that the chromatographic processes for the manufacture of plasma-derived albumin and immunoglobulin (including IVIg) are capable of removing prions should they be present in the plasma pool.

METHODS Two sets of studies have been conducted to evaluate the capacity of several albumin and immunoglobulin process steps to remove the abnormal isoform of scrapie prion (PrP^{Sc}) and scrapie infectivity. Studies were conducted on validated scaled-down models of the manufacturing process steps examined. The first set of studies was conducted using hamster-adapted scrapie agent strain 263K and used Western blot technology. The second set of studies was conducted using mouse-adapted scrapie agent strain ME7 for bioassays utilising C57 Black mice.

RESULTS Infectivity studies on the upstream delipidation process step that is common for both the albumin and immunoglobulin methods of manufacture demonstrated a logarithmic reduction factor (LRF) of at least 2.40 logs.

Substantial LRFs were also demonstrated across the anion and cation ion-exchange steps of the albumin and immunoglobulin processes with both PrP^{Sc} and scrapie infectivity.

Infectivity removal studies on the cold ethanol and depth filtration steps of the CSL intramuscular immunoglobulin (IMIG) process demonstrated a LRF of at least 5.63.

The studies conducted by CSL also demonstrated that infectious scrapie on chromatographic resin following the load-elution steps is removed to below the level of detection from the resins by the cleaning regime used in the manufacturing plant of solvent-detergent and NaOH treatment.

CONCLUSIONS These studies provide assurance that the chromatographic plasma fractionation processes utilised by CSL Bioplasma to manufacture plasma-derived albumin and immunoglobulin possess substantial capacity to remove putative prions during fractionation.

ABSTRACT NO.: 13

Increased Red Cell Susceptibility to Apoptotic Change is Associated with Duration of Packed Cell Storage

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Red cell apoptosis is characterized by loss of lipid asymmetry and cleavage of trans-membrane and cytoskeletal proteins by proteases such as caspase 3. Re-exposure of stored red cells to physiological concentrations of calcium and glucose when transfused may provide some of the signals required for initiation of apoptosis in susceptible cells.

AIM Whether red cells subjected to prolonged storage at 4 °C show increased rates of red cell apoptosis following transfusion has not been studied. In this study we measured susceptibility to apoptotic change after various periods of storage.

METHOD Flow cytometric methods were used to examine changes in annexin V binding to exposed phosphatidylserine, eosin-5'-maleimide binding to band 3 and expression of an adhesion receptor ICAM 4 (CD242, LW protein). Data were analysed using paired tests for non-parametric variables.

RESULTS After 30 minutes of exposure to physiological levels of calcium and glucose we found that the proportion of red cells binding annexin V increased with duration of red cell storage. On day zero 0.25% of calcium and glucose exposed red cells showed annexin V binding which increased 12.6 fold to 3.16% of red cells

after 42 days storage at 4 °C. These changes were enhanced by 1µM ionomycin; on day zero 6.38% of red cells showed annexin V binding which increased 3.3 fold to 21.19% of red cells after 42 days storage at 4 °C. Monoclonal antibody binding to ICAM 4 (LW protein) on red cells exposed to calcium and glucose was reduced with increased duration of red cell storage. Eosin-5'-maleimide binding also varied under these conditions.

CONCLUSIONS Our findings provide some insight into the factors that may influence differences in survival of transfused red cells related to period of storage. Interventions which minimize these changes, and may improve the outcome of transfusion with red cells after prolonged storage, will be the basis of future study.

ABSTRACT NO.: 14

Cryoprecipitate Audit Within Six Centres in New Zealand
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New Zealand Blood Service

INTRODUCTION An audit of apheresis cryoprecipitate transfusions and patients not receiving cryoprecipitate was conducted in 6 centres, covering approximately 84% of the country's cryoprecipitate use.

METHOD Transfusion Nurses Specialists prospectively collected data on cryoprecipitate transfusion episodes for 11 weeks. Where necessary, data was collected retrospectively to make a total of 30 episodes per centre. During each centre's study period, data on patients with fibrinogen levels below 1.0 g/L who did not receive cryoprecipitate was also collected (non-recipients). Two medical assessors reviewed each episode for clinical appropriateness.

RESULTS All centres except one captured at least 30 episodes of cryoprecipitate transfusion. Cryoprecipitate used for fibrin glue (7) and as part of paediatric cardiac bypass surgery protocol (22) were excluded leaving 152 episodes (316 units). Non-recipients involved 86 patients (134 episodes). The mean pre-transfusion fibrinogen level was 1.3 g/L with 46% below 1.0 g/L and 26% above 1.5 g/L. The median fibrinogen concentration of non-recipients was 0.7 g/L. The median number of units transfused per episode was 2.68% of episodes had the right dose (0.5 to 1.5 u/30 kg bodyweight) with underdosing in 24% and overdosing in 8%. Weight-adjusted dose and pre-transfusion fibrinogen level showed no association ($r = 0.19$, $p = 0.434$). Underdosing was associated with more episodes per 48 hours compared with the correct dosing (2.5 vs 1.1) ($p < 0.005$). 18% of cryoprecipitate episodes were considered inappropriate. 27% of non-recipients' episodes were considered inappropriate (cryoprecipitate indicated). Non-recipients showed a similar pattern of fibrinogen levels to recipients.

DISCUSSION Underdosing was more common than overdosing and required more cryoprecipitate transfusions than when the right dose was given. Education is needed regarding the correct dose, the importance of checking pre-transfusion fibrinogen levels and ensuring systems are available for monitoring fibrinogen levels in massive transfusion. It is recommended that Blood Banks ask for the patient's weight before issuing cryoprecipitate.

ABSTRACT NO.: 15

Identification of Proteins that Accumulate in the Supernatant of Platelet Concentrates During Storage

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Platelet transfusion has been implicated in adverse reactions. Soluble factors, including plasma proteins and bioactive molecules released from platelets during storage, are likely to play a critical role. Comprehensive maps of the proteins present in platelet concentrates (PCs) are not available. The aim of this study was to identify the soluble proteins that accumulate in the supernatant of PCs during storage by using a multifaceted proteomics approach.

Irradiated, prestorage leucocyte-filtered pooled buffy-coat PCs in additive solution (T-sol, Baxter) were prepared and stored according to standard blood bank procedures. Samples were collected at days 1, 3, 5, 6 and 7. Platelet membrane integrity was monitored by release of Annexin-V and platelet activation was determined by expression of CD62P. Proteins in the supernatant of PCs were identified by two-dimensional (2D) gel electrophoresis and mass spectrometry, and cytokine antibody microarrays. Cytokines and bioactive molecules were quantitated by ELISAs.

2D-gel maps of PC supernatant showed a number of plasma-derived proteins that appear to undergo alterations and degradation during storage. Several platelet-derived bioactive molecules, such as RANTES and derivatives of platelet basic protein (β -thromboglobulin, CTAP-III and NAP-2) accumulate to high levels. A number of proteins that have not previously been examined in the transfusion setting also accumulate during PC storage. Levels of leucocyte-derived cytokines (IL-1 β , IL-6, IL-8, TNF- α) remained relatively low.

These results provide an expanded view of the proteins that accumulate in the supernatant during storage of PCs and may lead to a greater understanding of the factors that contribute to adverse transfusion reactions and strategies to improve transfusion outcome.

ABSTRACT NO.: 16

Storage Related Increase in Red Blood Cell (RBC) Adhesion to Vascular Endothelium: Effects of Bacterial and Inflammatory Activation

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Physical and biochemical changes that occur to RBCs during storage may increase their adhesion to vascular endothelium thereby potentially impeding blood flow and decreasing oxygen supply in transfusion recipients. Although blood is often required during trauma or surgery where infection, inflammation and/or tissue damage is prevalent, it is unknown whether prior activation of vascular endothelium further affects RBC-endothelial cell (EC) interactions. The aim of this study was to determine whether prior activation of ECs affects adhesion of stored RBCs under conditions of continuous flow *in vitro*.

Human umbilical vein ECs were grown to confluence on gelatin-coated coverslips. Non-leucocyte-reduced, buffy-coat-reduced and leucocyte-filtered RBC products were prepared according to standard blood bank procedures. RBC samples were collected at weekly time points until product expiry. RBCs and ECs were incubated with saline, endotoxin (250 ng/ml) or TNF- α (5 ng/ml) for 4 hours. RBCs were then perfused across the EC monolayer using a parallel flow chamber mounted to an inverted microscope. Perfusion of RBCs was controlled for shear stress and temperature. RBC-EC interactions were recorded using a digital camera attached to the microscope. Activation of ECs was confirmed by immunohistochemical assay using E-selectin and VCAM-1.

Adherence of RBCs to unactivated and activated EC layers increased with product storage time. RBCs from products containing leucocytes were significantly more adherent to the unactivated and activated EC layers in the later stages of storage than RBCs from leucocyte-reduced products. Significantly increased numbers of RBCs adhered to the activated EC layers at Day 1 of storage compared to the unactivated EC layer. Interestingly, a response of this magnitude was not seen at other time point during storage in our model system. These results may lead to greater understanding of the interaction of transfused RBCs with recipient endothelium during inflammation and sepsis and the biological consequences of this adherence.

ABSTRACT NO.: 17

The Impact of Refractory Thrombotic Thrombocytopenia Purpura (TTP) on an Apheresis Unit and Resources – A Case Study

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In August 2004 Mrs X a 38 yr old female with a past history of Systemic Lupus Erythematosus (SLE) and diagnosed with TTP was referred for plasma exchange (PEX). Daily 3 L plasma exchange was commenced and after 6 days Mrs X's condition continued to deteriorate. PEX was increased to twice daily which continued for 36 consecutive days. PEX was gradually tapered off and continued weekly until January 2005. In total Mrs X received 126 PEX'S, 378 litres of cryodepleted plasma (CDP)/fresh frozen plasma (FFP). Conjunctive therapy included high dose steroids, cyclophosphamide, vincristine and rituximab. Rarely do we look at the impact ongoing treatments can have on staff and resources. Staff "cope", resources are stretched, and hopefully the patient recovers, but what are the physical, psychological and cost impacts on a small apheresis unit?

The three full time apheresis staff felt the greatest impact; they were required to work overtime on weekends, creating financial implications for the unit. Emotionally Mrs X became quite attached and reliant on the staff. Staff were faced daily with would Mrs X recover, and also providing emotional support to both Mrs X and her family. She spent up to three hours per day with these staff.

Staff were physically and mentally drained by the time Mrs X had started to recover. Fortunately she has made a full recovery. Staff supported each other, but the continued strain of the prolonged care of Mrs X also impacted on family, perhaps more than staff realised. Mrs X and her family were extremely grateful to the staff and came up with ingenious ways to thank them. As nurses we often say we are "just doing our job" which we are, but to our patients such as Mrs X we are doing a whole lot more and perhaps the "just doing our job line" is what gets us through the day.

ABSTRACT NO.: 18

Assessing and Maintaining Apheresis Competency in the Therapeutic Setting

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Apheresis skills are for the most part learned on the job. With the exception of the Graduate Certificate in Nursing Science (Apheresis Nursing) offered through Adelaide University, there is no formal recognition of the apheresis nurses' ability to perform the myriad of apheresis procedures currently being performed in our hospitals.

AIM In response to questions regarding assessment of new apheresis operators from apheresis units around the country and at the quarterly meetings of the Victorian Apheresis Interest Group, it was decided to attempt to formalise the assessment of competencies within the therapeutic setting, with the intention of creating a package that other units could easily adapt and implement.

METHOD An orientation and assessment package was created, which incorporated information related to apheresis procedures performed and commonly used terminology. Using an established competency based assessment tool (Bondy, K. 1983, 'Criterion-referenced definitions for rating scales in clinical evaluation', Journal of Nursing Education, 22) a checklist of core competencies to be achieved for each type of apheresis device was designed. These core competencies encompass the range of apheresis activity including kit installation, patient preparation, procedure monitoring, post procedure care and adverse reactions. The operator is rated as meeting one of five categories from Independent to Dependent. For each apheresis procedure the new operator is supervised and the checklist completed. A minimum of 15 supervised procedures should be undertaken prior to an initial evaluation. If the operator meets the requirement in the initial evaluation with a minimum 90% of the assessment in the 'Supervised' category they are deemed to be competent and able to undertake subsequent procedures unsupervised or with minimal assistance.

CONCLUSION The Apheresis Core Competency, Orientation and Assessment package has been implemented for new operators in both our units. So far this has proven to be a useful tool for new operators whilst the checklist provides the assessor with a point of reference to use when discussing the procedure with the operator.

ABSTRACT NO.: 19

Successful Mobilisation and PBSC Harvest of Stem Cells following AMD3100 in 4 Paediatric Patients The Children's Hospital at Westmead's Experience

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² Anormed, Canada

We report the successful mobilisation of four patients using granulocyte cell stimulating factor (G-CSF) and AMD3100 who had failed previous mobilisation attempts. AMD3100 is a potent and selective antagonist of the CXCR4 chemokine receptor, and blocks binding of its cognate ligand, encouraging the release of haemopoietic stem cells into the peripheral circulation.

AMD3100 was obtained on a compassionate basis from Anormed and given as per their protocol, with a pre-phase of 10 mcg/kg G-CSF for 4 days and AMD3100 240 mcg/kg given subcutaneously on day 5, 10–11 hours prior to apheresis and then given daily until the required number of cells were collected.

The patients had a variety of diagnoses; Ewings sarcoma, PNET, ALL and Medulloblastoma.

All were male patients aged between 10–17 years – the 10 year old being the youngest patient at the time of administration, to have received AMD3100. Each patient had a history of failed mobilisation requiring a range of remobilisation regimens and between one to four attempts at mobilisation prior to the commencement of AMD3100. All were successfully collected with adequate PBSC's to enable reinfusion after myeloablative chemotherapy.

No side effects of treatment with AMD3100 were noted.

These four cases illustrate that AMD3100, in combination with G-CSF, can achieve successful mobilisation and stem cell collection in previously poorly mobilising patients. This allowed subsequent myeloablative chemotherapy with autologous transplant/re-infusion of PBSC's as a treatment option for these patients.

ABSTRACT NO.: 20

Daily Granulocyte Colony Stimulating Factor (G-CSF) Versus Twice-Daily G-CSF Administration in Mobilisation of Peripheral Blood Stem Cells in Healthy Donors

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AIM Suboptimal mobilisation of CD34⁺ cells into the bloodstream results in longer peripheral blood stem cell (PBSC) collection times and increases the number of harvests required to collect the requested number of PBSC necessary for successful haematopoietic transplantation. The apheresis unit retrospectively analysed the CD34⁺ cell counts pre-harvest for all donors over a 42-month period, to determine if twice daily (BD) G-CSF administration versus daily (D) G-CSF administration yielded greater CD34⁺ cell concentrations.

METHOD From January 2002 to June 2005, 55 allogeneic donors (parent/sibling) and 31 Australian Bone Marrow Donor Registry donors (ABMDR) had peripheral blood stem cells mobilised and collected. All allogeneic donors routinely received G-CSF (granulocyte colony stimulating factor/filgrastin) 5 mcgs/kg/BD (twice per day) and all ABMDR donors received G-CSF 10 mcgs/kg/D (daily) as per ABMDR policy. All PBSC collections commenced on day 5 after 4 days of G-CSF for all donors. PB CD34⁺ cell concentrations were performed by dual platform flow cytometry prior to PBSC collections.

86 healthy donors were harvested, data from two ABMDR donors was omitted from the study due to incorrect G-CSF administration. RESULTS For allogeneic donors the mean pre-collection peripheral blood CD34⁺ stem cell concentration was $92.4 \times 10^6/L$ (range $13\text{--}234 \times 10^6/L$), and the mean pre-collection peripheral blood CD34⁺ concentration in ABMDR donors was $62.6 \times 10^6/L$ (range $14\text{--}202 \times 10^6/L$). 9% of allogeneic donors required more than one apheresis collection to harvest the requested number of PBSC. 21% of ABMDR donors required more than one apheresis collection to harvest the requested number of PBSC.

CONCLUSIONS Twice daily G-CSF administration results in higher peripheral blood CD34⁺ cell concentrations resulting in higher numbers of cells being collected in a single apheresis collection.

ABSTRACT NO.: 21

Low Density Lipoprotein (LDL) Apheresis : A Case Report using LDL Apheresis in Treatment for Familial Hypercholesterolaemia CM Scott¹, B O'Callaghan¹, PK Cannell², GF Watts³

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INTRODUCTION Familial Hypercholesterolaemia (FH) occurs in approximately 1–500 people and is characterized by a raised Low Density Lipoprotein (LDL) plasma concentration, due to reduced uptake by defective LDL receptors. LDL apheresis can selectively remove Low Density Lipoproteins using the Evaflux 5A (Kawasumi Laboratories, Inc., Japan) plasma fractionator.

CASE REPORT A 68 year old female with FH was referred for LDL apheresis; she had a presenting history of Coronary Artery Bypass Graft (CABG), coronary artery stenosis, carotid atherosclerosis and intolerance to anti-lipaeamic medication. On presentation her plasma Cholesterol level was 9.3 mmol/L (range <5.5 mmol/L), LDL 7.1 mmol/L (range <3.4 mmol/L), HDL 1.7 mmol/L (range 1.0–2.5 mmol/L). Following a baseline coronary angiogram and consultation with haematologist, a plan for LDL apheresis treatment was formulated.

METHOD LDL Apheresis is performed using a COBE[®] Spectra[™] apheresis machine; it is a dual needle procedure utilising software version 6.1. Access performed using a 16 G back eye needle via a cubital fossa vein and blood is returned via a 20G forearm cannula. A COBE[®] Spectra[™] Plasma Exchange Kit is modified to divert plasma from waste bag to the Evaflux 5A plasma fractionator (Kawasumi Laboratories, Inc., Japan) at an inlet rate of 40–60 ml/min. The filtered plasma is then diverted back to the patient via a transfer bag. Immediately prior to the procedure 2500 international units of Heparin was administered intravenously. During the procedure Acid Citrate Dextrose- Formula A (ACD-A) anticoagulant was used at a ratio of 1:25. A Manometer was positioned beneath the Evaflux 5A to measure back pressure; pressure throughout the procedures never exceeded 80 mm/Mg (Max pressure 500 mm/Mg). 3500 mls of Plasma can be processed using the Evaflux 5A; this equaled 1.5 plasma volumes for the abovementioned patient. Routine Baseline Blood samples including full blood count, urea and electrolytes, coagulation profile, liver function tests, and lipid profile including (Cholesterol, LDL & HDL). Lipid profile, cholesterol, LDL and HDL levels were taken post. Evaflux 5A was thoroughly flushed with 2 litres of normal saline solution prior to use. Treatment was scheduled for every third week.

RESULTS The patient has had a total of 12 procedures to date. Her mean total cholesterol pre-procedure is 8.2 mmol/L and 2.0 mmol/L post procedure. Average LDL pre-procedure is 7.0 mmol/L and 2.0 mmol/L post. The reduction in her total cholesterol and LDL post procedure indicate that LDL apheresis is an effective treatment option in the treatment of her familial hypercholesterolaemia. Total cholesterol and LDL rebound to elevated levels between procedures; treatment schedule has been altered to two weekly intervals. The patient has reported feeling better than she has in years, clear headed, brighter and has more energy.

ABSTRACT NO.: 22

TRALI: What Do We Know and What Can We Do About It?

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Transfusion-related acute lung injury (TRALI) has been the leading cause of transfusion-related deaths reported to the United States Food and Drug Administration (FDA) for the past 4 consecutive years, and is being increasingly recognized as a leading cause of transfusion-related morbidity and mortality worldwide. It is likely that TRALI is both under-recognized and under-reported—a situation complicated by the fact that there is no definitive laboratory test to confirm the diagnosis, and that, until very recently, there has not even been an international consensus definition of TRALI. In consequence, much of our knowledge about the incidence, epidemiology, pathophysiology, diagnosis and prevention of TRALI is substantially incomplete and/or controversial. Acute lung injury (ALI) unrelated to transfusion has long been recognized in intensive care (ICU) patients and is believed to have a 2-event pathogenesis involving adhesion of primed neutrophils (PMNs) to pulmonary endothelium with subsequent PMN activation by some inducing agent. Although TRALI has traditionally been thought of as having a one-event pathogenesis (passive donor anti-leukocyte antibody interacting with cognate recipient leukocyte antigen with resultant leukocyte activation), evidence has been accumulating that presence of cognate anti-leukocyte antibody/antigen pairs is neither necessary, nor often even sufficient in isolation, to provoke TRALI. Rather TRALI appears to be a multifactorial syndrome, and is likely a true 2-event subtype of ALI, with both recipient predisposition and biological response modifiers (BRMs) generated during storage of blood products also often playing major pathogenetic roles. This session will highlight recent advances in our knowledge of the pathophysiology of TRALI, both antibody-mediated and non-antibody mediated, and the public health implications regarding prevention implied by the two models (which are non exclusive). The recent international consensus definition of TRALI will be discussed and information provided to guide the attendee as to the recognition, investigation and clinical management of TRALI.

ABSTRACT NO.: 23

Lessons Learnt from Scotland

Sandra Gray

Scottish National Blood Transfusion Service, Edinburgh, Scotland, Effective Use of Blood Group

In 2000, we evaluated a programme of clinical effectiveness in transfusion practice utilizing the role of the transfusion nurse specialist (TNS). The study demonstrated that where the TNS was deployed there was a general trend towards improvement in practice, and also showed the importance of adopting a national co-ordinated quality improvement approach.

Following this study, in 2003, NHS Scotland (NHSS) launched a 3-year 'Better Blood Transfusion programme' (BBTP). Central to the BBTP programme, each hospital has access to a transfusion practitioner (TP) as part of a hospital based transfusion team. A standardized educational package has been introduced across NHSS (www.learnbloodtransfusion.org.uk) for all staff involved in the transfusion process, supported by a learner management system (ORASGOLD[™]). Over an 18-month period, approximately 30% of staff have successfully completed the Level 1: Safe Transfusion Practice module. Another key component of BBTP is to provide clinicians with transfusion data so they can review their clinical practice. The development of blood usage data has resulted in a range of reports, which will shortly be disseminated to users in NHSS. The BBTP teams have also instigated over 100 local blood saving initiatives and audits resulting in changes to regional blood ordering and cross-match policies. It has been agreed however, to explore a more nationally co-ordinated approach to blood saving initiatives. The BBTP team also works with the transfusion service to ensure that in times of blood shortages the available stock is managed efficiently.

We have demonstrated that a co-ordinated clinical effectiveness programme with the appropriate support and relevant data can improve transfusion practice. This collaborative approach between BBTP, Scottish hospitals and the Transfusion Service, is seen as the cornerstone of the NHSS response to recent European Union legislation (2002/98/EC/2004/33/EC) and will allow us to benchmark with the wider UK ensuring safe, efficient and effective transfusion practice.

ABSTRACT NO.: 24

The Hong Kong Experience
Lin Che-kit

Hong Kong Red Cross Blood Transfusion Service, Hong Kong
Transfusion medicine has evolved from a mostly laboratory-centred service with a focus on the serological aspects of blood, into a clinically oriented discipline that emphasizes patient care. With the rising cost of quality and safety, increasing number of blood borne pathogens, demand for more and enhanced plasma products, increasing litigations and drive for public sector efficiencies, blood centres are in general facing a serious issue of insufficient resources. For blood centres, the term 'resources' should not be limited to the traditional quantifiable definitions of financial support, infrastructure and investment record, but should also include the considerations of expertise, local social and cultural setting, community understanding and acceptance, and, most of all, a pool of willing and suitable blood donors that is sufficient to provide for the community needs. To maintain blood sufficiency and to ensure blood safety, blood centre has to be creative and be able to practise smarter.

The Hong Kong Red Cross Blood Transfusion Service (BTS) is the only organization responsible for donor recruitment, blood collection and supply of processed blood products to hospitals. Blood products are supplied free of charge to hospitals as the BTS is fully funded by the Government. Funding is allocated on an annual basis and based on previous year's allocation with adjustment. Since the regional economic downturn in 1997, there has been continual cut in the BTS budget. We have to be creative in resources management to maintain sustainability. Notable examples included the contracting out of delivery of blood products to hospitals, contracting out of NAT tests, implementation of automated component processors, introducing phlebotomists to perform venepuncture, etc.

In pursuing efficiency and cost-effectiveness, we have not sacrificed service quality service nor blood safety. The BTS is both ISO9001-2000 and TGA cGMP accredited. Recently, we have also achieved ISO14001 accreditation.

ABSTRACT NO.: 25

Female Genital Graft Versus Host Disease: A Case Study
Catherine Wood

Wellington Hospital (C & CDHB), New Zealand

Survival post allogeneic bone marrow transplantation has improved over the past few years as a result of enhanced supportive care.

Associated with better transplant outcomes are survivorship and quality of life issues. These issues have often been overlooked as the cure of the disease has been seen as the main focus. Over the last year one such quality of life issue has been brought to our attention. Female sexuality in our post transplant patients has been seriously overlooked.

This presentation will tell the story of one of our transplant patients who had problems with lower genital tract graft versus host disease and the changes that have been brought about in our transplant protocols as a result of her story.

ABSTRACT NO.: 26

Haemophilia Care for Nurses in Non-Specialist Centres
Penny McCarthy

Nurse Manager, Ronald Sawers Haemophilia Centre, The Alfred, Commercial Road, Melbourne Vic 3004, Australia

Haemophilia is a rare and potentially life-threatening disease. Meticulous management is important if the patients are to be

spared chronic disability and serious treatment complications. Comprehensive Haemophilia care has been defined as the continuing supervision of all medical and psychosocial factors affecting the person with haemophilia. Services offered by haemophilia treatment centres (HTC) adopting the comprehensive care model include establishing prophylaxis and other treatment protocols, development of psychosocial education and research programmes, maintenance of a patient registry, genetic and reference diagnostic services and orchestration and management of a wide variety of multidisciplinary interventions. (Evatt BL. *et al.*, 2004)

In Australia each state has a designated Haemophilia Centre, which offers a state-wide service and focuses on comprehensive care, providing a wide range of services to people with inherited bleeding disorders, including support for their families.

Each centre has a designated haemophilia nurse, be it full or part time. These specialist nurses are experts in the delivery of quality patient care, and management of costly treatment products.

For nurses working outside these specialist centres caring for a patient with an inherited bleeding disorder can be a daunting task. Advice and support is readily available at all times through the haemophilia centres and early contact is encouraged. By providing the non-specialist nurse with some basic education of inherited bleeding disorders, and providing information on how to access specialist advice and treatment products in a timely manner, we can spare the patient from serious complications of their diseases.

ABSTRACT NO.: 27

Development of a Tool to Measure the Acuity of Haematology and Stem Cell Transplant Patients
Juliija Sipavicius and Angela Booth

BMT Network NSW, Sydney, NSW, Australia

A long standing frustration for haematology/stem cell transplantation (SCT) nurses is the lack of recognition regarding the complex management of these patients. This is due largely to the inadequacy of existing generic tools to quantify specific haematology/SCT patient acuity. Lack of such documentation has led, over time to an erosion of staffing levels, poor skill mix and unsafe nurse/patient ratios. The BMT Network NSW appointed an experienced haematology nurse to develop a tool to demonstrate specific haematology/SCT patient complexity and measure acuity levels. Extensive literature searches revealed no published tools that measure haematology/SCT patient acuity in the past decade. A tool where signs and symptoms of potential adverse effects of treatments and diseases were each assigned scores was developed. Those scores relevant to each patient were totalled, which corresponded to an acuity level. Tool validation was performed by 10 senior haematology/SCT nurses within the BMT Network. Reliability was tested in 4 units. With validity and reliability proven, the tool was tested for 4 weeks within 6 units. Units included SCT, haematology/oncology and paediatric. Analysis of the first 650 patients scored by nurses demonstrated 67% believe the tool accurately reflected patient acuity. 10% found the tool did not accurately reflect patient acuity and in all cases the nurses would have rated the acuity level higher. Findings show that 11% of acutely unwell patients did not receive an accurate acuity score. A major limitation to the study was an inadequate time frame. The study demonstrated the difficulty of assessing the acuity of haematology/SCT patients, even with a tool specifically designed for this complex group of patients. Further refinement and more extensive trialling of this tool is essential to better define and accurately reflect the acuity of these patients if further erosion of staffing levels and degradation of skill mix is to be avoided.

ABSTRACT NO.: 28

Outpatient Based Bone Marrow Transplantation: The Royal North Shore Experience
Cassandra Reid

Royal North Shore Hospital, Sydney, Australia

The Northern Sydney Central Coast Area Health Service is responsible for providing health care to a population of 1.2 million

people. Royal North Shore Hospital also provides a tertiary referral BMT service for patients from the Northern Rivers, mid North Coast and the New England area. As BMT is now the standard of care for an increasing range of haematological malignancies, new strategies for coping with inpatient demand needed consideration. The feasibility and success of managing BMT care in the outpatient setting is well documented in the literature. As our Ambulatory Care unit provides a 7 day a week outpatient chemotherapy and transfusion service and a full time Haematology Registrar clinic on weekdays, the introduction of an early discharge began in late 2003, and evolved into a totally outpatient based transplant program by late 2004. Program eligibility is dependant on criteria such as haematologist recommendation, patient preference, carer availability and proximity to the hospital. By mid 2005 this service is now fully functional with 22% (n = 10) of transplants this year being undertaken in outpatient mode. To date a total of 5 patients (all ABMT) were discharged the day after HPC reinfusion, whilst 14 patients have received totally outpatient based transplants (12 ABMT, 2 mini allograft). All return second daily to the ambulatory care unit for medical and primary nurse review until engraftment. The majority received conditioning therapy of Melphalan alone (n = 13) whilst BEAM (n = 4), Fludarabine/Melphalan (n = 1) and Fludarabine/low dose TBI (n = 1) were also used. 30% (n = 6) of patients required admission for febrile neutropenia (n = 5) and epistaxis (n = 1). Average LOS = 4 days. Patients, their carers and primary nurses were asked to complete a descriptive satisfaction survey. Analysis of the surveys and clinical outcome data demonstrate that, for a select group of patients, outpatient transplantation is a safe and viable alternative to traditional BMT models of care.

ABSTRACT NO.: 29

A Collaborative Project to Improve Platelet Ordering and Administration

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BACKGROUND Platelets transfusions are a fundamental supportive therapy in clinical haematology. Significant issues found in relation to platelets for QEH hospital, IMVS QEH laboratory and ARCBS-SA staff included:

- Haematology-Oncology ward dependent on platelet availability to undertake invasive procedures or to allow patients day leave. Erratic platelet delivery led to procedures delayed and routine platelet transfusions occurring after hours.
- Laboratory staff received numerous phone calls as individual patient's platelets ordered by ward.
- The ad hoc ordering practices flowed through to ARCBS-SA resulting in multiple requests on collection, processing & dispatch staff.
- Clinical staff had limited understanding of platelet production.

METHOD Using clinical practice improvement methodology QEH Laboratory Scientists, Haematology-Oncology ward Clinical Nurse Consultant, Haematologist, Transfusion Clinical Nurse Consultant and ARCBS Transfusion Educator met to identify causes & trial solutions. Strategies focussed on improving communication to laboratories and education to alleviate identified knowledge gaps.

AIMS In QEH Haematology/Oncology Ward over 3 months:

- Prophylactic platelet transfusions are available & transfused before 5 pm 90% of the time. (stretch goal 95%).
- Platelet requirements are communicated to the Hospital Blood Transfusion Service daily (Mon-Fri) by 1000 hrs 90% of the time (stretch goal 95%).

RESULTS

- Significant decrease in after-hours routine platelets transfusions (Stretch goal >95% achieved).

- No procedures delayed due to platelet unavailability.
- Platelets requirements communicated to laboratory staff by 10 am 100% of the time (fewer phone calls and stress reduced).
- Ordering practices spread to other blood products & clinical areas.
- Improved communication between clinical laboratory & ARCBS.
- Improved clinician knowledge.

CONCLUSION A review of platelet ordering practices achieved significant gains for patients, QEH Haematology-Oncology ward, QEH-IMVS laboratory staff and ARCBS-SA.

Staff knowledge of platelet issues including production and administration has been improved by education package.

Aspects of these strategies may be implemented in other institutions.

ABSTRACT NO.: 30

A Transfusion Reaction with a Tingle

Karen O'Shea

Royal Hobart Hospital (RHH), Hobart, TAS, Australia

Transfusion reaction reporting at the RHH was historically low approximately 3% year, which was also consistent with the literature stating that transfusion reactions are under reported. An education strategy was developed at the RHH to improve knowledge on symptoms of transfusion reactions. The strategy involved extensive education sessions, lanyard cards for all medical and nursing staff and the development of a new transfusion reaction reporting form. The transfusion reaction lanyard cards highlight the signs and symptoms of the more common transfusion reactions and act as a guide to immediate management. It emphasises the need for nursing staff to report changes in the condition of a patient during and/or after a transfusion as the list for reactions is diverse.

A 79 year old lady was admitted to the ambulatory care day unit with a history of cancer of the anal canal. Her pre-transfusion Haemoglobin was 8 g/L with no other clinical indications documented. The treatment planned was for transfusion of three units packed red cells. Two units were transfused uneventfully. During transfusion of the third unit the patient developed symptoms of tingling in both arms and hands, and also decreased hearing. Blood was infused via a blood warmer due to the presence of an anti-Lewis b antibody. Observations were stable except for a temperature rise of 36.3–37.3 °C. Baseline observations of Pulse 82 per minute, Respirations 16 per minute and Blood Pressure 116/63 mmHg at 1505 hrs were stable with the reaction at 1715 hrs with observations of Pulse 85 per minute, Respirations 20 per minute and blood pressure 112/60 mmHg. The laboratory investigated the transfusion reaction and found no evidence of red cell incompatibility or haemolysis due to the use of the blood warmer. Her pre-transfusion calcium was 2.07 mmol/L Ca (adj) Normal Range 2.10–2.55 mmol/L. A post transfusion calcium level could not be obtained due to discharge of the patient. The patient quoted to the medical officer that "many years ago I had a transfusion reaction with tingling all over that lasted minutes". Unfortunately no documented evidence exists to review this.

On consultation with the Australian Red Cross Blood Service web page they were able to support our judgement that this may have been a citrate reaction to the transfusion.

ABSTRACT NO.: 31

Best Practice in Transfusion Specimen Labelling – Declaration of Positive Patient Identification

Trudi Verrall

Children, Youth and Women's Health Service, North Adelaide, South Australia, Australia

Inadequate patient identification during the specimen labelling process instigated involving consumers to check their details on the tube and form.

Evidence reveals that incorrectly labelled Transfusion Specimen can be fatal. Miscollection of blood is commonly referred to as 'Wrong Blood in Tube' (WBIT). A WBIT occurs when identical patient details are found on the tube and form however it is not that patient's blood in the tube. WBIT are detected when comparing current blood group results with an historical blood group and they differ. Majorities of WBIT go undetected as a high proportion of patients do not have an historical record.

As a result of a fatal incident due to mis-labelling of blood, a recent Australian Coroner recommended that all organisations undertaking Pre-Transfusion testing follow the Australian New Zealand Society of Blood Transfusion (ANZSBT) Specimen Labelling Guidelines. These guidelines state the collector must sign a declaration verifying positive patient identification at the time the blood is taken. It is also recommended that a second witness sign verifying the patient's identity. The coroner stated that where possible, especially with the young and aged that a relative or carer be involved in identifying the patient.

Reviewing the organisations pre-transfusion specimen labelling standards revealed that they did not meet the ANZSBT requirements regarding a declaration of positive patient identification by the collector or a second witness. Wrong blood in tube incidents and a recent root cause analysis in the hospital exposed that a problem did exist.

A small project team, involving a consumer was formed and Clinical Practice Improvement (CPI) methodology was utilised to resolve the declaration of positive patient identification when collecting blood samples in an emergency service. A mission statement aiming to eliminating miscollection and mislabelling was created. The specimen labelling process was studied and a detail flow chart developed highlighting areas of concern. Team voting revealed the inadequate transfusion request form was a priority.

Interventions such as a new sticker with a collectors and second witness declaration of patient identification was developed and placed on the original form. Staff education occurred and posters were developed explaining the new process. Each new intervention followed the CPI, Plan, Do, Study, Act cycle and interventions where adapted and changed according to the results.

Results were tallied weekly and Patient Identification Declaration by the collector was completed 100%, except for twice when it dropped to 97%. No WBIT have been detected during the 28 week trial and a second witness signature has been provided 74% to 100%.

Staff have embraced the practice change with consumers stated they are comfortable being included in the labelling process.

ABSTRACT NO.: 32

Transfusion Practice Improvement Strategies, Raising Awareness and Knowledge of Transfusion Issues in Victoria and Tasmania Nadine Gilby¹, Marcus Kennedy², David Beilby³, Karen Botting¹, Neil Boyce⁴, Peter Bradford⁵, Wallace Crellin, Twanny Farrugia, Sue Graham⁶, Chris Hogan², Annette Holian⁷, Geoff Magrin⁷, Alison McMillan¹, Larry McNicol⁸, Tania Nallathambiy¹, Miles Prince⁹, Helen Savoia¹⁰, Carole Smith⁸, Lisa Stevenson¹¹, Elizabeth Wilson⁵, Erica Wood⁴

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AIM The Better Safer Transfusion (BeST) Program of the Department of Human Services Victoria, aims to improve transfusion safety and practice in hospitals.

The goal is to achieve practical and sustainable improvements in four main areas:

- (1) improve awareness and knowledge of transfusion practice within hospitals;
- (2) implement appropriate and best practice for clinical decision making and blood administration;
- (3) develop and implement a state-wide haemovigilance system; and
- (4) engage and support the private and rural sectors.

METHOD The program is overseen by a multidisciplinary departmental advisory committee, supported by four expert working parties and a secretariat.

Improvement of knowledge and safety of transfusion within health services is through:

- support of existing transfusion nurse roles
- support and promotion of the transfusion practice course; an on-line course accredited as equivalent to a graduate certificate
- audits encouraging review of practice against evidence-based guidelines have been sent to health services
- development of a haemovigilance system
- information on BeST program initiatives has been discussed in regional forums
- improvement practice tools and information for transfusion have been made available through the BeST website at: <http://www.health.vic.gov.au/best>

RESULTS

- on-line transfusion practice course continues with national & international participants in 2005, resulting in knowledgeable transfusion practitioners (medical, scientific, nursing) who can lead transfusion improvement
- 19 Transfusion Nurses working in Victorian & Tasmanian public hospitals
- education of health professionals by Transfusion Nurses within health services
- development of consumer education material
- response to transfusion audits (initial audit looking at storage & handling of blood products had >80% response rate)

CONCLUSION Enthusiastic participation in transfusion improvement initiatives in Victoria and Tasmania continues to improve transfusion practice and patient outcomes. Building on the initial success of the Blood Matters program, rural and private hospitals are now also involved in this program.

ABSTRACT NO.: 33

Adapt, Adopt or Augment: Strategies for Change in the Blood Service?

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² CO-LEARNZ, Auckland, New Zealand

Safety and supply are two issues that universally concern Blood Services throughout the world, whatever their configuration. Measures to safeguard against the risks of HIV, Hep C and vCJD have required radical changes in many Blood Service organisations in recent years. In New Zealand, risk management was a major driver leading to the inauguration of a 'vein-to-vein' National Blood Service (NZBS) in 1998. In Australia the change imperatives were different, but the challenges nonetheless pressing. Prior to the establishment of the Australian Red Cross Blood Service (ARCBS) in 1996, each Australian State and Territory had an autonomous Red Cross Blood Service. Our purpose in presenting this paper is to discuss ways in which strategies for making radical changes were identified, appropriate to the specific contexts of change within the two countries. We discuss the merits and benefits of the strategies adopted. We believe the organisational development lessons learnt Down Under have wide application in Blood Services undergoing change around the world. There are, we believe, very real merits in adapting, adopting

or augmenting other Services' experiences. Successful choice of change strategies, however, for the design, development and implementation of service and organisational change, depends on appropriate analysis of the tasks and challenges being confronted, and the careful marrying of theoretical principles with effective best practice in both clinical and organisational development areas.

ABSTRACT NO.: 34

A Retrospective Audit of Irradiated Component Use in New Zealand
Christopher Corkery, Richard Charlewood

New Zealand Blood Service, Hamilton, New Zealand

BACKGROUND Transfusion Associated Graft vs. Host Disease (TA-GvHD) is a fatal complication of blood transfusion. Practically, there is no treatment for TA-GvHD. The disease is prevented by providing irradiated components to at-risk patients. These patients are treated by a diverse group of health professionals. The challenge is to ensure that such patients always receive irradiated blood components.

AIM To ascertain if patients with an absolute indication for irradiated components (as per ANZSBT Guideline) received only irradiated components. To assess whether patients who have an irradiated components protocol in place have appropriate diagnoses.

METHOD Transfusion Nurse Specialists at six main centres across New Zealand retrospectively collated lists of patients with absolute indications for irradiated components for 2004. The clinical data included the diagnosis and treatment dates. Sources included case mix analysts, paediatricians, haematologists, pharmacists and blood banks. The units transfused to these patients were sourced from Progesa.

INTERIM RESULTS 498 patients were identified as having attended hospital in 2004 with an indication for irradiated components. 294 (59%) received transfusions. 4580 units in total were transfused of which 341 (7%) were not irradiated. 71 (24%) transfused patients received a mean of 4.8 non-irradiated units (range: 1–34). The diagnosis most strongly associated with patients receiving non-irradiated components was Hodgkin's Disease followed by Aplastic Anaemia and purine analogue therapy (23 and 20 patients respectively). Of the 338 patients with irradiated protocols in place, 68% were absolute indications and 25% were probable indications as per ANZSBT guidelines. Some neonatal units had standing policies to provide irradiated components to all patients. No cases of TA-GVHD were reported in 2004.

CONCLUSION A significant proportion of patients with absolute indications for irradiated components received unirradiated components. Where protocols are in place, indications were mainly appropriate.

ABSTRACT NO.: 35

Iron Metabolism and Assessment of Iron Stores

John K Olynyk

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Type 1 hereditary hemochromatosis is a common disorder of iron overload occurring in individuals homozygous for the C282Y *HFE* gene mutation. It can be a progressive and fatal condition. Early detection and phlebotomy prior to the onset of cirrhosis can reduce morbidity and normalize life expectancy. It is readily identified through biochemical testing for iron overload using serum transferrin saturation and genetic testing for C282Y homozygosity. Recent advances in non-invasive magnetic resonance imaging have substantially improved the diagnosis and risk stratification of patients with iron overload. General population screening has been waived in preference to targeting high-risk groups such as first degree relatives of affected individuals and those with clinical features suggestive of iron loading. This screening strategy is likely to continue until uncertainties regarding the natural history of the

disease, age-related penetrance, and management of asymptomatic individuals are clarified.

ABSTRACT NO.: 36

Iron Nutrition in Pregnancy and Early Life

Maria Makrides

Child Health Research Institute, North Adelaide, South Australia

AIM Iron deficiency is a relatively common problem in pregnancy in both developed and developing countries. In industrialised countries there is some debate about usefulness of routine iron supplementation in pregnancy. The aim of the Adelaide Mothers' and Babies' Iron Trial was to investigate whether routine low dose iron supplementation in pregnancy has beneficial effects for the mother and child.

METHODS Randomised controlled trial of a daily iron tablet (20 mg) vs placebo from 20 weeks gestation until birth. Primary outcomes included maternal iron status at the end of the pregnancy and at 6 months post partum, as well as childhood IQ at 4 years of age. Other outcomes included pregnancy outcome, maternal health and well-being and childhood behaviour at 4 years of age.

RESULTS 431 women (215 in the control group and 216 in the iron group) were recruited from the Women's and Children's Hospital, Adelaide. The prevalence of iron deficiency anaemia (IDA) in the iron group (3%) was lower than the control group (11%). By 6 months post-partum, the frequency of IDA did not differ between the two groups but women in the iron group had less iron deficiency compared with control. There were no differences between the groups in pregnancy outcome, or any indices of maternal mood and well-being. Similarly the mean IQ and mean behavioural scores of children born to mothers in the iron and control groups did not differ. However, the percentage of children with abnormal total behavioural scores was higher in the iron group compared with the control group.

CONCLUSIONS Although routine iron supplementation in pregnancy was associated with improvements in maternal iron status and a lower risk of iron deficiency, there were no improvements in indices of maternal well-being or childhood development in this well-nourished population.

ABSTRACT NO.: 37

Every Evening at The Bun Shop

Robyn Barlow

ARCBS, Sydney, NSW

Characters:

Sir Ronald Fisher- mathematician and geneticist

Dr Robert Race- red cell serologist

Scene I: The Bun Shop – Cambridge, 1943

Corner, Downing & Corn Exchange Streets

Ronald Fisher and Robert Race are seen, as they are every evening, discussing the latest interpretations of Race's serological findings.

They are drinking beer. Fisher is writing on a sheet of paper.

Scene II: Caius College – Cambridge University

Fisher has returned to his rooms in College. He works for two hours, studying the sheet of paper. He then retires to bed.

Scene III: The Bun Shop

At Bun Shop time the next evening, Fisher shows Race the sheet of paper. On it is revealed a quantum leap towards safer blood transfusion and a defining step in the prevention of a devastating disease.

ABSTRACT NO.: 38

Novel Approaches for Blood Group Genotyping

Jill R Storry

Blood Centre, Lund, Sweden

Studies show that approximately 1–2% of transfused patients will produce one or more alloantibodies to antigens absent on their RBCs but present on the transfused RBCs. The incidence is greatly

increased in those patients who are regularly and/or multiply transfused and it is estimated that 35–50% of all transfused patients with sickle cell disease become alloimmunized. Additionally, alloimmunization may occur during pregnancy with potentially fatal consequences for the unborn foetus. Aside from any morbidity caused by the antibody, the provision of antigen-negative blood often requires the screening of hundreds of RBC units to find compatible blood. Current practice in many blood centres and transfusion services is to maintain a limited inventory of RBCs that have been typed for common antigens in the Rh, Kell, Duffy, MNS and Kidd systems. This testing is costly and the availability of test reagents is dwindling.

The molecular bases of 28 of the 29 blood group systems have been elucidated. Techniques to detect genetic polymorphisms that encode blood group antigens have been developed and refined. These are used primarily for the resolution of weak or unusual antigens, as an adjunct to serological investigations of samples from previously alloimmunized patients and for genotyping foetal DNA to determine at-risk fetuses. These mostly manual techniques are not practical for mass screening such as donor testing.

Most blood group antigens are encoded by single nucleotide polymorphisms (SNPs) and have the potential to be analyzed by more high-throughput technology e.g. microarray. Several groups have been exploiting the use of SNP microarrays for the rapid detection of blood group polymorphisms. Although in their infancy, initial results look promising and microarray platforms look likely to be the way of the future for genotyping donor blood.

ABSTRACT NO.: 39

Toward *In Vitro* Production of Blood Products

Lars Keld Nielsen

Australian Institute of Bioengineering and Nanotechnology, The University of Queensland, Brisbane, QLD, Australia

Transfusion of blood products is a common, life saving medical procedure used in trauma and surgery as well as in treating blood clotting disorders, cancer, sickle cell anaemia, and during organ transplantation. The Australian Red Cross Blood Service is responsible for the collection, processing and testing of almost one million blood donations a year from healthy volunteer donors. The challenges and costs of sustaining this very successful system are ever increasing. The potential contamination of blood products by adventitious agents is driving up costs for performing detailed donor interviews and testing of products, while reducing the number of eligible and willing donors.

Production of blood products in bioreactors mimicking haematopoiesis is an emerging alternative strategy. It is already technically possible to produce fully functional red blood cells (RBC) and neutrophils from haematopoietic stem cells and technologies are emerging to replace haematopoietic stem cells with embryonic stem cells, thus providing an abundant, well-characterized starting material. Irrespective of starting material, however, a substantial hurdle to *in vitro* production is the cost and logistic of producing meaningful numbers of cells. A single unit of red blood cells contains 2 trillion cells and with current technologies it would require 20 days operation of a 2,000 L bioreactor to produce a single unit of RBC. In scale, this corresponds to the production of 1 kg of monoclonal and an expected price of \$1 million per unit.

This talk will highlight some of the engineering challenges as well as some solutions illustrated with examples from our own work on both RBC and neutrophil production.

ABSTRACT NO.: 40

The Introduction of a National Haemovigilance Programme

Simon Benson

New Zealand Blood Service, Auckland, New Zealand

On 1 May 2005 following a four-month pilot at three North Island hospitals, New Zealand Blood Service (NZBS) introduced its national haemovigilance programme.

The programme, which embraces Council of Europe requirements, is modelled on similar schemes in the UK and Eire and aims to capture data on the prevalence of all types of transfusion-related adverse events not only so-called transfusion reactions.

The reporting process centres on hospital 'Transfusion Safety Officers' (TSO), nominally a hospital's blood bank charge scientist, who is responsible for ensuring all events are reported.

Events are initially reported using a dedicated haemovigilance form. An accompanying 'user guide' provides information for completing the form, definitions of types of events and anticipates some frequently asked questions. Completed forms are then submitted to NZBS National Office where they are entered into a Microsoft Access™ database for subsequent analysis. If further investigation is indicated additional event-specific forms are sent to the hospital concerned.

From the start of the pilot until 30 June 2005, 112 reports of events have been received from 22 hospitals. Whilst the majority of events have been non-haemolytic febrile transfusion reactions or allergic reactions (58% and 30% respectively) other events include 'transfusion associated circulatory overload' (TACO; 4%), 'incorrect blood component transfused' (IBCT; 4%) and possible 'transfusion associated acute lung injury' (TRALI; 3%).

The Haemovigilance Programme appears to have gained acceptance within the New Zealand transfusion sector with 16 of the country's 21 District Health Boards represented in the reports received. However there is still much effort required in raising awareness of the programme particularly among nursing, medical and quality personnel. Regular communication with these groups, hospital transfusion committees, newsletters, the NZBS website and ultimately release of an annual report are all tools that are being, or will be used, to raise the profile of haemovigilance in New Zealand.

ABSTRACT NO.: 41

The NZBS 'DHB Clinical Oversight Programme'

Simon Benson

New Zealand Blood Service, Auckland, New Zealand

On 1 January 2005 New Zealand Blood Service (NZBS) introduced its District Health Board (DHB) 'Clinical Oversight Programme'.

NZBS has statutory responsibility for collection and distribution of blood being appointed by Parliament to ensure DHBs maintain efficient blood banking systems.

In discharging this responsibility NZBS has traditionally supported DHB transfusion activities with clinical audits, site visits and regional blood bank meetings. However delivery was inconsistent from region to region and not nationally coordinated.

To gauge what support was expected consultation with blood banks was undertaken, responses shaping the eventual proposal for a formalised 'clinical oversight programme'. Subsequent feedback showed support for the proposal providing a clear mandate for its introduction. The proposal was also endorsed by International Accreditation New Zealand (IANZ; the accrediting agency for medical testing laboratories) in line with requirements of standard NZS/ISO15189.

The programme is now well established. DHBs participate by formal agreement with NZBS and so far only one DHB has chosen not to participate.

12 site visits have been performed with 7 sites receiving corrective action requests. Copies of site visit reports are forwarded to IANZ and issues raised in them have been used by IANZ during their assessment visits to other blood banks.

Regular regional meetings are held by the four main NZBS centres. Attendance particularly by the smaller, geographically isolated blood banks is inconsistent, as staff cannot be spared to attend. However IANZ expects blood banks to attend at least 2 (out of 3) meetings per year which they will monitor. Finally 20 sites are scheduled to receive their (biannual) clinical audits.

It is hoped the NZBS 'DHB Clinical Oversight Programme' offers DHBs the support required to provide transfusion activities which are consistently practiced across the country, which meet the expectations of NZBS and IANZ and which conform to best international practice.

ABSTRACT NO.: 42

First Yield for HIV-1 NAT in the Australian Blood Donor Population – A Repeat Donor with Acute HIV-1 Infection

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² Serology Laboratory SEALS, Prince of Wales Hospital, Sydney, NSW, Australia

³ Centre for Immunology, St Vincent's Hospital, Sydney, NSW, Australia

BACKGROUND In June 2000, the Australian Red Cross Blood Service (ARCBS) implemented nucleic acid amplification testing (NAT) of all blood donations for HIV-1 and HCV to reduce the residual risk of transfusion-transmitted viral infections.

CASE REPORT On 7 May 2004, a 55-year-old donor made a blood donation that was negative on routine screening for anti-HIV-1/2. However, the donation was within a minipool of 24 that tested positive for HIV-1/HCV RNA (Chiron, Procleix TMA HIV-1/HCV Multiplex Assay). Testing of individual donations resolved the pool to this donor. The donation was subsequently confirmed HIV-1 RNA positive and HCV RNA negative (Chiron, Procleix TMA HIV-1 and HCV Discriminatory Assays). A follow-up sample collected 7 days post index donation remained anti-HIV-1/2 negative and HIV-1 RNA positive. Seroconversion was documented on the testing of a sample collected 12 days post index donation.

At the time of donation, the donor reported being in good health and had responded negatively to all ARCBS risk factor screening questions. Two days post index donation, the donor became mildly unwell, with lethargy, cough and nasal congestion. These symptoms persisted for one week and resolved spontaneously. The donor subsequently remained well and was not commenced on anti-retroviral therapy.

Sequencing of the HIV-1 reverse transcriptase and protease genes from a follow-up sample collected 124 days post index donation demonstrated no drug resistance mutations and identified the HIV-1 subtype as CRF01_AE.

The donor had visited Thailand during the period of March 2004 and April 2004 for the purpose of elective surgical treatment and had returned to Australia 11 days prior to the index donation. The donor specifically denied risk behaviour in Thailand, including sexual activity or injecting drug use, suggesting the possibility of iatrogenic transmission.

CONCLUSION This case represents the ARCBS' first NAT yield for HIV-1 and demonstrates the improved safety of the blood supply following the introduction of NAT.

ABSTRACT NO.: 43

Reducing the Risk of HBV Transmission

Richard Charlewood, Peter Flanagan, Melanie Dravitsky, Helen Hollis, Rebecca Horder

New Zealand Blood Service, Auckland, New Zealand

AIM Hepatitis B infection is endemic within New Zealand. Blood donations are tested for HBsAg (Abbott Prism CLIA). Approximately 160 000 donations are transfused each year with 1–2 reports of probable transfusion-transmitted Hepatitis B received annually. The feasibility and likely benefits of additional Hepatitis B testing were investigated using either HBV DNA (Chiron Ultrio assay) or Hepatitis B Core antibody (Abbott Prism CLIA).

METHOD 10 000 donations were tested for HBV DNA (single donation testing). Reactive specimens were investigated with anti-HBc, anti-HBs (IMx Abbott) and real-time HBV DNA PCR (ESR, NZ). 10 000 separate donations were tested for anti-HBc with repeat reactive donations tested with anti-HBs and an independent anti-HBc assay (Murex). HBV DNA testing was undertaken on anti-HBc positive donations with anti-HBs less than 100 IU/L.

RESULTS 6.8% of donations were anti-HBc reactive. 64% of anti-HBc reactive donations had anti-HBs greater than 100 IU/L. 6.3% had no detectable anti-HBs. 1 anti-HBc reactive donation showed low level Ultrio reactivity. Anti-HBs was 8.6 IU/L with

negative Ultrio HBV discriminatory and repeat Ultrio testing. To date, 7820 donations have been tested by Ultrio. 13 donations were reactive but only 1 was reactive in both discriminatory and confirmatory assays. This donation was anti-HBc strongly reactive and anti-HBs was less than 10 IU/L. Only 1 stored aliquot from 12 previous donations tested was Ultrio reactive but HBV discriminatory assay was negative. Lookback on the recipients of components from this donor's donations found 7 of 15 recipients alive. None of 4 tested had markers of past Hepatitis B infection.

CONCLUSION The high level of anti-HBc reactivity suggests this would not be an appropriate screening test to reduce the risk of transfusion-transmitted HBV in New Zealand. Further study is needed to improve understanding of the significance of occult HBV infection and the utility of the Ultrio assay to reduce the risk of HBV transmission.

ABSTRACT NO.: 44

Transfusion Risk Perceptions of the Australian Public
Tessa Hillgrove¹, Ross Savvas¹, Kathleen Doherty¹, Margaret Dorsch²

¹ Australian Red Cross Blood Service, Adelaide, SA

² Australian Red Cross Blood Service, Canberra, ACT

AIM Transfusion risk perceptions of the general public have been investigated in Canada and the USA. However, to the best knowledge of the authors, there have been no studies investigating how the Australian population perceives the safety of transfusion.

METHODS A questionnaire was adapted for the study based on that developed by R.D. Davenport and D. Henrard from the University of Michigan, USA (unpublished, used with permission). The questionnaire included questions to explore both elective and emergency surgery transfusion preferences, risk perceptions, personal experience with blood transfusion and adverse transfusion outcomes, blood donor status, and demographic information. The questionnaire was distributed by mail to a proportionally stratified random sample of 3014 adults drawn from the Australian electoral roll during October–December of 2003. Data was analysed with chi-square tests using the Stata8 software package.

RESULTS When asked to rate their level of concern with the safety of blood transfusion today, 69% of respondents stated a low level of concern. The majority of respondents (60%) were able to correctly estimate the risk of contracting a serious viral infection as very low (1 in 100 000 or 1 in 1000 000). When asked about transfusion preference during elective surgery, similar proportions preferred autologous blood (43%) and homologous blood (42%), 13% preferred a directed donation, and less than 3% said they would not accept a blood transfusion at all. For a transfusion in a medical emergency, 60% of respondents indicated they would prefer blood from the "blood bank", with 40% seeking alternatives. These results can be directly compared to those of the University of Michigan study.

CONCLUSION Relative to the general public of the USA, the Australian public has lower levels of concern about blood safety. This is reflected in the higher proportion willing to accept a homologous transfusion in elective and emergency surgery.

ABSTRACT NO.: 45

Neonatal Alloimmune Thrombocytopenia – A 5 year Victorian Experience

Marija Borosak, Rhonda Holdsworth, Erica Wood
Australian Red Cross Blood Service, Melbourne, Victoria

AIM To review cases referred for investigation of neonatal alloimmune thrombocytopenia (NAIT) in Victoria from 2000–2004. NAIT is a major cause of morbidity and mortality in the thrombocytopenic foetus and neonate. It results most commonly from parental human platelet antigen (HPA) incompatibility.

METHODS Retrospective review of database results (HPA, HLA, crossmatch investigations etc) and clinical records.

RESULTS Over the last 5 years we have seen increased numbers of referrals for investigation of NAIT: n = 14 in 2000, 23 in 2001, 26 in 2002, 45 in 2003 and 40 in 2004. Confirmed cases averaged 7/year,

corresponding to an incidence in Victoria of 11/100,000 live births. Other cases were unable to be confirmed or excluded because of inadequate specimens, confounding HLA antibodies, or complex clinical scenarios including other potential diagnoses. Intravenous immunoglobulin (IVIg), infused to the mother antenatally, is used to augment the foetal platelet count prior to delivery and reduce the risk of intracranial haemorrhage. From March 1999 to June 2005, requests for 161 doses of IVIg were made to treat NAIT (0.4% all requests for IVIg), corresponding to 0.75% grams IVIg issued, with the average antenatal dose being 58 grams on each occasion. Many of these pregnancies also required HPA-matched intrauterine platelet transfusions and neonatal platelet and IVIg support.

CONCLUSIONS Our referral centre provides coordinated testing, specialised platelet therapy and clinical advice for the management of these complex cases. Referrals for labour intensive and time-consuming NAIT investigation and support have increased in recent years. Optimal therapy remains to be defined. Development of a national registry would further the knowledge about pathogenesis and management of NAIT and may provide new insights into the course and prognosis of the disease.

ABSTRACT NO.: 46

The Mystery of the Missing Miltenberger – What is the Basis of Non-B-A-B Mi(a)-Pos Phenotypes in Asia?

Robert Lewis Flower

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AIM The antigens of the MNS system are located on erythrocyte glycoproteins. GpMur (Mi III) results from gene conversion. A GPA sequence is inserted into GPB, leading to reactivation of a splice site and expression of a hybrid of GpA and a GpB pseudoexon. Other variant MNS glycoproteins such as GpHop, GpBun, or GpHF (Mi types IV, VI or X) have been thought to be low frequency with GpMur the basis of Mi(a)-pos types in Asia. A-B crossovers, such as GpHil (MiV) have not been thought to contribute to Asian Mi(a)-pos phenotypes. The aim of this study was to study the frequency of B-A-B gene conversions and A-B crossovers in specimens serologically defined as Mi(a)-pos.

METHOD A PCR for detection of B-A-B conversions and a PCR designed by Shih et al for detection of A-B crossovers were optimised. **RESULT** A 145 bp product consistent with B-A-B gene conversion was detected in 80% (16/20) of serologically Mi(a)-pos samples from Vietnamese subjects. B-A-B gene conversion was detected in 3% of Australian Chinese tested. A-B crossovers were detected only in two individuals of GpJL (Mi XI) phenotype.

CONCLUSION Nadarajan *et al* found 60% of Mi(a)-pos Malaysian Chinese were B-A-B pos but that less than 50% of Mi(a)-pos Malays were B-A-B pos and while up to 2% of Indians were Mi(a)-pos none were B-A-B PCR-pos. Lin has reported that 60% of serologically Mi(a) pos Taiwanese are of types associated with B-A-B conversion. For Australian Chinese the proportion that was B-A-B positive was similar to those reported by Nadarajan and Lin. For Mi(a)-pos samples from Vietnamese the 80% B-A-B pos frequency was a higher than for other Asian ethnic groups. Whether some of these Mi(a)-pos individuals react with anti-Vw found in some batches of intravenous immunoglobulin is a question currently under investigation.

ABSTRACT NO.: 47

First Example of the Para-Bombay Phenotype in an Australian Proband is Encoded by a Novel *FUT1* Allele

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AIM The rare H-deficient Bombay and para-Bombay RBC phenotypes are characterised by sporadic mutations in the *FUT1* gene

in different populations. Samples were investigated from a pregnant Australian female with a suspected para-Bombay phenotype. Our aim was to characterise the RBCs and to identify *FUT1*, and possibly *FUT2*, mutations responsible for the phenotype.

METHODS Standard serological methods were used, including adsorption/elution of human anti-H. Genomic DNA was isolated from the proband and a cord sample drawn at delivery. Routine *ABO* genotyping assays and sequencing analyses of *FUT1* and *FUT2* were performed.

RESULTS The proband's RBCs typed as O, Le(a-b+), and were H- by direct testing with human, monoclonal and lectin anti-H. However, an eluate from her RBCs following adsorption contained anti-H that was reactive with papain-treated RBCs only. Analysis of *FUT1* in the proband revealed homozygosity for a mutation 661C > T, predicted to change Arg221Cys. No mutations were identified in her *FUT2* sequence. *FUT1* from the cord sample demonstrated heterozygosity: 661C/T, and this sample was also heterozygous for a common silent mutation in *FUT2*, 357C > T. The *ABO* genotype of both proband and cord sample was O¹O¹.

CONCLUSIONS We report the identification of a para-Bombay phenotype in an Australian woman that is due to a novel mutation in *FUT1* (661C > T). Weak H antigen was detected on her RBCs. This might be due to low activity levels of a fucosyltransferase encoded by the mutated *FUT1*, but it is more likely due to adsorbed soluble H antigen produced by the normal fucosyltransferase encoded by *FUT2*.

ABSTRACT NO.: 48

Alloimmunisation in the Setting of Apheresis Single Donor Platelets. A Review of Haematology Patients Undergoing Chemotherapy or Bone Marrow Transplantation

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INTRODUCTION Foreign antigen exposure may result in alloimmunisation leading to immune platelet transfusion refractoriness with requirements for specialised platelet products.

AIMS Study aims were to assess the alloimmunisation rate in haematology bone marrow transplant (BMT) and chemotherapy patients, the impact of alloimmunisation on transfusion requirements and address the value of continued routine platelet antibody (PAI) screening in the setting of leucodepleted apheresis-collected single donor platelets (PSD).

METHOD Routine PAI screens performed over 33 months on BMT and chemotherapy patients for pre-emptive detection of alloimmunisation were reviewed. Patients were classified using clinical and laboratory information.

RESULTS The audit identified 283 patients; 148 BMT patients (autologous 71, allogeneic 77) and 135 chemotherapy patients. 43 patients had positive PAI tests at any stage; 21 BMT and 22 chemotherapy patients. Alloantibodies were detected in 40 patients (alloimmunised 22 patients, transient 18 patients) and autoantibodies in 3. Alloantibody specificity was anti-HLA (20 patients) with anti-HPA/platelet glycoprotein detected in 2. Three times as many female patients were alloimmunised. Alloimmunisation was detected earlier in females, usually on first PAI screen and before transfusion. The overall alloimmunisation rates were 8.8% in BMT patients (allogeneic 13.0%, autologous 4.2%) and 6.7% in chemotherapy patients. Risk factors for secondary alloimmunisation were present in 86.4%; 13 of 17 alloimmunised women had children, 2 of 4 nulliparous women and 4 of 5 male patients had extensive transfusion histories. Alloimmunised patients required specialised platelet products including crossmatched, HLA, directed donations or frozen-thawed PSD. Four BMT patients lost evidence alloimmunisation post-engraftment allowing return to standard PSD.

CONCLUSION Alloimmunisation continues to occur despite the use of leucodepleted transfusion products. Evidence for secondary alloimmunisation was present in the great majority of patients. Early detection of alloimmunisation assists transfusion decision-making.

ABSTRACT NO.: 49**Incidence of Red Cell Alloantibodies in Two Multi-transfused Populations**

Christina Brown, Amber Madden, Andrew Webb, Heather Cleland, Merrole Cole-Sinclair, Stephen Opat

Alloimmunisation to red cell antigens is a recognised risk of transfusion and may complicate provision of blood products to multi-transfused patients. We hypothesised that patients with haematological malignancies, either as a result of chemotherapy and/or underlying disease process, may have impaired immune responses and therefore a lower rate of alloimmunisation. To address this hypothesis we performed a retrospective analysis comparing red cell alloantibody formation in two cohorts of multi-transfused patients. The first cohort consisted of 424 patients with haematological malignancies (HM) admitted between July 2003 and June 2005; the second included 116 patients with major burns (MB) admitted between April 2002 and April 2005. Both cohorts had similar demographics: 44% female, median age 55 (17–93) for HM; 36% female, median age 42 (16–85) for MB. Alloantibodies developed in a lower proportion of HM patients compared with MB patients; 16/424 (3.7%) versus 9/114 (7.9%), however this did not reach statistical significance ($p \leq 0.1$). Overall, the median number of red cell units received was greater for HM than MB; 16 (1–185) versus 5 (1–109). Preliminary results suggest the rate of alloimmunisation per red cell unit transfused to be less for HM than MB; 0.19% versus 0.46%. Rhesus antigens (E, D, C, Cw c, e) were the commonest specificity in both groups with Kell being the second commonest specificity in the MB group. Kell alloantibodies rarely developed in HM patients due to widespread use of Kell negative units in this cohort. Universal usage of Rh D and E compatible blood may have reduced the rate of alloimmunisation by 40% in patients with HM. This policy with the addition of Kell negative blood in MB patients would have reduced alloimmunisation to a similar degree. These data support the practise of Rh phenotyping of all patients where recurrent transfusion is anticipated.

ABSTRACT NO.: 50**Laboratory Investigation of Drug Induced Thrombocytopenia**A Hughes¹, J Crawford^{1,2}, J Lown¹, P Cannell¹¹ Department of Haematology, Royal Perth Hospital² PathCentre, Western Australian Centre for Pathology Research, Nedlands

INTRODUCTION Drug induced thrombocytopenia (DITP) is commonly caused by antibody specific destruction of platelets. Evidence of causation is provided by detection of drug-specific platelet reactive antibodies.

Aims Drug specific platelet antibody detection (DRG), (excluding heparin) at RPH was reviewed to examine the range of drugs tested, methods used and detection rate. The adequacy of current investigation protocols was also examined.

METHOD DRG investigations performed over 5 years were reviewed (hospital & referred tests). Testing involved a solid phase red cell adherence method (SPRCA) in presence and absence of drug plus flow cytometric assessment of platelet surface immunoglobulin. For patients with indeterminate results suggesting presence of circulating drug, testing was repeated once the drug had cleared the circulation.

RESULTS For 67 suspected DITP episodes, 77 drugs were tested on 66 patients (36 quinine, 11 glycoprotein IIb/IIIa inhibitors, 20 antibiotics, 10 others). Initial DRG test results were 33 negative, 9 indeterminate and 25 positive. Eight patients with indeterminate results had strong evidence of DITP on repeat testing (6 SPRCA, 1 MAIPA, 1 clinical). Positive DRG tests included: quinine(15), quinidine(1), abciximab(10), tirofiban(1), beta-lactams(3), phenytoin(2), thiazide(1). Only 5 patients with negative DRG results had repeat testing (0/5 were positive). 5 patients with negative DRG results had strong clinical evidence of DITP. Drugs tested in these cases were quinine, rifampicin, ceftazidime, cephalosporin, urokinase, timentin, cephalosporin.

CONCLUSION Positive DRG testing provides strong laboratory evidence of DITP. DRG testing should be performed when DITP

is first suspected, as timing of testing is critical. Patients with negative or indeterminate DRG results should be reassessed within 24–48 hours. The laboratory investigation protocol has been amended to improve drug detection in suspected DITP by adapting current methodology to better suit the proposed drug mechanism of action.

ABSTRACT NO.: 51**The Challenges of Providing Safe Blood in Africa**

Roger Dodd

American Red Cross Holland Laboratory, Rockville, Maryland, USA

Africa is a large and varied continent with some 46 different countries at varying levels of development. As of 2002, only 30% of the countries had drawn up their transfusion policies and, of these, not all were in place. Only a minority of systems has a fully voluntary donor system and fully 60% of blood is collected from family or replacement donors. In most parts of the continent, there is a critical shortage of blood, amounting to 70% of estimated need, exacerbated by shortages of qualified staff. The majority of blood is used to treat anemia of parasitism and maternal bleeding. Blood is most often collected in the hospital environment. In many countries, the cold-chain is fragile or non-existent. Financial issues create additional difficulties, particularly as equipment, supplies and test kits have to be imported. Testing may be incomplete, despite the very high prevalence rates for transfusion transmissible infections. In sub-Saharan countries, the prevalence of HIV infection may reach 20% and there may also be very high frequencies of infection with HBV. HCV prevalence rates vary, but are generally lower. Consequently, testing loss is very high, further affecting the blood supply. Additionally, much of the continent is highly endemic for malaria. In some locations, creative approaches have been developed for donor management (such as the "Club 25" concept) and these have been effective in reducing the prevalence of HIV among the donor population. Additionally, significant efforts are being made to introduce and maintain quality systems throughout the continent. There is an increasing trend towards provision of financial and technical support from developed nations and the US PEPFAR program will be outlined. The major difficulty with such external support is establishing sustainable outcomes.

ABSTRACT NO.: 52**Bloody Obstetrics: A Developing Country Perspective**

Miriam O'Connor

University of Papua New Guinea, Port Moresby, PNG

Maternal mortality: ~ 600 000 women die worldwide each year as a direct result of a pregnancy or childbirth complication. Most are preventable using cheap and simple measures. For every woman who dies, there are another 100 who are permanently disabled from the complications they survive.

Where does Obstetric Haemorrhage fit in with respect to Maternal Morbidity and Mortality?

Aetiologies

- Miscarriage and ectopic
- Unsafe abortion
- Antepartum haemorrhage
- Intrapartum haemorrhage
- Postpartum haemorrhage

Aided and abetted by high levels of intercurrent diseases and high prevalence of anaemias, eg, malaria, nutritional anaemias, chronic diseases, eg, TB and HIV.

Aided and abetted by:

- Corruption: funding and supply lines
- Social Determinants of Health eg Low education levels in the community, Low status of women, transport issues, cultural taboos

- Lack of resources: appropriately trained health staff, equipment, communication infrastructure, IT, appropriately banked and tested blood, medications, infrastructure, Standard Treatment Guidelines, agreed an measurable benchmarks, audit, monitoring and evaluation

Some examples:

The way forward, at least for obstetric haemorrhage

- Addressing the Social Determinants
- Health Promotion
- Blood banking and distribution
- Service planning, funding, audit and M&E
- Partnerships

Poverty Reduction and The Millennium Development Goals

- Poverty reduction strategies
- The debt crisis
- Global trade policies
- Science for Development
- Effective international development assistance

Myths and magic bullets

ABSTRACT NO.: 53

China – The New Blood Service

Dr Lin Che-kit

Hospital Chief Executive Hong Kong Red Cross Blood Transfusion Service

Before 1978, most clinical blood in China were sourced from paid donations. As an effort to eliminate paid blood donations, from 1978 to 1998, the provincial governments coordinated annual blood donation plan and assigned blood donation quota to all enterprises and organizations for them to fulfill. It was the prototype of non-remunerated blood donation in China. However, it was not voluntary as individuals had to take turns to be selected for blood donation. In deed, there were many instances of enterprise paying money or offering incentives such as promotion or vacation to selected staff to donate blood, or one person paying another person a good sum of money to make a donation instead of oneself.

On 1 October 1998, China enacted a blood donation law which laid down the system of voluntary non-remunerated blood donation and stipulated the duty of healthy citizens to donate blood. Since then, voluntary non-remunerated blood donation has been rapidly increasing across the country. According to the statistics released by the Ministry of Health, China, non-remunerated donations had jumped from for 22% in 1998 to 91.3% of clinical blood in 2004, and voluntary non-remunerated from 5% to 71.5%.

% of clinical blood	1998	1999	2000	2001	2002	2003	2004
Voluntary non-remunerated	5	13.6	21.1	39.4	58.6	61	71.5
Directed non-remunerated	17	32.1	37.5	33.1	29.9	24	19.8
Paid	78	54.3	41.4	27.5	11.5	15	8.7

The development of blood programme has also stimulated the establishment of blood centres. In 1978, there were only 40 blood centres in China. In 2000, there were 441 blood centres. Except Tibet, every province in China has established a provincial blood centre in the capital city. The roles of provincial blood centres are to provide technical support and training to prefecture blood centre as well as co-ordinating local blood supply.

ABSTRACT NO.: 54

Voluntary Non-Remunerated Blood Donation – Establishing Cultural Change in Developing Countries

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AIM Voluntary, non-remunerated blood donation is the cornerstone of a safe blood supply. As part of the World Health Organisation’s (WHO) Global Blood Safety Programme, a collaborative partnership has been established with the International Federation of Red Cross and Red Crescent Societies (the Federation) to effect cultural change in those developing countries where blood services are not currently based on voluntary blood donor programmes.

METHOD WHO and the International Federation have jointly prepared a set of materials for training in the basic principles of education, motivation, recruitment and retention of voluntary blood donors. The *Developing a Blood Donor Programme* curriculum has been produced to provide the basis on which workshops can be conducted. These workshops are designed to be interactive and focus on practical issues and the challenges associated with the phasing out paid or family replacement donor systems.

RESULT Since November 2004, regional and local workshops have been conducted in Singapore, the PR China, Nigeria, Vietnam and Egypt. Throughout the workshops, participants learn about the key elements of a successful blood donor programme, including identification of safe donor populations, audience-specific strategies for donor education, motivation and recruitment as well as pre-donation counselling, donor care and retention. They also undertake a Gap Analysis of their blood donor programmes. The ultimate outcome is the development of a Plan of Action for their local, regional or national Blood Service.

Progress on with the implementation of their plans is monitored regularly and ongoing support is provided by a regional contact, usually one of the workshop facilitators. Follow-up workshops after 12 months are planned to measure the results to date, in particular, improvements in the number of voluntary, non-remunerated blood donors.

CONCLUSION *The Developing a Blood Donor Programme* is an important initiative in developing countries, ultimately contributing to the delivery of safe and adequate blood supplies, a situation which is largely taken for granted in the developed world.