INTRODUCTION

In order to effectively manage the nation’s blood supply, modification of blood utilization practices has become a high priority, but that objective has been made especially challenging by society’s evolving demographics. The population of the United States is aging,1 and the first wave of Baby Boomers will reach the full retirement age of 65 in 2011. From then on, 10,000 new retirees will be added to Social Security and Medicare every day for 20 years.1 One unrecognized consequence of this demographic shift is the impact on our blood supply. According to the American Association of Blood Banks (AABB), the cutoff age for potential blood donors is 60 years of age. Therefore, as our population ages, the number of potential blood donors decreases and the number of potential blood recipients increases. A severe shortage of blood and blood components may develop in the foreseeable future, unless it is offset by a significantly increased supply, or by reduced usage of blood and blood components.2,3

To offset this concern, a systematic approach to Blood Management has evolved with an emphasis on quality, safety and cost efficiency of blood component therapy. The cornerstones of blood management programs are the implementation of evidence-based transfusion guidelines to reduce variability in transfusion practice, and the employment of multidisciplinary teams to study, implement, and monitor local blood management strategies. At Lancaster General Hospital, the Blood Utilization Review Committee has established evidence-based transfusion guidelines, and functions as the multidisciplinary team that monitors hospital blood management strategies. One of those strategies is Peri-Operative Autotransfusion, or the collection, processing, and reinfusion of the patient’s own blood that is lost during the peri-operative period.

Autotransfusion is uniquely advantageous because it directly reduces the demand for banked blood, while it simultaneously eliminates the risks of allogeneic blood transfusions.

CONSEQUENCES OF TRANSFUSION

In spite of the fact that the U.S. blood supply is the safest it has ever been, allogeneic blood transfusion is still associated with significant risks. The public mistakenly believes that most of the risk of allogeneic transfusion involves transmission of human immunodeficiency virus (HIV), but in fact, the most significant risks in 2006 were unrelated to viral transmission. Improved donor screening and testing of donated units has decreased the risk of hepatitis and HIV to less than 1 out of every 1,000,000 transfusions.4 Mistransfusion—administration of blood products to the wrong patient—is now one of the leading causes of transfusion complications. In spite of increased awareness and vigilance, mistransfusion still occurs in approximately 1 of every 14,000 units transfused.5 Death occurs in this group in 1 of 600,000 – 800,000 transfusions.6

Worldwide, the leading cause of transfusion related morbidity and mortality is Transfusion Related Acute Lung Injury, (TRALI), with an estimated frequency of 1 of 500 platelet transfusions and 1 of 1,000 – 5,000 plasma and red blood cell transfusions.[7-11] It is likely that the actual incidence of TRALI is higher than reported, due to a lack of awareness of the syndrome on the part of many clinicians.

PATHOPHYSIOLOGY OF TRANSFUSION-RELATED INJURY

TRALI and the Systemic Inflammatory Response Syndrome, (SIRS) are related to the buildup of mediators of inflammation in stored blood. Cytokines released from the residual leukocytes appear to be the primary concern, although complement activation has been implicated also.12 In addition, hemolysis of RBCs releases intracellular contents and raises serum potassium levels. This “storage lesion” is cumulative and worsens with prolonged storage. Several studies have associated “older” blood with higher morbidity and mortality.12,13,14 Currently, the AABB allows RBCs to be stored for up to 42 days.

TRALI can also be caused by the presence in donor plasma of antibodies to human leukocyte antigen.
These antibodies, which attack recipient white blood cells, are most prevalent in plasma products donated by multiparous females. In fact, the UK has recently converted to male-only plasma donations, and the US is likely to follow this lead.\textsuperscript{15}

Prolonged storage also impairs the ability of stored RBCs to deliver oxygen to the tissues.\textsuperscript{16,17} 2,3 diphosphoglyceric acid, (2,3,DPG), a substance produced in RBCs to aid in transfer of oxygen to tissues, is depleted quickly in stored blood and it can take 24 to 48 hours to replenish 2,3,DPG levels. Until 2,3,DPG levels are restored, transfused red cells have very limited participation in tissue oxygenation.

Another under-recognized complication of allogeneic transfusion is Transfusion Related Immunomodulation (TRIM). TRIM-invoked immunologic changes include both stimulation of humoral immunity resulting in production of allo-antibodies, and down-regulation of cellular immunity resulting in altered host defenses.\textsuperscript{18,19,20} Since transfusions generally only occur in patients already stressed by surgery or illness, TRIM is thought to contribute to the consistent finding of stepwise increases in infection rates,\textsuperscript{21-26} ventilator support times,\textsuperscript{25} ICU and hospital length of stays\textsuperscript{27,28,29} and short term and long term mortality in patients who receive transfusions.\textsuperscript{34,26,29} Also, several studies have shown an increase in cancer recurrence rates in transfused vs. non-transfused patients.\textsuperscript{30,31,32}

**CRITERIA FOR TRANSFUSION**

Despite mounting evidence that unnecessary transfusions can cause serious harm, several studies have documented a lack of compliance with appropriate transfusion guidelines, as well as tremendous variations in transfusion practices among different institutions and even among individual physicians within the same institution.\textsuperscript{33-37}

Lancaster General Hospital’s Blood Utilization Review Committee has established Blood Product Usage Screening Criteria based on the simple and unexceptionable principle that the benefit of a transfusion should outweigh its risk. To that end, the criteria state that patients should not receive donor banked blood unless the hemoglobin level is less than 7, except with extenuating circumstances. (Table 1)

**COST OF ALLOGENEIC TRANSFUSIONS**

The “true” cost of an RBC transfusion is difficult to estimate. Aside from the “direct” costs of acquiring, cross-matching, testing and infusing the blood, there are also “indirect” costs, such as the time physicians spend explaining the risks of transfusion and getting informed consent, or the time nurses spend monitoring the transfusion. There is also the time spent transporting blood around the hospital and cleaning up the waste, and the overhead cost of storing and controlling the blood in the blood bank and discarding any unused blood. Shander’s group at the The New Jersey Institute for Bloodless Medicine and Surgery, Englewood Hospital and Medical Center, Englewood, NJ, attempted to measure all these direct, indirect and overhead costs at their institution, and found the total cost of transfusing a unit of RBC in 2007 was $1,158.\textsuperscript{38} This total was the cost to the institution, not the patient charge, and it consisted of: indirect overhead cost - 40.6%, transfusion processing cost - 34.0%, weighted average acquisition cost - 21.5%, and direct overhead cost - 3.9%.

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**Table 1: Guidelines for Review of Blood Product Usage Screening Criteria:**

<table>
<thead>
<tr>
<th>Packed red blood cells or whole blood, homologous and autologous – One of:</th>
</tr>
</thead>
<tbody>
<tr>
<td>1. Hgb &lt; 7 gm/dl</td>
</tr>
<tr>
<td>2. Hgb &lt; 10 gm/dl with:</td>
</tr>
<tr>
<td>- symptoms of anemia and/or unstable vital signs, or</td>
</tr>
<tr>
<td>- extenuating circumstances (e.g. significant cardiac and/or pulmonary disease/condition; significant, rapid blood loss)</td>
</tr>
<tr>
<td>3. Patient age &gt; 65 with Hgb &lt; 8 gm/dl (&lt; 10 gm/dl for autologous transfusion)</td>
</tr>
<tr>
<td>4. Preoperative age &gt; 65 with Hgb &lt; 10 gm/dl</td>
</tr>
</tbody>
</table>
Shander’s analysis did not include the cost of treating any adverse events related to allogeneic transfusions. (There were 21 mild or moderate reactions in the 461 patients who received transfusions.) Nonetheless, the cost of treating even a minor transfusion reaction adds significantly to the indirect costs of allogeneic transfusions, not to mention the cost of treating any infection that results from immuno-modulation. Several studies have shown that patients who receive no transfusions, or autotransfusion RBC’s alone, have shorter LOS, and lower overall costs.\textsuperscript{38-40}

PERIOPERATIVE AUTOLOGOUS BLOOD MANAGEMENT AND AUTOTRANSFUSION

Surgical procedures account for a high percentage of blood transfusions,\textsuperscript{41} and anything that can be done to reduce blood utilization during and after surgery will have a substantial impact on transfusion requirements. Autotransfusion is one of the most important such interventions.

Autotransfusion involves the collection, washing, and reinfusion of blood shed at the surgical site. Intraoperatively, suction is used to collect shed blood in a controlled manner to a dedicated device rather than to a discard circuit. Heparin or Anticoagulant Citrate Dextrose is infused into the shed blood to keep it from clotting. Once enough blood has accumulated, it is centrifuged in the autotransfusion device to separate the RBCs from the waste products, and the RBCs are washed with 0.9% NaCl to remove any unwanted contaminants. Finally, the washed RBCs are filtered and reinfused to the patient. Since the patients receive their own fresh RBCs, the cells have high levels of 2,3DPG, and will be immediately active in tissue oxygenation. The RBC’s never leave the surgical area, are never put into the banked blood pool, do not incur storage expense, and have virtually no chance of a clerical error or mistransfusion. Moreover, problems with undesirable immune responses are mitigated.

Autotransfusion is contraindicated in some cancer operations, any operation with bacterial contamination, and in sickle cell disease. Otherwise, any operation in which there is the possibility of significant blood loss has the potential for autotransfusion. Intraoperative autotransfusion is performed in cardiac, vascular, orthopedic, neurosurgical, gynecological, and general surgery procedures, as well as during the management of trauma.

A major application of postoperative autotransfusion is in orthopedic surgery, primarily in joint replacement surgery. Drains are placed in the surgical site and blood shed postoperatively is collected and anticoagulated with Anticoagulant Citrate Dextrose. From that point the process is similar to intraoperative autotransfusion, with centrifugation, washing, filtering and reinfusion.

AUTOTRANSFUSION AT LANCASTER GENERAL HOSPITAL

Following are data about the number of patients and the autotransfusion volumes of RBCs infused over the last three calendar years (2007-2009) at Lancaster General Hospital:

<table>
<thead>
<tr>
<th>Year</th>
<th>Intraoperative</th>
<th>Postoperative</th>
<th>Total</th>
</tr>
</thead>
<tbody>
<tr>
<td>2007</td>
<td>1051</td>
<td>1029</td>
<td>2080</td>
</tr>
<tr>
<td>2008</td>
<td>1128</td>
<td>1020</td>
<td>2148</td>
</tr>
<tr>
<td>2009</td>
<td>952</td>
<td>1102</td>
<td>2054</td>
</tr>
</tbody>
</table>

A “unit” of allogeneic banked RBCs is approximately 250 cc. The total volume of reinfused RBCs is the equivalent of:

10,672 “units” of RBCs.

This is an average of 425 cc’s per patient.
CONCLUSION
Peri-operative Autotransfusion is an essential part of a comprehensive Blood Management program. It is a safe and cost effective technique to reduce the strain on the blood bank, and to ensure patients get their own blood back. It has been used effectively at Lancaster General Hospital, salvaging a massive amount of blood that would have otherwise been lost, and returning it to its rightful owner, the patient.

REFERENCES
PERIO-OPERATIVE AUTOTRANSFUSION


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