

score ranged from 0 to 17 points. Four risk categories of increasing severity (scores 0 to 2, 3 to 5, 6 to 8, and 9 to 13) were determined arbitrarily by the authors. The frequency of ARF among these categories varied between 0.4% for the lowest risk score to 21.5% for the highest score.

This study involved a large cohort of patients, sufficient to generate and validate a postoperative ARF score that incorporated multiple independent risk factors. Limitations included the single-center data source and the retrospective observational design. Nevertheless, the study provided a valuable tool for both determining the risk of ARF for individual patients and planning future clinical trials.

Clinical score needed for noncardiac surgery

Currently, a risk score for ARF has only been developed for patients who have had cardiac surgery; no sufficiently powered study has yet been done for those undergoing noncardiac surgery. The Section of Hospital Medicine and the Department of Nephrology and Hypertension at the Cleveland Clinic are currently conducting a large retrospective cohort study in patients undergoing elective noncardiac surgery.

Importance of identifying patients at risk

Identification of patients likely to develop ARF after surgery is important, as it enables physicians to improve care and to inform patients about their individual risk. Future intervention-based trials can be conceived to target high-risk populations to decrease length of stay, morbidity, and mortality.

REFERENCES

1. Schreiber MJ Jr. Minimizing perioperative complications in patients with renal insufficiency. *Cleve Clin J Med* 2006; 73(Suppl 1):S116–S120.
2. Chertow GM, Lazarus JM, Christiansen CL, et al. Preoperative renal risk stratification. *Circulation* 1997; 95:878–884.
3. Mangano CM, Diamondstone LS, Ramsay JG, Aggarwal A, Herskowitz A, Mangano DT. Renal dysfunction after myocardial revascularization: risk factors, adverse outcomes, and hospital resource utilization. *Ann Intern Med* 1998; 128:194–203.
4. Loeff BG, Epema AH, Smilde TD, et al. Immediate postoperative renal function deterioration in cardiac surgical patients predicts in-hospital mortality and long-term survival. *J Am Soc Nephrol* 2005; 16:195–200.
5. Thakar CV, Arrigain S, Worley S, Yared J-P, Paganini EP. A clinical score to predict acute renal failure after cardiac surgery. *J Am Soc Nephrol* 2005; 16:162–168.

Address: Vesselin Dimov, MD, Section of Hospital Medicine, Department of General Internal Medicine, Cleveland Clinic, 9500 Euclid Avenue, S70, Cleveland, OH 44195; dimovv@ccf.org.

Q: Why treat anemia in the preoperative period of joint replacement surgery with erythropoietin?

AJAY KUMAR, MD*

Section of Hospital Medicine
Department of General Internal Medicine
Cleveland Clinic, Cleveland, OH

VESELIN DIMOV, MD*

Section of Hospital Medicine
Department of General Internal Medicine
Cleveland Clinic, Cleveland, OH

A: Recombinant human erythropoietin (epoetin alfa) is an effective therapy approved by the US Food and Drug Administration (FDA) to treat preoperative anemia in patients undergoing knee or hip replacement surgery.

Anemia is linked to poor outcomes

Anemia in the preoperative period is a known predictor of adverse outcomes in surgical patients. Carson et al¹ studied 125 consecutive patients who declined blood transfusions and found that operative mortality was 16 times greater in patients with hemo-

globin levels less than 8 mg/dL than in patients with higher hemoglobin levels. Other studies suggest that it is not the anemia itself that is associated with increased mortality, but the practice of treating anemia with blood transfusions.²

Blood transfusion used most often, alternatives needed

Allogenic blood transfusion is the most commonly used technique in the United States to correct perioperative anemia. Although several methods are available to reduce the need for blood transfusion, the use of allogenic blood remains very high; for example, 47,456 units of blood products were used in the perioperative period of various orthopedic surgeries at the Cleveland Clinic in 2004.

In the next decade, with the aging of the population, the number of major joint replacements to be performed annually will increase, likely leading to an

* Both authors reported that they have no commercial affiliations or financial interests that pose a potential conflict of interest with this article.

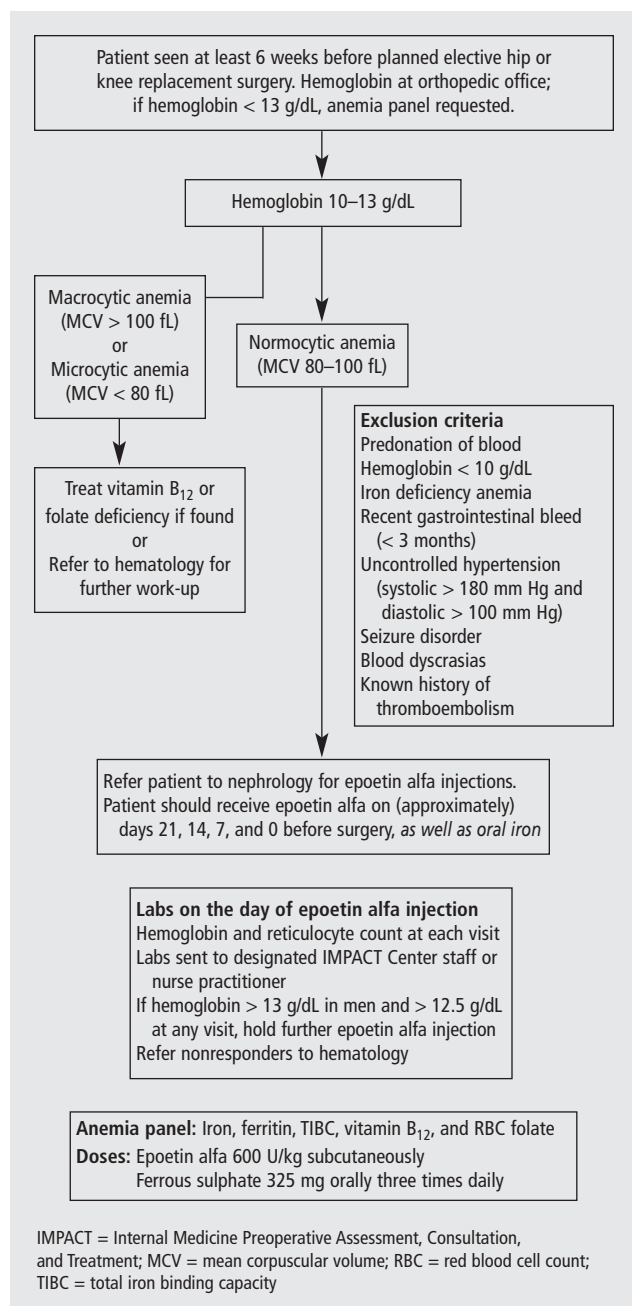


FIGURE. Protocol for management of preoperative anemia at the Cleveland Clinic.

increase in the demand for blood products. Due to the fluctuating supply and the costs and risks associated with the administration of blood products, searching for potential alternatives to transfusions is imperative.

Because blood loss associated with orthopedic surgery is predictable, the application of a carefully designed blood management program is appropriate. However, the usefulness of autologous donations and

cell saver machines is limited by incomplete utilization of predonated blood and high cost. Furthermore, allogenic blood transfusions carry certain risks, in particular ABO incompatibility caused by administrative error and transfusion-related lung injuries.³ Exposure to leukocytes in allogenic blood can also cause immunosuppression.⁴ In a large prospective study of 6,301 patients undergoing noncardiac surgery, Dunne et al⁵ concluded that the incidence of perioperative anemia in surgical patients is high and is related to an increase in blood utilization. These factors are associated with an increased risk for perioperative infection and other adverse outcomes (including death) in surgical patients.

In a large study by Bierbaum et al,⁶ blood management data were collected prospectively on patients who had undergone total hip replacement and knee replacement. Fifty-seven percent of the patients undergoing hip replacement and 39% of the patients undergoing knee replacement received blood transfusions. Patients who were transfused were more likely to have infections, fluid overload, and an increased length of hospitalization compared with patients who did not receive transfusions. The Orthopedic Surgery Transfusion Hemoglobin European Overview study from 225 centers in Europe produced similar results.⁷ In this study, allogenic transfusion was associated with a higher rate of wound infection than autologous transfusion (4.2% vs 1%, respectively).

Treatment of anemia and blood conservation

Treatment of perioperative anemia has been shown to decrease the need for transfusion and to improve perioperative outcomes such as postoperative infections, length of stay, and mortality in patients undergoing joint replacement surgery. The efficacy of preoperative erythropoietin therapy for increasing patients' hemoglobin concentrations and reducing exposure to allogenic red blood cell transfusion in orthopedic surgery has been demonstrated in several double-blind randomized clinical trials.^{8–11} Synthetic erythropoietin was approved by the FDA and has been used for almost 9 years in orthopedic surgeries as a method to improve hemoglobin levels in anemic patients undergoing surgery, and thus to decrease blood transfusions. Several centers in the United States have adopted this novel therapy to reduce the use of blood transfusions.

At the Cleveland Clinic, patients selected for a blood conservation protocol (**Figure**) with erythropoietin are eligible for four subcutaneous injections of epoetin alfa (600 U/kg) at days 21, 14, and 7 before surgery and on the day of surgery. Exclusion criteria for preop-

erative erythropoietin treatment are hemoglobin of less than 10 g/dL, iron deficiency anemia, recent gastrointestinal bleed (within 3 months), uncontrolled hypertension, seizure disorder, predonation of blood, blood dyscrasias, and history of thromboembolism.

Reticulocyte count, hemoglobin, and blood pressure should be checked prior to each injection. Iron deficiency may occur during erythropoietin therapy. Normal ferritin but low transferrin saturation may be observed due to an inability to mobilize iron stores rapidly enough to keep pace with the increased erythropoiesis. Supplemental oral or intravenous iron supports erythropoiesis and prevents iron store depletion.

Summary

Treatment of anemia in the perioperative period of major orthopedic surgery decreases the need for blood transfusion and improves perioperative outcomes. Use of epoetin alfa in this setting is FDA-approved and provides significant benefit to qualified and carefully chosen patients.

REFERENCES

1. Carson JL, Poses RM, Spence RK, et al. Severity of anemia and operative mortality and morbidity. *Lancet* 1988; 1:727–729.
2. Benoist S, Panis Y, Pannegon V, Alves A, Valleur P. Predictive factors for perioperative blood transfusions in rectal resection for cancer: a multivariate analysis of a group of 212 patients. *Surgery*

2001; 129:433–439.

3. Goodnough LT. Risks of blood transfusion. *Anesthesiol Clin North America* 2005; 23:241–252.
4. Goodnough LT, Brecher ME, Kanter MH, AuBuchon JP. Transfusion medicine. First of two parts—blood transfusion. *N Engl J Med* 1999; 340:438–447.
5. Dunne JR, Malone D, Tracy JK, et al. Perioperative anemia: an independent risk factor for infection, mortality, and resource utilization in surgery. *J Surg Res* 2002; 102:237–244.
6. Bierbaum BE, Callaghan JJ, Galante JO, et al. An analysis of blood management in patients having a total hip or knee arthroplasty. *J Bone Joint Surg Am* 1999; 81:2–10.
7. Rosenthal N, Kerkkamp HE, Macheras G, et al. Orthopedic Surgery Transfusion Hemoglobin European Overview (OSTHEO) study: blood management in elective knee and hip arthroplasty in Europe. *Transfusion* 2003; 43:459–469.
8. de Andrade JR, Jove M, Landon G, et al. Baseline hemoglobin as a predictor of risk of transfusion and response to epoetin alfa in orthopedic surgery patients. *Am J Orthop* 1996; 25:533–542.
9. Goldberg MA, McCutchen JW, Jove M, et al. A safety and efficacy comparison study of two dosing regimens of epoetin alfa in patients undergoing major orthopedic surgery. *Am J Orthop* 1996; 25:544–552.
10. Faris PM, Ritter MA, Abels RI. The effects of recombinant human erythropoietin on perioperative transfusion requirements in patients having a major orthopaedic operation. The American Erythropoietin Study Group. *J Bone Joint Surg Am* 1996; 78:62–72.
11. Laupacis A, Fergusson D. Erythropoietin to minimize perioperative blood transfusion: a systematic review of randomized trials. The International Study of Perioperative Transfusion (ISPOT) Investigators. *Transfus Med* 1998; 8:309–317.

Address: Ajay Kumar, MD, Section of Hospital Medicine, Department of General Internal Medicine, Cleveland Clinic, 9500 Euclid Avenue, A13, Cleveland, OH 44195; kumara@ccf.org.

Q: Obstructive sleep apnea: What to do in the surgical patient?

ROOP KAW, MD*

Section of Hospital Medicine
Department of General Internal Medicine
Cleveland Clinic, Cleveland, OH

JOSEPH GOLISH, MD*

Department of Pulmonary, Allergy and
Critical Care Medicine
Cleveland Clinic, Cleveland, OH

A: Adverse surgical outcomes appear to be more frequent in patients with known obstructive sleep apnea syndrome (OSAS). Anesthetic, sedative, and analgesic drugs should be used with extreme caution in patients with known or suspected OSAS, and close perioperative monitoring of high-risk patients is recommended.

Epidemiology

In Western countries, the prevalence of OSAS is about 5%.¹ The estimated prevalence in surgical

patients is 1% to 9%, though it may be even more common in certain populations.²

Disruption of sleep architecture

Sleep studies in patients who undergo major abdominal or cardiac surgery have demonstrated suppression of rapid eye movement (REM) sleep and slow-wave sleep after surgery. The REM sleep returns or rebounds in the late postoperative period (when oxygen may have been discontinued). This return of REM sleep was linked to significant respiratory abnormalities in a group of elderly patients who underwent abdominal vascular surgery.³ In REM sleep, the neural drive to the pharyngeal muscles is at a minimum, and the atonia of antigravity muscles predisposes to airway instability, causing episodic hypoxemias. Reductions in REM and slow-wave sleep

* Both authors reported that they have no commercial affiliations or financial interests that pose a potential conflict of interest with this article.