

About Dosage Schemes and Safety of Tranexamic Acid in Cardiac Surgery

To the Editor:—Dowd *et al.*¹ have provided important information about the pharmacokinetics of tranexamic acid (TA) in cardiac surgery with cardiopulmonary bypass. Particularly, they demonstrated the necessity of a continuous infusion of TA to obtain stable therapeutic concentrations. Then, they proposed two different dosage schemes in low- and high-risk patients for bleeding, to obtain TA plasma concentrations of 334 μM and 800 μM , respectively. Considering the patients with low-risk for bleeding, they recommended a loading dose of 12.5 mg/kg (or greater) over 30 min, a continuous infusion of 6.5 mg \cdot kg⁻¹ \cdot hr⁻¹, and 1 mg/kg (or greater) added to the priming, whereas in patients with high-risk for bleeding they proposed doses about 2.5 higher.

My group published various studies²⁻⁴ proposing an original pharmacologic protocol for TA that seem very similar to that proposed by Down *et al.* For patients with low-risk for bleeding; that is, a loading dose of 1 g over 20 min before sternotomy (and not 1 g, 20 min before sternotomy, as erroneously reported in the work of my group cited by Dowd *et al.*⁵), followed by a continuous infusion of 400 mg/h, and 500 mg added to pump priming. We also applied the same protocol in high-risk patients for bleeding, obtaining a significant reduction of blood loss and allogeneic transfusions.^{4,5} One criticism of the study of Dowd *et al.* is that the need to increase the doses of TA in this type of patient requires further clinical demonstrations, particularly considering (as the same authors report) that TA plasma concentrations of about 200 μM completely inhibit fibrinolysis.

Concerning the administration of TA after surgery, I agree with Dowd *et al.* regarding benefits in the postoperative period depending on intraoperative dosing techniques, but I do not understand why the authors claim that the efficacy of prolonged TA administration in the postoperative period is an open question. In reality, the authors, applying their pharmacokinetic model to our TA protocol, confirmed the conclusions of our study.⁵ Our intraoperative TA dosage scheme guarantees therapeutic concentrations for about 12 h, rendering unnecessary postoperative infusion of the drug.

It also seems that the potential thrombotic risk intrinsic to antifibrinolytic drugs is underestimated by Down *et al.* I particularly disagree with their statement, "TA appears to be a very safe medication. . . . Thus our attempt to avoid excessive TA concentrations may not be necessary."¹ An extensive literature search showed cases of thrombosis following the use of hemostatic drugs such as ϵ -aminocaproic acid, aprotinin, and TA, but it would be sufficient to cite a recent case report⁶ describing two fatal cases of thrombosis after the use of ϵ -aminocaproic acid (very similar to TA with

a potency 10 times lower) in cardiac surgical patients operated on with deep hypothermic circulatory arrest, wherein postmortem laboratory analysis revealed the presence of Factor V Leiden. Because it is impossible to identify patients with a preoperative prothrombotic state, it is appropriate to use the minimal effective doses of hemostatic drugs to avoid amplifying these thrombotic complications. Furthermore, the same authors' group reported in a previous study⁷ three cases of stroke in cardiac surgical patients with known peripheral vascular disease, treated intraoperatively with high doses of TA. One would speculate that high concentrations of TA facilitated the formation of a thrombus in the presence of blood flow reductions in a diseased vessel.

In conclusion, only large, prospective, blinded studies will establish the real safety and efficacy of the various doses of tranexamic acid in cardiac surgery. Currently, caution is required when using a hemostatic drug.

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References

1. Dowd NP, Karski JM, Cheng DC, Carroll JA, Lin Y, James RL, Butterworth J: Pharmacokinetics of tranexamic acid during cardiopulmonary bypass. *ANESTHESIOLOGY* 2002; 97:390-9
2. Casati V, Guzzon D, Oppizzi M, Cossolini M, Torri G, Calori G, Alfieri O: Hemostatic effects of aprotinin, tranexamic acid and ϵ -aminocaproic acid in primary cardiac surgery. *Ann Thorac Surg* 1999; 68:2252-7
3. Casati V, Guzzon D, Oppizzi M, Bellotti F, Franco A, Gerli C, Cossolini M, Torri G, Calori G, Benussi S, Alfieri O: Tranexamic acid compared with high-dose aprotinin in primary elective heart operations: Effects of perioperative bleeding and allogeneic transfusions. *J Thorac Cardiovasc Surg* 2000; 120:520-7
4. Casati V, Sandrelli L, Speziali G, Calori G, Grasso MA, Spagnolo S: Hemostatic effects of tranexamic acid in elective thoracic aortic surgery: A prospective, randomized, double-blind, placebo controlled study. *J Thorac Cardiovasc Surg* 2002; 123:1084-91
5. Casati V, Bellotti F, Gerli C, Franco A, Oppizzi M, Cossolini M, Calori G, Benussi S, Alfieri O, Torri G: Tranexamic acid administration after cardiac surgery: A prospective, randomized, double-blind, placebo-controlled study. *ANESTHESIOLOGY* 2001; 94:8-14
6. Fanashawe MP, Shore-Lesserson L, Reich DL: Two cases of fatal thrombosis after aminocaproic acid therapy and deep hypothermic circulatory arrest. *ANESTHESIOLOGY* 2001; 95:1525-7
7. Karski JM, Teasdale SJ, Norman P, Carroll J, Van Kessel K, Wong P, Glynn MF: Prevention of bleeding after cardiopulmonary bypass with high-dose tranexamic acid. *J Thorac Cardiovasc Surg* 1995; 110:835-42

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In Reply:—We appreciate your interest in our work and would like to take this opportunity to reply to your comments. First, we would like to apologize for misquoting your tranexamic acid (TA) administration regimen. However, we think that the principal behind the administration of a loading dose remains the same (1 g of TA over 20 min).

Regarding the criticism with respect to validity of increased TA dosage for patients undergoing complex cardiac surgery, we would like to emphasize that our previous studies (*J Thorac Cardiovasc Surg* 1995; 110:835-42) showed that higher TA dose regimens were more effective in reducing postoperative bleeding compared to a lower single-dose regimen.

With respect to safety aspects of TA, we must admit that we have never claimed that the use of any antifibrinolytics during cardiac surgery is a safe practice in the face of circulatory arrest. All information in the literature about vascular thrombosis after the use of antifibrinolytics in cardiac

surgery is anecdotal. We have recently reviewed stroke rates (as an indicator of vascular thrombosis) in our prospectively collected database of 18,000 primary coronary artery bypass graft patients with respect to utilization of TA. Stroke rates of 1.2-1.4% were similar between the two groups of patients, regardless of TA assignment. Furthermore, we have conducted a prospective randomized placebo controlled trial that demonstrated that there was no difference in early coronary graft patency between high-dose TA and placebo groups (*Can J Anaesth* 1999; 46:A29).

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Measurement of the Low-frequency Component of Blood Pressure Variability Can Assist the Interpretation of Heart Rate Variability Data

To the Editor:—We congratulate Hogue *et al.*¹ for their excellent investigation of the autonomic effects of dexmedetomidine in humans. Although a good understanding of the hemodynamic effects of different anesthetic drugs in patients is the daily routine for most anesthesiologists, the origin and mechanisms of these drug effects are still confusing and complex. The author's use of heart rate variability (HRV) in addition to baroreflex responses and systemic catecholamine concentrations in these volunteers exemplifies the need to use multiple measurement techniques when examining the sympathetic and parasympathetic nervous system. We suggest that the addition of blood pressure variability (BPV) to this study would help strengthen the HRV data and expand their measurement techniques. As the authors stated, the low-frequency oscillatory component of the respiratory rate interval is considered a marker of sympathetic activity. Similarly, one can identify low-frequency oscillations that emanate from a different end organ, arterial smooth muscle, known as Mayer waves.² The simultaneous measurement of HRV and systolic BPV has been described and used by Malliani *et al.* and showed increases in *both* low-frequency components with tilt,³ exercise,⁴ mental stress,⁵ physical exercise,⁶ moderate hypotension,³ and coronary artery occlusion.⁷ In conditions that directly affect one of the respective end organs, *i.e.*, heart (HRV) *versus* peripheral arteries (BPV), a unilateral response may be identified.⁸ Thus, the low-frequency component of systolic BPV, in addition to that generated by HRV, represents a good marker of sympathetic activity.⁹ Unfortunately, although the high-frequency component of HRV is considered a marker of parasympathetic activity, the high-frequency component of BPV is not. This is because the high-frequency oscillations in blood pressure are largely influenced by the mechanical changes induced by respiration.⁹ Therefore, we believe that analysis of the changes in low-frequency oscillations of the systolic blood pressure simultaneously with low-frequency and high-frequency oscillations measured in HRV will help investigators better explore and

quantitate the effects of anesthetic drugs on neural regulatory mechanisms.

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References

1. Hogue CW, Talke P, Stein PK, Richardson C, Domitrovich PP, Sessler DI: Autonomic nervous system responses during sedative infusions of dexmedetomidine. *ANESTHESIOLOGY* 2002; 97:592–8
2. Mayer S: Studien zur Physiologie des Herzens und der Blutgefäße: 5. Abhandlung: Über spontane Blutdruckschwankungen. *Sber Akad Wiss Wien* 1876; 74:281–307
3. Pagani M, Lombardi F, Guzzetti S, Rimoldi G, Furlan R, Pizzinelli P, Sandrone G, Malfatto G, Dell'Orto S, Piccalugga E, Turicci M, Baselli G, Cerutti S, Malliani A: Power spectral analysis of heart rate and arterial pressure variabilities as a marker of sympatho-vagal interaction in man and conscious dog. *Circ Res* 1986; 59:178–93
4. Furlan R, Dell'Orto S, Crivellaro W, Pizzinelli P, Cerutti S, Lombardi F, Pagani M, Malliani A: Effects of tilt and treadmill exercise on short-term variability in systolic arterial pressure in hypertensive men. *J Hypertens* 1987; 5:S423–S425
5. Pagani M, Furlan R, Pizzinelli P, Crivellaro W, Cerutti S, Malliani A: Spectral analysis of R-R and arterial pressure variabilities for the assessment of sympatho-vagal interactions during mental stress in humans. *J Hypertens* 1989; 6:S14–5
6. Pagani M, Somers V, Furlan R, Dell'Orto S, Conway J, Baselli G, Cerutti S, Sleight P, Malliani A: Changes in autonomic regulation induced by physical training in mild hypertension. *Hypertension* 1988; 12:600–10
7. Rimoldi G, Pierini S, Ferrari A, Cerutti S, Pagani M, Malliani A: Analysis of short term oscillations of R-R and arterial pressure in conscious dogs. *Am J Physiol* 1990; 258:H967–76
8. Guzzetti S, Cogliati C, Broggi C, et al.: Heart period and arterial pressure variabilities in quadriplegic patients. *Am J Physiol* 1994; 266:H1112–20
9. Malliani A, Pagani M, Lombardi F, Furlan R, Guzzetti S, Cerutti S: Spectral analysis to assess increased sympathetic tone in arterial hypertension. *Hypertension* 1991; 4(suppl III):36–42

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In Reply:—We thank Introna *et al.*¹ for their kind comments regarding our manuscript.² The authors correctly point out that a comprehensive approach to assessing the autonomic nervous system effects of anesthetic and sedative drugs provides clinically useful insight into their cardiovascular effects. Measurement of variations in systolic blood pressure is a well-characterized method for measuring peripheral sympathetic nervous system responses to various autonomic stressors, and such data might have furthered our findings.^{2,3} We chose, however, to focus on the cardiac autonomic effects of sedative doses of dexmedetomidine during clinically common thermal stress. Future investigation along these lines that incorporate measurement of systolic blood pressure variations might prove to be useful.

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References

1. Introna R, Blair J, Martin DC: Measurement of the LF component of blood pressure variability can assist the interpretation of HRV data. *ANESTHESIOLOGY* 2002; 97:592–8
2. Hogue CW Jr, Talke P, Stein PK, Richardson C, Domitrovich PP, Sessler DI: Autonomic nervous system responses during sedative infusions of dexmedetomidine. *ANESTHESIOLOGY* 2002; 97:592–8
3. Pagani M, Lombardi F, Guzzetti S, Rimoldi G, Furlan R, Pizzinelli P, Sandrone G, Malfatto G, Dell'Orto S, Piccalugga E, Turicci M, Baselli G, Cerutti S, Malliani A: Power spectral analysis of heart rate and arterial pressure variabilities as a marker of sympatho-vagal interaction in man and conscious dog. *Circ Res* 1986; 59:178–193

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Pain Significantly Influences Postoperative Wound Oxygen Tension

To the Editor:—Buggy *et al.* conducted a very interesting study investigating the influence of epidural or intravenous analgesia on postoperative wound oxygen tension.¹ They found significantly increased values in patients who received epidural analgesic techniques when compared with patients who received intravenous analgesia. In my opinion, the two groups were not comparable, because the patients without neuraxial blockade had significantly higher pain scores. It is known that pain is a major determinant of postoperative subcutaneous oxygen partial pressures.² It causes an autonomic response with increased adrenergic nerve activity and plasma-catecholamine concentrations. Hence, according to the presented data it seems unjustified to conclude that epidural techniques improve in general postoperative oxygen wound tension.

The role of neuraxial analgesia to improve postoperative outcome remains to be debated.³ Randomized trials comparing different perioperative analgesic concepts are often unable to demonstrate the superiority of one of the concepts. This is partially attributable to lack of power and a focus on major morbidities. Buggy *et al.* must be congratulated for the aim of their study, but I suggest repeating the

investigation while ensuring that patients in both groups receive comparably effective pain treatments. This is not an easy task because patients recovering from surgical interventions often consider pain as normal. They should, therefore, be instructed to use the patient-controlled devices not only to achieve a tolerable level of analgesia but also to aim for an *a priori* defined level of analgesia.

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References

1. Buggy DJ, Doherty WL, Hart EM, Pallett EJ: Postoperative wound oxygen-tension with epidural or intravenous analgesia. *ANESTHESIOLOGY* 2002; 97:952–8
2. Akça O, Melischek M, Scheck T, Hellwagner K, Akiliç CF, Kurz A, Kapral S, Heinz T, Lackner FX, Sessler DI: Postoperative pain and subcutaneous oxygen tension. *Lancet* 1999; 354:41–2
3. Kehlet H, Holte K: Effects of postoperative analgesia on surgical outcome. *Br J Anaesth* 2001; 87:62–72

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In Reply:—We thank Dr. Hübler for his interest in our manuscript, but we think he may have missed the point. Our stated hypothesis was to evaluate whether combined general anesthesia and epidural anesthesia and postoperative analgesia would increase subcutaneous tissue oxygen tension in the wound (P_{sqO₂}) compared with general anesthesia alone followed by postoperative opioid analgesia. It is well recognized that epidural analgesia provides superior analgesia compared with intravenous morphine analgesia, and as Dr. Hübler points out, minimizing pain increases P_{sqO₂}. Therefore, *of course* our patients with epidural analgesia had less pain: The issue we evaluated was whether the *method* of analgesia (epidural or intravenously administered opioid) also influences P_{sqO₂}. Furthermore, randomization took

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place preoperatively, and the distinction between our two groups commenced before the induction of anesthesia, not on emergence.

We believe Dr. Hübler's proposed repeat study is unworkable, because postoperative patients on intravenously administered opioid patient-controlled analgesia can never realistically expect to achieve similar levels of analgesia after major laparotomy compared with functional epidural analgesia.

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It's the Colloid, Not the Esophageal Doppler Monitor

To the Editor:—The debate over whether crystalloid or colloid is a better replacement fluid in the perioperative period has been of interest for decades. Gan *et al.*¹ have introduced new data in this debate with their study of 100 patients undergoing major elective surgery with an anticipated 500-ml blood loss. The authors used esophageal Doppler monitor (EDM)-guided colloid administration to improve preload, cardiac output, tissue perfusion, and patient outcome in the treatment group.

In effect, the treatment group has two variables: use of the EDM and use of colloid boluses. The EDM provided guidance for "goal-directed" fluid therapy. Although the EDM may not give an accurate measurement of preload or cardiac output,^{2,3} as a clinical monitor it enables an assessment of changes in corrected flow time and preload. With regard to the use of colloid boluses, the treatment group received 200 ml of

6% hydroxyethyl starch in saline every 15 min to maximize corrected flow time and stroke volume. The control group received 200-ml "fluid" boluses (more likely crystalloid) to treat changes in hemodynamic parameters (heart rate, blood pressure, central venous pressure) or decreased urinary output.

The better outcomes in the treatment group can be attributed solely to those patients receiving 847 ml of hydroxyethyl starch compared to 282 ml for the control group. Both groups received 4400 ml of lactated Ringer's solution during roughly 4 h of surgery. Other studies have demonstrated that EDM-guided use of supplemental colloid boluses in cardiac⁴ and orthopedic surgical patients⁵ was associated with improved outcomes, including shorter hospital stays. Taken together, the three studies demonstrate significantly improved patient outcomes when larger amounts of colloid were given to patients in the treatment group.

An esophageal Doppler monitor costs \$8000, with an additional

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\$100 for each esophageal probe (verbal communication, Deltex Medical, Branford, Connecticut, October 2002). The EDM provides a better assessment of preload than does a pulmonary artery or central venous catheter,⁶ but contrary to the BBC interviews⁷ that resulted from the present study, we suggest that none of these three methods may be required to achieve the benefits of earlier return to bowel function, lower incidence of postoperative nausea and vomiting, and decrease in length of postoperative hospital stay. Based on the findings of Gan *et al.*, we suggest that the intraoperative administration of 500–1000 ml of hydroxyethyl starch will provide a more cost-effective patient benefit for relatively healthy patients, without significant cardiovascular disease, who are undergoing major elective surgery.

We agree with the authors that “intraoperative fluid augmentation appears to confer significant benefits compared with customary practice.” Further investigation is required to test whether additional colloid administration without the use of EDM would result in similar patient benefits, because Gan *et al.* do not directly address this issue. Crystalloid or colloid? The results of this study imply that administration of both may improve patient outcomes after major elective surgery.

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In Reply:—Thank you for allowing us the opportunity to respond to the letter by Horowitz and Kumar. In our study,¹ we investigated whether goal-directed intraoperative plasma volume expansion guided by the esophageal Doppler monitor would shorten the length of hospital stay and improve postoperative outcomes. The control and protocol groups consisted of patients undergoing similar procedures. The general anesthetic for both groups was standardized. We did not regiment the type of fluid to be administered in the control group, because we wanted this group to receive, as closely as possible, a routine standard of care. The protocol group received 6% hetastarch in saline to a maximum of 20 ml/kg, based on a fluid challenge algorithm. Patients in the protocol group received, on average, 500 ml more colloid than the control group. Hence, we could not completely rule out that additional colloid may have contributed to the findings of our study. This was stated in the discussion of the original manuscript. Goal-directed fluid administration is a strategy. Using the fluid challenge algorithm with esophageal Doppler monitoring minimizes the risk of over-resuscitation, because stroke volume is reassessed before each additional fluid bolus.

A number of confounding factors in this study must be addressed in future studies. First is the timing of fluid administration. A significantly greater volume of intravenous fluid was administered toward the beginning of the surgical procedure in the protocol group than in the control group. It may be that the earlier administration of fluid resulted

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Alternative Method to Deflate the Operated Lung when Using Wire-guided Endobronchial Blockade

To the Editor:—In his excellent review, Dr. Campos¹ describes clinically relevant lung isolation methods. To deflate the lung when using a wire-guided endobronchial blocker (WEB®; Cook, Bloomington, IN), Dr. Campos proposes withdrawing the loop wire, thus opening a 1.4-mm air channel to the atmosphere. Because the procedure as proposed can be very time-consuming, we use an alternative method to deflate the operated lung when using WEB®: After conventional intubation, we introduce the WEB® catheter in the main stem bron-

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References

1. Gan TJ, Soppitt A, Maroof M, El-Moalem H, Robertson KM, Moretti E, Dwane P, Glass PSA: Goal-directed intraoperative fluid administration reduces length of hospital stay after major surgery. *ANESTHESIOLOGY* 2002; 97:820–826
2. Guzzetta NA, Ramsay JG, Bailey JM, Palmer-Steele C: Clinical evaluation of the esophageal Doppler monitor for continuous cardiac output monitoring. *Anesth Analg* 1998; 86:SCA82
3. Lefrant JY, Bruelle P, Aya AGM, Saissi G, Dauzat M, de La Coussaye JE, Eledjam JJ: Training is required to improve the reliability of esophageal Doppler to measure cardiac output in critically ill patients. *Intensive Care Med* 1998; 24:347–52
4. Mythen MG, Webb AR: Perioperative plasma volume expansion reduces the incidence of gut mucosal hypo perfusion during cardiac surgery. *Arch Surg* 1995; 130:423–9
5. Sinclair S, James S, Singer M: Intraoperative intravascular volume optimization and length of hospital stay after repair of proximal femoral fracture: Randomized controlled trial. *BMJ* 1997; 315:909–12
6. Madan AK, UyBarreta VV, Aliabadi-Wahle S, Jespersen R, Hartz RS, Flint LM, Steinberg SM: Esophageal Doppler ultrasound monitor versus pulmonary artery catheter in the hemodynamic management of critically ill surgical patients. *J Trauma* 1999; 46: 607–11; discussion 611–2
7. BBC News Broadcast. Sound waves to cut up risk. October 7, 2002, 01:23 GMT. Source: Deltex Medical, Branford, CT at ASA Annual Meeting, Orlando, Florida, October 12–16, 2002 verified at: <http://news.bbc.uk/1/hi/health/2295111.stm>

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in better perfusion of the gastrointestinal tract and, hence, earlier resumption of gastrointestinal motility and return to normal diet.

The type of fluid administered may also be important in postoperative patient outcome. In a recent study,² we found an improvement in the quality of recovery in patients receiving a combination of colloid (6% hetastarch) and crystalloid (lactated Ringer's) *versus* crystalloid alone. Specifically, the colloid/crystalloid patients had a lower incidence of postoperative nausea and vomiting, severe pain, and peripheral edema. Further investigations are therefore needed to study the contribution of these factors.

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References

1. Gan TJ, Soppitt A, Maroof M, El-Moalem H, Robertson KM, Moretti E, Dwane P, Glass PSA: Goal-directed intraoperative fluid administration reduces length of hospital stay after major surgery. *ANESTHESIOLOGY* 2002; 97:820–6
2. Moretti E, Robertson KM, El-Moalem H, Gan TJ: Intraoperative colloid administration reduces postoperative nausea and vomiting and improves postoperative outcomes compared to crystalloid administration. *Anesth Analg* 2003; 96:611–7

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chus of the operated lung but do not inflate it. After patient positioning and thoracotomy, we check and, when needed, correct the position of the WEB® catheter using a fiberoptic. To deflate the lung, we then disconnect the ventilator from side port of the WEB® connector and thus open the main airway to the atmosphere. After verifying lung collapse, we inflate the WEB® under fiberoptic control and reconnect the ventilator to the endotracheal tube. This way we avoid removing the flexible wire loop, which is essential for repositioning the WEB® if it

dislodges during the operation. Only if the surgical field is in close proximity to¹ the WEB® balloon do we recommend removing the wire loop during the operation. For most operative procedures, the probability that the WEB® wire loop could be included in the stapling line is very small. Of note, it is essential to inflate the WEB® balloon under fiberoptic control because the amount of air need to properly seal the airway may be otherwise mostly underestimated but occasionally overestimated.

With regard to this letter and the subsequent two letters and reply, Richard B. Weiskopf, M.D., is the Editor for this exchange.

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Endobronchial Blocker Response

To the Editor:—I would like to respond to Dr. Campos' excellent article on options for one-lung ventilation.¹ The use of a conventional single-lumen endotracheal tube is more efficient in providing ventilation relative to a double-lumen tube, or Univent® endotracheal tube (Vitaid, Lewiston, NY). This difference is even greater during one-lung ventilation with a double-lumen endotracheal tube. The cross-sectional area will decrease by an additional 50%. The 8.0 and 8.5 single-lumen endotracheal tubes are the most common sizes of wire-guided endobronchial blocker (WEB®; Cook, Bloomington, IN) systems placed. The equivalent outer diameter double-lumen and Univent® tube would be 32 French, 35 French, 6.0-mm internal diameter, and 7.0-mm internal diameter, respectively. As compared to a single lumen-endotracheal tube, the inner diameter is small relative to the outer diameter. When comparing the 32 French and 35 French double-lumen tubes, they have a combined internal diameter of 5.0 to 6.0 mm, respectively. The internal diameter for ventilation decreases to 3.4 to 3.5 mm and 4.3 to 4.5 mm, respectively. The equivalent diameter Univent® tubes

Dr. Arndt has received compensation for the development of the WEB® blocker from Cook (Bloomington, Indiana).

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An Alternative Way to Use Fogarty Balloon Catheter for Perioperative Lung Isolation

To the Editor:—We read with great interest the review of current techniques for perioperative lung isolation by Dr. Campos.¹ Use of the Fogarty® balloon catheter (FBC®) (Edwards Lifesciences, Irvine, CA) as a bronchial blocker to achieve lung isolation is reviewed. It is suggested that a standard endotracheal tube (ETT) of at least 6.0-mm internal diameter is required to use the FBC® for lung isolation.

We suggest an alternate way to use the FBC® for lung isolation. In our experience, under direct laryngoscopy it is possible to place the FBC independently through the vocal cords alongside the ETT. A fiberoptic bronchoscope may or may not be required to guide the FBC into the desired position. This alternate technique obviates the need for an ETT larger than 6.0-mm internal diameter and a right-angle connector with self-sealing diaphragm. Introducing the FBC independently alongside the ETT may also preclude the need for ventilation interruption for placement of the FBC in critically ill mechanically ventilated patients requiring high positive end expiratory pressure. Use of the FBC alongside the ETT instead of through the ETT also decreases the risk of dislodging the FBC that may occur when the fiberoptic bronchoscope is withdrawn alongside the FBC.

The FBC is considerably cheaper than double-lumen tubes, Univent® endotracheal tubes, and wire-guided endobronchial blockers

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Reference

1. Campos JH: Current techniques for perioperative lung isolation in adults. *ANESTHESIOLOGY* 2002; 97:1925-301

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have an internal diameter of 6.0 to 7.0 mm. The WEB® allows patient management with a large internal diameter endotracheal tube as compared to the Univent® endotracheal tube or double-lumen endotracheal tube with the same approximate outer diameter. The WEB® is now made in a 7 French adult diameter, which makes placement through 7.0 or 7.5 endotracheal tubes relatively easy.

The advantages of the WEB® are several; however, the ability to conserve internal diameter as compared to outer diameter is an important advantage. Both the Univent® and double-lumen endotracheal tube lose a large part of their internal diameter because of the extra plastic material used, which is inherent in their construction.

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Reference

1. Campos JH: Current techniques for perioperative lung isolation in adults. *ANESTHESIOLOGY* 2002; 97:1295-301

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(WEB®) for lung isolation and is readily available in operating rooms. The disadvantages include a low-volume, high-pressure cuff, and its lack of hollow center. The FBC, therefore, may increase the risk of tracheal mucosal damage if used for a prolonged time, and it cannot be used to insufflate oxygen, evacuate air, or perform pulmonary toilet. We also disagree with the author about using a fiberoptic bronchoscope suction port to evacuate air from the lung. We believe that using active suction may collapse the proximal bronchioles and bronchi and leave the distal alveoli distended, thus deteriorating the situation.

We recommend that the size of the ETT may not limit the selection of the FBC for one-lung ventilation. Introducing the FBC independently alongside the ETT may be considered for critically ill mechanically ventilated patients, and to avoid risk of FBC dislodgment.

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Reference

1. Campos, JH: Current techniques for perioperative lung isolation in adults. *ANESTHESIOLOGY* 2002; 97:1295-1301

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In Reply:—I appreciate the suggestion of Dr. Karzai of an alternative method to facilitate lung collapse while using a wire-guided endobronchial blocker (WEB®; Cook, Bloomington, IN). The various methods I discussed in my article¹ for facilitating and/or expediting lung collapse when the WEB® is in use include: (1) Removal of the wire loop after the patient is turned to a lateral decubitus position and fiberoptic confirmation demonstrate that the WEB® is correctly placed in the optimal position, so that air can be evacuated spontaneously along with the absorption atelectasis in the nondependent lung. (2) Use of the low-suction-assisted method, which basically consists of connecting the barrel of a 3-ml syringe to the wire loop site (after the wire loop has been removed) then attaching it to a low-suction device after the cuff of the blocker has been fully inflated. I do not recommend continuous suction through this channel, because of the risk of developing negative pressure pulmonary edema reported with the Univent® endotracheal tube (Vitaid, Lewiston, NY).²

Regardless of the method used to facilitate or expedite lung collapse, variability among time to lung collapse exists when using double-lumen endotracheal tubes (DLT) or bronchial blockers (Univent® and WEB®). In a recent study,³ when comparing the DLT *versus* Univent® and WEB®, it was found that the majority of patients assigned to the WEB® group required assisted suction and took longer to achieve complete lung collapse when compared to the DLT or Univent® groups (by an average of 7 min). The method proposed by Dr. Karzai raises some concerns: How long do we need to stop ventilating the patient until complete collapse is achieved? Has this method been tested in a prospective, randomized, and scientific fashion? I will assume that some patients with severe chronic obstructive pulmonary disease may not tolerate prolonged periods of apnea. Until Dr. Karzai's method to expedite lung collapse is tested in a scientific fashion defining length of time and oxygenation/saturation measurements, I would not propose using it clinically.

Another important issue that Dr. Karzai mentioned relates to maintaining the wire loop through the operation as long as is needed. In a recent study³ when comparing the left-sided DLT with the Univent® and WEB®, we reported nine total malpositions in 32 patients in the WEB® group. The first six malpositions occurred while changing from the supine to the lateral decubitus position. The wire loop was kept until the WEB® was confirmed to be in its optimal position with the fiberoptic bronchoscope after the blocker cuff was fully inflated. The other three malpositions occurred during one-lung ventilation, after the wire loop was already removed. The WEB® was repositioned under fiberoptic bronchoscopy with no difficulty and was placed in its optimal position. In addition, removing the wire loop allows the opening channel to apply continuous positive airway pressure ventilation or intermittent inflation, if required. Maintaining the wire loop in the WEB® during the operation may potentially damage the membranous airway, or the airway may accidentally be stapled into the bronchial closure, as reported when the bronchial blocker of the Univent® was stapled during a right upper lobectomy.⁴

I appreciate the comments by Dr. Arndt and would like to address the issue he has raised. I agree with the importance of comparing the inner and outer diameters of the DLT, Univent®, and single-lumen endotracheal tubes. As mentioned in his letter, one of the advantages of the WEB® is the ability to preserve the inner diameter of a single-lumen endotracheal tube while using the WEB®. However, one must

consider that a reduction of the cross-sectional area occurs when a WEB® is placed inside a single-lumen endotracheal tube. I would speculate that the presence of the WEB® may also affect air-flow resistance, which at the present time has not been tested scientifically, as opposed to DLT, Univent®, and single-lumen endotracheal tubes.⁵

I also appreciate the comments by Munir *et al.* and would like to address the alternative method for using the Fogarty® embolectomy balloon catheter (Edwards Lifesciences, Irvine, CA) through the vocal cords alongside the single-lumen endotracheal tube under direct laryngoscopy. This method of placement has been previously reported in the pediatric literature and is commonplace in some practices.⁶ I disagree with Munir *et al.* that a fiberoptic bronchoscope may not be needed for guidance while advancing the Fogarty catheter. I believe it would not be a safe practice to omit the use of the bronchoscope. If a Fogarty catheter is advanced with its stylet in place, there is the potential for tracheobronchial laceration, as previously reported.⁷ Regarding the cost, Fogarty catheters are cheap, \$4.20, when compared to the DLT (the University of Iowa acquisition cost for a left-sided DLT is \$79.20, a Univent® is \$137, a WEB® is \$107 plus \$1 for a single-lumen endotracheal tube, and a Fogarty catheter No. 8/14 F is \$75). One of the advantages that the Fogarty catheter may offer over the DLT is its balloon durability when compared with DLTs in which the tracheal cuff is more prone to tears in some cases.⁸

I believe that the use of the fiberoptic bronchoscope suction port to facilitate and expedite lung collapse is justified while using a bronchial blocker. It is doubtful that the negative pressure and flow characteristics used would be sufficient to collapse the cartilage-supported bronchi and bronchioles. Finally, to advance a Fogarty catheter independently alongside the single-lumen endotracheal tube in a patient who is critically ill and mechanically ventilated, as suggested by Munir *et al.* (usually these patients are intubated for days and have severe edema in the vocal cords), would be difficult because the upper airway would likely be edematous.

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References

1. Campos JH: Current techniques for perioperative lung isolation in adults. *ANESTHESIOLOGY* 2002; 97:1295-1301
2. Baraka A, Nawfal M, Kawkabani N: Severe hypoxemia after suction of the non-ventilated lung via the bronchial blocker lumen of the Univent tube. *J Cardiothorac Vasc Anesth* 1996; 10:694-5
3. Campos JH, Kernstine KH: A comparison of a left-sided broncho-cath® with the torque control blocker Univent and the wire-guided blocker. *Anesth Analg* 2003; 96:283-9
4. Thielmeir KA, Anwar M: Complication of the Univent tube (letter). *ANESTHESIOLOGY* 1996; 84:491
5. Slinger PD, Lesiuk L: Flow resistances of disposable double-lumen, single-lumen and Univent tubes. *J Cardiothorac Vasc Anesth* 1998; 12:142-4
6. Tan GM, Kendrick TPA: Bronchial diameters in children: Use of the Fogarty catheter for lung isolation in children. *Anaesth Int Care* 2002; 30:615-8
7. Schwartz DE, Yost CS, Larson MD: Pneumothorax complicating the use of a Univent® endotracheal tube. *Anesth Analg* 1993; 76:443-5
8. Lewis JW, Serwin JP, Gabriel FS, Bustanar M, Jacobsen G: The utility of a double-lumen tube for one-lung ventilation in a variety of noncardiac thoracic surgical procedures. *J Cardiothorac Vasc Anesth* 1992; 6:705-10

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The Laryngeal Mask Airway™ Audio Monitor. Obtaining Breath Sounds from the Laryngeal Mask Airway™: A New Device for Patient Monitoring

To the Editor:—The Laryngeal Mask Airway™ (LMA™) is a well-known method of clinical airway management now in worldwide use.¹ This letter describes the design, construction, and preliminary clinical evaluation of a new medical instrument for respiratory monitoring that attaches to the LMA™ cuff inflation line (LMA™ Audio Monitor). The device is essentially a new form of electronic stethoscope and is intended for qualitative respiratory monitoring during general anesthesia when the LMA™ is in use.

A special, leakfree microphone assembly was designed, fabricated, and pressure-tested to 300 mmHg using a miniature electric microphone (Radio Shack 30-3013, Radio Shack, Fort Worth, TX) and a shortened 3-ml plastic syringe with a Luer lock end. Epoxy glue was used to secure the microphone into the barrel of the syringe. A high-gain monaural audio amplifier was used to amplify the microphone signal for use with a headset or for computer analysis.

Following institutional approval, the system was evaluated clinically in 10 patients undergoing general anesthesia using the classic LMA with spontaneous ventilation. In all cases, attaching the device to the pilot line of the LMA was straightforward. Clearly identifiable breath sounds were heard in all cases; these were thought to be suitable for qualitative respiratory monitoring. It was observed that with normal breathing, the sounds were regular and smooth. In one patient who

developed partial airway obstruction with the LMA *in situ*, the sounds become chaotic, irregular, and intense. A sample normal recording is available as a Web Enhancement at <http://www.anesthesiology.org>.

Esophageal and precordial stethoscopes are often used during surgery for monitoring breath sounds. Some of these devices are electronic in design.^{2,3}

Potential drawbacks of these methods include relatively poor acoustic properties, awkward placement, and, in the case of esophageal stethoscopes, the remote possibility of injury to the patient. The LMA Audio Monitor allows anesthesia providers to hear clear respiratory sounds, and it works with all forms of the LMA™. Because the unit is battery-operated, and because there is no direct connection of the device to the patient, the device poses no risk to the patient. The expected value of the device is that, with experience, anesthesia providers will be able to interpret the obtained sounds to detect various normal and pathologic states. These include normal breathing, tachypnea, phonation, partial airway obstruction, wheezing, and ventilation leaks with positive pressure ventilation.

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Additional material related to this article can be found on the ANESTHESIOLOGY Web site. Go to the following address, click on Enhancements Index, and then scroll down to find the appropriate article and link. <http://www.anesthesiology.org>

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Cricoid Pressure Is Effective in Preventing Esophageal Regurgitation

To the Editor:—Cricoid pressure (Sellick's maneuver) is a routinely used technique for the prevention of pulmonary aspiration of gastric contents during anesthesia in patients at high risk of aspiration. Its effectiveness in preventing regurgitation of esophageal contents has been demonstrated in several studies^{1,2} and in cadavers.³⁻⁵ However, its effectiveness in preventing aspiration in clinical practice is debated.⁶ The following case scenario clearly demonstrates its effectiveness during clinical anesthesia.

A 21-yr-old, 75-kg man presented for repositioning of an intraocular lens under general anesthesia. He had a history of esophageal reconstruction by gastric pull-through to treat childhood achalasia. General anesthesia about 2 yr earlier for eye surgery had resulted in pulmonary aspiration of gastric contents during induction of anesthesia. No further details were available. Preoperatively, he was apprehensive about recurrent aspiration. On the day of surgery, he was given metoclopramide 10 mg and ranitidine 20 mg intravenously; he had taken nothing by mouth for 15 h previously. In the operating room, he was positioned supine with extension of the neck. The cricoid cartilage was identified while the patient was awake. Following adequate preoxygenation, continuous cricoid pressure was applied immediately before the induction of anesthesia with propofol 200 mg and succinylcholine 120 mg in a rapid sequence manner. Laryngoscopy and tracheal intubation with a 7.5 endotracheal tube were easily accomplished, the endotracheal tube cuff was inflated with 10 ml of air,

and cricoid pressure was released. This was followed immediately by the appearance of copious, greenish fluid in the mouth, which was suctioned. There were no further sequelae.

Although no prospective randomized controlled clinical studies can be done to prove its clinical efficacy, the above case illustrates that proper application of cricoid pressure is effective, at least in some patients, in the prevention of gastric aspiration from passive regurgitation.

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References

1. Wraight WJ, Chamney AR, Howells TH: The determination of an effective cricoid pressure. *Anaesthesia* 1983; 38:461-6
2. Vanner RG, O'Dwyer JP, Pryle BJ, Reynolds F: Upper oesophageal sphincter pressure and the effect of cricoid pressure. *Anaesthesia* 1992; 47:95-100
3. Salem MR, Wong AY, Fizzoti GF: Efficacy of cricoid pressure in preventing aspiration of gastric contents in paediatric patients. *Br J Anaesth* 1972; 44:401-4
4. Salem MR, Joseph NJ, Heyman HJ, Belani B, Paulsian R, Ferrara TP: Cricoid compression is effective in obliterating the esophageal lumen in the presence of a nasogastric tube. *ANESTHESIOLOGY* 1985; 63:443-6
5. Vanner RG, Pryle BJ: Regurgitation and oesophageal rupture with cricoid pressure: A cadaver study. *Anaesthesia* 1992; 47:732-5
6. Brimacombe JR, Berry AM: Cricoid pressure. *Can J Anaesth* 1997; 44:414-25

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