

JOINT TRAUMA SYSTEM CLINICAL PRACTICE GUIDELINE (JTS CPG)



Anesthesia for Trauma Patients (CPG ID: 40)

A method of anesthesia that incorporates the induction and maintenance of anesthesia into an ongoing resuscitation during surgery for a trauma patient in extremis.

Contributors

CDR Joshua Tobin, MC, USNR
 COL William Barras, AN, USA
 Capt Stephen Bree, MC, UK
 CAPT Necia Williams, MC, USN
 LtCol Craig McFarland, MC, USAR

LtCol Claire Park, RAMC
 LtCol David Steinhiser, MC, USAF
 MAJ Craig Stone, MC, CF
 CAPT Zsolt Stockinger, MC, USN

First Publication Date: 16 Jun 2014

Publication Date: 23 Jun 2016

Supersedes CPG dated 16 Jun 2014

TABLE OF CONTENTS

Background	2
Specific Considerations for Trauma Anesthesia.....	2
Pre-induction	2
Induction of Anesthesia.....	2
Maintenance of Anesthesia.....	3
Resuscitation	4
Post-operative/emergence	4
Performance Improvement (PI) Monitoring.....	5
Intent (Expected Outcomes).....	5
Performance/Adherence Measures	5
Dat Source	5
System Reporting & Frequency	6
Responsibilities	6
References	6
Appendix A: Trauma Anesthesia Checklist.....	9
Appendix B: Additional Information Regarding Off-Label Uses in CPGs.....	10

BACKGROUND

Resuscitation goals for trauma patients have undergone significant change in the past decade. Appropriate blood product transfusion ratios, use of pharmacologic adjuncts (e.g.; Tranexamic acid) and other modalities have improved survival for the wounded combatant. In the operating room, this resuscitation occurs in the context of providing an anesthetic that minimizes hemodynamic instability in the severely injured patient. It is imperative, therefore, that the anesthesiologist understands their role in the management of this resuscitation continuum. While recent review articles, checklists and textbooks have drawn attention to the role of the anesthesiologist as resuscitation consultant, there is currently no guideline for the induction, maintenance and transfer of anesthetic care of the military trauma patient in extremis.¹⁻⁴

SPECIFIC CONSIDERATIONS FOR TRAUMA ANESTHESIA

PRE-INDUCTION

- **Hypothermia** is one of the arms of the lethal triad of coagulopathy, acidosis and hypothermia.⁵ It is important, therefore, to warm the OR to greater than 30C and have a warmed intravenous (IV) line, forced air warmer, and rapid infuser with warming capability immediately available. Standard checks (e.g., anesthesia machine check, verification that airway equipment is in good working order) assure that vital equipment is ready for immediate use.
- **Establishment of a massive transfusion protocol and effective communication with the blood bank** is essential and can improve survival.⁶ The Damage Control Resuscitation CPG⁷ defines the massive transfusion protocol for the combat theater. At all roles of care, awareness of the individual MTF's on-hand resources (including walking blood bank) and applicable protocols are key considerations.
- The presence of anesthesia in the trauma bay is necessary for smooth transition of care to the OR and offers the opportunity to assist with invasive procedures. Identification of team roles prior to patient arrival facilitates effective transfer from the delivering team.

INDUCTION OF ANESTHESIA

1. Induction of anesthesia in the exsanguinating patient can be disastrous. Ongoing volume resuscitation to prevent this from occurring is critical.
2. After a patient is identified for surgery, **verification of functioning vascular access** (either intravenous or intraosseous) and placement of monitoring devices (e.g., oxygen saturation, blood pressure, and electrocardiogram) must occur quickly.
3. **Do not delay induction of the patient in extremis for placement of central access or invasive monitoring.** Placing monitors at the same time as the surgical prep and drape can save time in a crisis. A wide draping procedure with "arms out" ensures adequate surgical exposure, while affording access to the arms as needed after the start of surgery. Pre-oxygenation with four full vital capacity breaths can "de-nitrogenate" the end alveoli sufficiently to optimize oxygenation prior to rapid sequence induction. In the obtunded patient, it may not be possible to achieve four vital capacity breaths prior to induction, and one must proceed with induction relying upon apneic oxygenation.
4. There are a variety of **sedative hypnotics** available for induction of anesthesia. Standard induction dosages should be reduced and titrated to balance the induction of anesthesia with hemodynamic changes. Ketamine (1 mg/kg) will not decrease the systemic vascular resistance to the same extent as other sedative hypnotics. While Propofol is a standard induction agent, it can decrease the systemic

vascular resistance significantly. It is prudent, therefore, to use reduced doses of Propofol (0.5-1 mg/kg) in hypotensive patients. Ongoing volume resuscitation is vital to prevent vascular collapse.

5. **Neuromuscular relaxation** sufficient to facilitate endotracheal intubation can be achieved in approximately 45 seconds with succinylcholine in a standard rapid sequence induction dose (1mg/kg). Rocuronium is a non-depolarizing neuromuscular relaxant that is useful in cases where succinylcholine may be contraindicated (e.g.; burns, spinal cord injury, hyperkalemia). An increased dose of Rocuronium (1-1.2 mg/kg) can produce intubating conditions similar to succinylcholine in approximately 60 seconds.
6. **Prompt endotracheal intubation of the trachea following induction** mitigates the risk of aspiration. Rapid sequence induction (RSI) with direct laryngoscopy is a safe and effective method to secure the airway of the trauma patient.^{8,9} The efficacy of in-line stabilization during RSI is somewhat controversial; however, it remains prudent to minimize the manipulation of the cervical spine to the extent possible during laryngoscopy. Regardless, it is re-assuring to know that spinal cord injury following direct laryngoscopy rarely causes or worsens cervical spine injury.¹⁰
7. A variety of **airway adjuncts** are available to the laryngoscopist. The gum elastic bougie can be helpful in securing a challenging airway and is a low-cost, effective airway adjunct.¹¹ Video laryngoscopy can provide an improved view of the vocal cords during intubation. This does not, however, necessarily improve successful first pass intubation or result in faster time to intubation.¹² It remains prudent to have a limited number of immediately available airway adjuncts with which one is familiar, rather than a larger selection of less familiar equipment.¹³ An alternate plan, including equipment for surgical airway management, must also be immediately available. (See also [Airway Management of Traumatic Injuries](#) CPG)¹⁴
8. After endotracheal intubation of the trachea and verification of end tidal carbon dioxide, communication with the surgeon ensures that the operation proceeds in a timely fashion. **Placement of an orogastric tube** at this point may potentially decrease the risk of aspiration.

MAINTENANCE OF ANESTHESIA

- **Maintenance of anesthesia can be accomplished via an inhalational volatile agent or via a total intravenous anesthetic (TIVA).**¹⁵ Both approaches must be carefully titrated to the hemodynamic profile while assuring adequate sedation/hypnosis and analgesia. Awareness during anesthesia and the acute pain response can be mitigated during TIVA by assuring that both a sedative hypnotic (e.g., Propofol, benzodiazepine) and an analgesic (e.g.; narcotic) are being administered. Narcotic dose can be titrated to hemodynamics.
- **Adequate IV access** must be assured immediately (e.g.; large bore peripheral IV, intraosseous). Placement of additional IV access or an arterial line (if indicated for continuous monitoring of beat-to-beat blood pressure) can be undertaken without delaying the start of the operation.
- Sending a **baseline set of labs**, to include coagulation studies and base excess, at the start of the case can set a reference point for the remainder of the resuscitation. Consider validation of Point of Care testing (i.e.; iSTAT values) with traditional laboratory assays.
- The maintenance of anesthesia and the resuscitation can be guided by following the trend in **mean arterial pressure (MAP)**. While the ideal blood pressure is controversial, a MAP < 55 mmHg has been associated with acute kidney injury and myocardial injury during anesthetics for non-cardiac surgery.¹⁶ Maintaining a MAP > 55 mmHg will facilitate end organ perfusion without exacerbating any unsecured bleeding.
- **Traumatic brain injury (TBI)** represents a unique situation in which isolated episodes of hypotension can worsen mortality.¹⁷ It is, therefore, advisable to maintain systolic blood pressure > 90 mmHg in patients

with documented or suspected TBI. (See also Neurosurgery and Medical Management of Severe Head Injury CPG)¹⁸

RESUSCITATION

NOTE: See also the Damage Control Resuscitation CPG⁷

- **Ratios of FFP: PRBC** approaching 1:1 have been demonstrated to confer a survival benefit in military and civilian trauma patients.^{19,20} While the ideal ratio of FFP: PRBC remains somewhat controversial; it is fair to say that early administration of plasma and platelets is appropriate for the trauma patient in extremis.²¹ A more exhaustive discussion of **damage control resuscitation** is found elsewhere in the CPGs and is recommended reading for this subject. Communication with the surgical team regarding the progress of the resuscitation and the stage of the surgery is an important factor in overall success.
- **Tranexamic acid** is a potent synthetic lysine derivative that functions as an anti-fibrinolytic. Administration of 1 gm of tranexamic acid over 10 minutes within three hours of injury has been demonstrated to improve survival in a highly powered, randomized trial of international trauma patients.²² The initial bolus dose was followed by an infusion of 1 gm over 8 hours. A survival advantage was also demonstrated with the use of tranexamic acid in military trauma.²³
- **Hydrocortisone** is a potent mineralocorticoid that can augment blood pressure during shock states when the HPA axis is suppressed and unable to mount an effective stress response. Administration of hydrocortisone 100 mg can improve vasopressor responsiveness in critically ill trauma patients.^{24,25}
- **Massive blood transfusion** can result in hypocalcemia due to chelation of calcium by the citrate preservative in PRBCs. Administration of 1 gm calcium chloride can correct this potentially life-threatening hypocalcemia, and the hypotension associated with it.²⁶
- **Use of vasopressors** in trauma is generally associated with higher mortality.²⁷ In one analysis evaluating trauma patients who received vasopressor support, however, vasopressin was found to be the only vasopressor in which the 95% confidence interval for mortality crossed unity, suggesting non-significance.²⁸ Vasopressin is now the subject of an on-going clinical trial. The Arginine Vasopressin During the Early Resuscitation of Traumatic Shock (AVERT) Study is a phase 2 clinical trial that will evaluate the use of vasopressin supplementation in the resuscitation of trauma patients, as well as the utility of using copeptin as a biomarker for vasopressin (available at: <http://clinicaltrials.gov/ct2/show/study/NCT01611935>). In cases of refractory hypotension, a vasopressin bolus (5-10 units) followed by infusion (0.04 U/min) can be given in concert with aggressive blood product administration.
- **Timely administration of antibiotics** can decrease the incidence of post-operative infections and is part of the anesthetic resuscitation. Consider agents that will be effective against skin flora (Gram positive organisms) or, in the event of bowel injury, gastrointestinal flora (anaerobes and Gram negative organisms). The Infection Prevention CPG identifies the optimal antibiotics for multiple clinical scenarios.

POST-OPERATIVE/EMERGENCE

Low lung volume ventilation (6mL/kg) can decrease mortality in critically ill patients with the acute respiratory distress syndrome.²⁹ Even in patients who have not developed the acute respiratory distress syndrome; initiation of low lung volume ventilation can improve outcome.³⁰ Consider initiation of low lung volume ventilation in the OR.

Communication with the next role of care is vital to maintaining continuity of care. In the deployed setting this may entail a face-to-face conversation with the intensive care unit (ICU) team, or a report transmitted to a critical care air transport (CCAT) team. A detailed written report/anesthetic record documents the operative resuscitation and facilitates transition to the next role of care. Being immediately available in the post-operative period to answer any questions can clarify any issues that may arise.

PERFORMANCE IMPROVEMENT (PI) MONITORING

INTENT (EXPECTED OUTCOMES)

1. Trauma patients in the OR will maintain a body temperature > 36°C.
2. Anesthesia following major trauma will be induced and maintained with < 20% drop in initial blood pressure.
3. Any incidence of awareness under anesthesia during TIVA or GETA will be documented.
4. No trauma patient will experience a sustained MAP < 50 mmHg during maintenance of anesthesia.
5. Patients undergoing massive transfusion will receive blood products in a PRBC: FFP: Platelet ratio approaching 1:1:1.
6. Calcium chloride will be administered to hypotensive patients who have received more than 4 units of PRBC.
7. Antibiotics will be administered to all patients within 30 minutes of incision.
8. The anesthesia record will reflect an accurate description of the resuscitation that is undertaken in the OR.

PERFORMANCE/ADHERENCE MEASURES

1. Trauma patients in the OR maintained a body temperature > 36°C.
2. Anesthesia following major trauma will be induced and maintained with < 20% drop in initial blood pressure.
3. Any incidence of awareness under anesthesia during TIVA or GETA was documented.
4. Patients undergoing massive transfusion received blood products in a PRBC: FFP: Platelet ratio approaching 1:1:1.
5. Calcium chloride was administered to hypotensive patients who received more than 4 units of PRBC.
6. Antibiotics were administered to all patients within 30 minutes of incision.

DATA SOURCE

- Patient Record
- Department of Defense Trauma Registry (DoDTR)

SYSTEM REPORTING & FREQUENCY

The above constitutes the minimum criteria for PI monitoring of this CPG. System reporting frequency will be performed annually; additional PI monitoring and system reporting may be performed as needed.

The system review and data analysis will be performed by the Joint Trauma System (JTS) Director and the JTS Performance Improvement Branch.

RESPONSIBILITIES

It is the Chief of Trauma or equivalent's responsibility to ensure familiarity, appropriate compliance and PI monitoring at the local level with this CPG.

REFERENCES

1. Tobin JM, Grabinsky A, McCunn M, Pittet JF, Smith CE, Murray MJ, et al. A checklist for trauma and emergency anesthesia. *Anesth analg.* 2013;117(5):1178-84. Epub 2013/10/11. (APPENDIX A)
2. Tobin JM, Varon AJ. Review article: update in trauma anesthesiology: perioperative resuscitation management. *Anesth analg.* 2012;115(6):1326-33. Epub 2012/07/06.
3. Russell R, Bess A, eds. Joint Service Publication 999 - Clinical Guidelines for Operations. 3rd ed. Leeds, UK: UK Ministry of Defence; 2012.
4. Buckenmaier C, Mahoney PF, eds. Combat Anesthesia: The First 24 Hours. In: Banks DE, ed. Textbooks of Military Medicine. Fort Sam Houston, TX: Borden Institute; 2015.
5. Rotondo MF, Zonies DH. The damage control sequence and underlying logic. *Surg Clin North Am.* 1997;77(4):761-77. Epub 1997/08/01.
6. Riskin DJ, Tsai TC, Riskin L, Hernandez-Boussard T, Purtill M, Maggio PM, et al. Massive transfusion protocols: the role of aggressive resuscitation versus product ratio in mortality reduction. *J Am Coll Surg.* 2009;209(2):198-205. Epub 2009/07/28.
7. Joint Trauma System, Damage Control Resuscitation, 02 Feb 2017. [http://jts.amedd.army.mil/assets/docs/cpgs/JTS_Clinical_Practice_Guidelines_\(CPGs\)/Damage_Control_Resuscitation_03_Feb_2017_ID18.pdf](http://jts.amedd.army.mil/assets/docs/cpgs/JTS_Clinical_Practice_Guidelines_(CPGs)/Damage_Control_Resuscitation_03_Feb_2017_ID18.pdf) Accessed Mar 2018
8. Sollid SJ, Lossius HM, Soreide E. Pre-hospital intubation by anaesthesiologists in patients with severe trauma: an audit of a Norwegian helicopter emergency medical service. *Scand J Trauma Resusc Emerg Med.* 2010;18:30. Epub 2010/06/16.
9. Stephens CT, Kahntroff S, Dutton RP. The success of emergency endotracheal intubation in trauma patients: a 10-year experience at a major adult trauma referral center. *Anesth analg.* 2009;109(3):866-72. Epub 2009/08/20.
10. Shatney CH, Brunner RD, Nguyen TQ. The safety of orotracheal intubation in patients with unstable cervical spine fracture or high spinal cord injury. *Am J Surg.* 1995;170(6):676-9; discussion 9-80. Epub 1995/12/01.
11. Tobin JM. Usage and efficacy of airway adjuncts in an emergency intubation kit. *Emerg Med Australas.* 2011;23(4):514-5. Epub 2011/08/10.

12. Griesdale DE, Liu D, McKinney J, Choi PT. Glidescope((R)) video-laryngoscopy versus direct laryngoscopy for endotracheal intubation: a systematic review and meta-analysis. *Can J Anaesth.* 2012; 59(1):41-52. Epub 2011/11/02.
13. Marco CA, Marco AP. Airway adjuncts. *Emerg Med Clin North Am.* 2008;26(4):1015-27, x. Epub 2008/12/09.
14. Joint Trauma System, Airway Management of Traumatic Injuries, 17 Jul 2017. [http://jts.amedd.army.mil/assets/docs/cpgs/JTS_Clinical_Practice_Guidelines_\(CPGs\)/Airway_Management_Traumatic_Injuries_17_Jul_2017_ID39.pdf](http://jts.amedd.army.mil/assets/docs/cpgs/JTS_Clinical_Practice_Guidelines_(CPGs)/Airway_Management_Traumatic_Injuries_17_Jul_2017_ID39.pdf) Accessed Mar 2018.
15. Barras P, McMasters J, Grathwohl K, Blackbourne LH. Total intravenous anesthesia on the battlefield. *US Army Medical Department journal.* 2009:68-72.
16. Walsh M, Devereaux PJ, Garg AX, Kurz A, Turan A, Rodseth RN, et al. Relationship between intraoperative mean arterial pressure and clinical outcomes after noncardiac surgery: toward an empirical definition of hypotension. *Anesthesiology.* 2013;119(3):507-15. Epub 2013/10/19.
17. Chesnut RM, Marshall LF, Klauber MR, Blunt BA, Baldwin N, Eisenberg HM, et al. The role of secondary brain injury in determining outcome from severe head injury. *J Trauma.* 1993;34(2):216-22. Epub 1993/02/01.
18. Joint Trauma System, Neurosurgery and Severe Head Injury, 02 Mar 2017, [http://jts.amedd.army.mil/assets/docs/cpgs/JTS_Clinical_Practice_Guidelines_\(CPGs\)/Neurosurgery_Severe_Head_Injury_02_Mar_2017_ID30.pdf](http://jts.amedd.army.mil/assets/docs/cpgs/JTS_Clinical_Practice_Guidelines_(CPGs)/Neurosurgery_Severe_Head_Injury_02_Mar_2017_ID30.pdf) Accessed Mar 2018.
19. Borgman MA, Spinella PC, Perkins JG, Grathwohl KW, Repine T, Beekley AC, et al. The ratio of blood products transfused affects mortality in patients receiving massive transfusions at a combat support hospital. *J Trauma.* 2007;63(4):805-13. Epub 2007/12/20.
20. Holcomb JB, Wade CE, Michalek JE, Chisholm GB, Zarzabal LA, Schreiber MA, et al. Increased plasma and platelet to red blood cell ratios improves outcome in 466 massively transfused civilian trauma patients. *Ann Surg.* 2008;248(3):447-58. Epub 2008/09/16.
21. Holcomb JB, Tilley BC, Baraniuk S, Fox EE, Wade CE, Podbielski JM et al. Transfusion of plasma, platelets, and red blood cells in a 1:1:1 vs a 1:1:2 ratio and mortality in patients with severe trauma: the PROPPR randomized clinical trial. *JAMA.* 2015;313(5):471-82. doi:10.1001/jama.2015.12
22. Shakur H, Roberts I, Bautista R, Caballero J, Coats T, Dewan Y, et al. Effects of tranexamic acid on death, vascular occlusive events, and blood transfusion in trauma patients with significant haemorrhage (CRASH-2): a randomised, placebo-controlled trial. *Lancet.* 2010;376(9734):23-32. Epub 2010/06/18.
23. Morrison JJ, Dubose JJ, Rasmussen TE, Midwinter MJ. Military Application of Tranexamic Acid in Trauma Emergency Resuscitation (MATTERs) Study. *Arch Surg.* 2012;147(2):113-9. Epub 2011/10/19.
24. Hoen S, Asehnoune K, Brailly-Tabard S, Mazoit JX, Benhamou D, Moine P, et al. Cortisol response to corticotropin stimulation in trauma patients: influence of hemorrhagic shock. *Anesthesiology.* 2002;97(4):807-13. Epub 2002/10/03.
25. Hoen S, Mazoit JX, Asehnoune K, Brailly-Tabard S, Benhamou D, Moine P, et al. Hydrocortisone increases the sensitivity to alpha1-adrenoceptor stimulation in humans following hemorrhagic shock. *Crit Care Med.* 2005;33(12):2737-43. Epub 2005/12/15.

26. Ho KM, Leonard AD. Concentration-dependent effect of hypocalcaemia on mortality of patients with critical bleeding requiring massive transfusion: a cohort study. *Anaesth Intensive Care*. 2011;39(1):46-54. Epub 2011/03/08.
27. Plurad DS, Talving P, Lam L, Inaba K, Green D, Demetriades D. Early vasopressor use in critical injury is associated with mortality independent from volume status. *J Trauma*. 2011;71(3):565-70; discussion 70-2. Epub 2011/09/13.
28. Sperry JL, Minei JP, Frankel HL, West MA, Harbrecht BG, Moore EE, et al. Early use of vasopressors after injury: caution before constriction. *J Trauma*. 2008;64(1):9-14. Epub 2008/01/12.
29. Ventilation with lower tidal volumes as compared with traditional tidal volumes for acute lung injury and the acute respiratory distress syndrome. The Acute Respiratory Distress Syndrome Network. *N Engl J Med*. 2000;342(18):1301-8. Epub 2000/05/04.
30. Serpa Neto A, Cardoso SO, Manetta JA, Pereira VG, Esposito DC, Pasqualucci Mde O, et al. Association between use of lung-protective ventilation with lower tidal volumes and clinical outcomes among patients without acute respiratory distress syndrome: a meta-analysis. *JAMA*. 2012;308(16):1651-9. Epub 2012/10/25.

APPENDIX A: TRAUMA ANESTHESIA CHECKLIST

BEFORE PATIENT ARRIVAL

- Room temperature > 30°C
- Warm IV line
- Machine check
- Airway equipment check
- Emergency medication check
- Blood Bank notified to have blood available per unit SOP

PATIENT ARRIVAL

- Patient identified for surgery as soon as possible
- Blood Bank notified to deliver blood per unit SOP
- Ensure large bore IV or CVC access
- Monitors (SaO₂, BP, ECG)
- Pre-oxygenation

INDUCTION

- Sedative hypnotic (Ketamine vs. Propofol)
- Neuromuscular blockade (Rocuronium vs. succinylcholine)

INTUBATION

(Per Airway Management CPG)

- (+) ETCO₂
- Place orogastric tube

ANESTHETIC

- Consider TIVA
- (Volatile anesthetic and/or benzodiazepine) + narcotic
- Insert additional IV access and/or arterial line if needed

RESUSCITATION

(per Damage Control Resuscitation CPG)

- Send baseline labs, type and cross if not yet done
- Follow MAP trends
- Goal FFP: PRBC: Plt 1:1:1 if Massive Transfusion
- Goal urine output 0.5-1.0 mL/kg/hr
- Consider TXA if <3 hours from injury and indicators for Massive Transfusion identified
- Consider calcium chloride 1 gm
- Consider hydrocortisone 100 mg
- Consider vasopressin 5-10 IU
- Administer appropriate antibiotics
- Special considerations for TBI as indicated in *Severe Head Injury CPG*

CLOSING/POST-OPERATIVE

- Low volume ventilation per *Acute Respiratory Failure CPG*

APPENDIX B: ADDITIONAL INFORMATION REGARDING OFF-LABEL USES IN CPGS

PURPOSE

The purpose of this Appendix is to ensure an understanding of DoD policy and practice regarding inclusion in CPGs of “off-label” uses of U.S. Food and Drug Administration (FDA)–approved products. This applies to off-label uses with patients who are armed forces members.

BACKGROUND

Unapproved (i.e., “off-label”) uses of FDA-approved products are extremely common in American medicine and are usually not subject to any special regulations. However, under Federal law, in some circumstances, unapproved uses of approved drugs are subject to FDA regulations governing “investigational new drugs.” These circumstances include such uses as part of clinical trials, and in the military context, command required, unapproved uses. Some command requested unapproved uses may also be subject to special regulations.

ADDITIONAL INFORMATION REGARDING OFF-LABEL USES IN CPGS

The inclusion in CPGs of off-label uses is not a clinical trial, nor is it a command request or requirement. Further, it does not imply that the Military Health System requires that use by DoD health care practitioners or considers it to be the “standard of care.” Rather, the inclusion in CPGs of off-label uses is to inform the clinical judgment of the responsible health care practitioner by providing information regarding potential risks and benefits of treatment alternatives. The decision is for the clinical judgment of the responsible health care practitioner within the practitioner-patient relationship.

ADDITIONAL PROCEDURES**Balanced Discussion**

Consistent with this purpose, CPG discussions of off-label uses specifically state that they are uses not approved by the FDA. Further, such discussions are balanced in the presentation of appropriate clinical study data, including any such data that suggest caution in the use of the product and specifically including any FDA-issued warnings.

Quality Assurance Monitoring

With respect to such off-label uses, DoD procedure is to maintain a regular system of quality assurance monitoring of outcomes and known potential adverse events. For this reason, the importance of accurate clinical records is underscored.

Information to Patients

Good clinical practice includes the provision of appropriate information to patients. Each CPG discussing an unusual off-label use will address the issue of information to patients. When practicable, consideration will be given to including in an appendix an appropriate information sheet for distribution to patients, whether before or after use of the product. Information to patients should address in plain language: a) that the use is not approved by the FDA; b) the reasons why a DoD health care practitioner would decide to use the product for this purpose; and c) the potential risks associated with such use.