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Severinghaus and the author developed as physician-scientists from initially addressing concrete goals to realizing failures, establishing networks, and finally asking questions more about *why* rather than *how*. This memorial lecture highlights this progression and their contributions to science.

### edicine

#### INCE

#### omuscular Monitoring in Clinical Practice: actice Change Initiative L. Williams, N. A. Hanson, C. C. Blackmore,

fessional practice initiative began with the goal of n-of-four ratio greater than or equal to 0.90 for all ndepolarizing neuromuscular blocking drug. This sment of the implementation of documenting trainer than or equal to 0.9 before extubation improved of cases in November 2016 to 93% (250 of 269) of 2020. Attaining this endpoint required not only placing tor in each anesthetizing location but also ongoing and follow-up. SUPPLEMENTAL DIGITAL CONTENT IS TFXT

# or without N-Acetvlcvsteine for the Prevention of after Thoracic Surgery: A Double-blind, Randomized

D. Amar, H. Zhang, M. K. Chung, K. S. Tan, D. Desiderio, B. J. Park,	
A. Pedoto, N. Roistacher, J. M. Isbell, D. Molena, G. L. Milne,	
B. F. Meyers, G. W. Fischer, V. W. Rusch, D. R. Jones	916

d randomized trial of noncardiac thoracic surgery patients was done to test the hypothesis that the addition of N-acetylcysteine to concurrent amiodarone administration would reduce the incidence of postoperative atrial fibrillation when compared with placebo being concurrently administered with amiodarone. The study was halted midway for futility, as there was no difference in postoperative atrial fibrillation in the patients who received N-acetylcysteine plus amiodarone versus the patients who received placebo plus amiodarone.

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ON THE COVER: Residual neuromuscular blockade can be avoided with quantitative neuromuscular monitoring. In this issue of ANESTHESIOLOGY, Weigel et al. detail a professional practice initiative they conducted to attain documented train-of-four ratios greater than or equal to 0.90 in all patients to improve patient outcomes through reducing residual paralysis. In an accompanying editorial, Lane-Fall tells us how this new article demonstrates much about what anesthesiology has to learn from implementation science and quality improvement. Cover Illustration: A. Johnson, Vivo Visuals Studio.

- Weigel et al.: Quantitative Neuromuscular Monitoring in Clinical Practice: A Professional Practice Change Initiative, p. 901
- Lane-Fall: What Anesthesiology Has to Learn from Implementation Science and Quality Improvement, p. 875

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$\oplus$	Intraoperative Hypotension and Acute Kidney Injury, Stroke,
◙	and Mortality during and outside Cardiopulmonary Bypass:
ら	A Retrospective Observational Cohort Study

M. A. de la Hoz, V. Rangasamy, A. B. Bastos, X. Xu, V. Novack, B. Saugel, B. Subramaniam......927

Among 4,984 patients undergoing cardiac surgery at a single tertiary care center between 2008 and 2016, 256 (5.1%) experienced the primary outcome of stroke (66, 1.3%), acute kidney injury (125, 2.5%), or mortality (109, 2.2%). Each 10 min of hypotension (mean arterial pressure less than 65 mmHg) during, before, or after cardiopulmonary bypass was associated with an increased odds ratio of 1.06 (95% Cl, 1.03 to 1.10; P = 0.001). Intraoperative hypotension, even if it occurs outside of cardiopulmonary bypass, is independently associated with stroke, acute kidney injury, or death after cardiac surgery. *SUPPLEMENTAL DIGITAL CONTENT IS AVAILABLE IN THE TEXT* 

# Referral Indications for Malignant Hyperthermia Susceptibility Diagnostics in Patients without Adverse Anesthetic Events in the Era of Next-generation Sequencing

The hypothesis that there is an increased referral to malignant hyperthermia units of patients without a personal or family history of adverse anesthetic events suspected to be malignant hyperthermia was tested in a retrospective multicenter cohort study. The proportion of patients referred without a personal or family history of adverse anesthetic events increased from 28.4% (61 of 215) between 2010 and 2014 to 43.6% (133 of 305) between 2015 and 2019. Patients with a personal or family history of adverse anesthetic events were more frequently diagnosed as malignant hyperthermia– susceptible (133 of 220; 60.5%) than those without (47 of 120; 39.2%). *SUPPLEMENTAL DIGITAL CONTENT IS AVAILABLE IN THE TEXT* 

#### **BASIC SCIENCE**

# Midazolam at Low Nanomolar Concentrations Affects Long-term Potentiation and Synaptic Transmission Predominantly *via* the α,-γ-Aminobutyric Acid Type A Receptor Subunit in Mice

X. Puig-Bosch, S. Bieletzki, H. U. Zeilhofer, U. Rudolph, B. Antkowiak, G. Rammes.......954

Using a combination of  $\gamma$ -aminobutyric acid type A (GABA<sub>A</sub>)  $\alpha$ -receptor subunit knock-in mice revealed that low concentrations (10 nM) of midazolam blocked long-term potentiation in the hippocampal slice preparation predominantly *via*  $\alpha_1$ -GABA<sub>A</sub> receptors. Electrophysiologic recordings in neocortical slice cultures imply a dominant role for the  $\alpha_1$  subtype in governing inhibitory postsynaptic current kinetics at nanomolar concentrations, midazolam enhances synaptic transmission of GABA<sub>A</sub> receptors *via* targeting  $\alpha_1$  subtypes and provides mechanistic explanation for the drug's sedative and amnestic action. *SUPPLEMENTAL DIGITAL CONTENT IS AVAILABLE IN THE TEXT* 

### Pain Medicine

CME

#### **CLINICAL SCIENCE**

	Basal Infusion versus Automated Boluses and a Delayed Start Timer	
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	L Finneran IV E T Said B P Curran M W Swisher T B Black	

J. J. FINNERAN IV, E. I. Sald, B. P. Curran, IVI. W. Swisner, J. R. Black,	
R. A. Gabriel, J. F. Sztain, W. B. Abramson, B. Alexander,	
M. C. Donohue, A. Schaar, B. M. Ilfeld	970

Patients undergoing foot or ankle surgery received popliteal–sciatic catheter-reservoir systems delivering ropivacaine by continuous infusion or by a bolus of anesthetic every 2 h. Those patients receiving bolus anesthetic experienced better pain control and effects of longer duration than those receiving continuous infusions.

#### **Education**

#### IMAGES IN ANESTHESIOLOGY

Labeled Surgical Caps: A Tool to Improve Perioperative Communication

B. J. Wong, A. K. Nassar, S. N. Goldhaber-Fiebert	3
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## **CLINICAL FOCUS REVIEW**



Age-adjusted fraction of minimum alveolar concentration derived from end-tidal anesthetic partial pressure measurement remains a useful drug advisory display to help prevent awareness if interpreted with proper understanding of the quantal and probabilistic nature of minimum alveolar concentration, semantics, drug interactions, and hysteresis.

#### **REVIEW ARTICLE**

#### Historical and Modern Evidence for the Role of Reward Circuitry in Emergence

M. Heshmati, M. R. Bruchas	
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This review explores the integration of advanced systems neuroscience approaches into translational anesthesia research to elucidate the important role of mesolimbic brain reward circuitry in emergence from general anesthesia.

#### Advances in Neuroimaging and Monitoring to Defend Cerebral Perfusion in Noncardiac Surgery

J. P. Fanning, S. F. Huth, C. Robba, S. M. Grieve, D. Highton......1015

The authors present an introduction to the emerging roles of neuromonitoring in optimizing perioperative care through guiding intraoperative hemodynamic management, improving surgical risk stratification, and enhancing diagnosis of postoperative neurologic sequelae.

#### MIND TO MIND

#### Swan Song

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