Journal on Cataract and Refractive Surgery

Judicious Benzodiazepine Administration for Older Patients with Preoperative Anxiety

Katie J. Schenning

Associate Professor of Anesthesiology and Perioperative Medicine, School of Medicine

Corresponding author:

Katie J. Schenning,

Associate Professor of Anesthesiology and Perioperative Medicine, School of Medicine

Received Date: 03 Janu 2024 Accepted Date: 20 Janu 2024 Published Date: 26 Janu 2024

Citation:

Katie J. Schenning. Judicious Benzodiazepine Administration for Older Patients with Preoperative Anxiety. Journal on Cataract and Refractive Surgery 2024.

1.Abstract

In this Pro-Con commentary article, we discuss the risks and benefits of giving older patients with preoperative anxiety benzodiazepines. The pro side starts by emphasizing how important it is to treat anxiety before surgery and how benzodiazepines are the best way to do so. The opposing argument made by the opponents is that there are numerous options for treating anxiety before surgery without simultaneously raising the risk of devastating complications like postoperative delirium. Both sides call for more high-quality studies to find the best ways to reduce preoperative anxiety in older people, improve outcomes, and cut down on morbidity. Preoperative anxiety is emotionally distressing, and patients desire anxiolytic medications and the support of their anesthesiologist in coping with the anxiety.[1,10] This may be why preoperative benzodiazepine administration to older patients remains commonplace despite many multidisciplinary clinical guidelines recommending avoidance.[11-13] In fact, recent studies have shown that 38% to 65% of older patients received benzodiazepines during anesthesia care.[14,15] The recommendation to avoid pre Postoperative delirium is the most common complication in older patients and is associated with severe outcomes such as prolonged hospitalization, functional decline, institutionalization, mortality, a worse cognitive trajectory after surgery, and the development of dementia.[16-19] Recent research has questioned whether preoperative benzodiazepine administration for anxiolysis is associated with increased delirium.[20] Unfortunately, there is a lack of high-quality evidence to determine whether a single dose of a preoperative benzodia Based on the current state of the literature, we discuss two opposing arguments for and against the

judicious administration of benzodiazepines for preoperative anxiolysis in older patients in this Pro-Con commentary article. Pro: Preoperative Benzodiazepines for Anxiolysis in Older Patients Should Be Given Carefully There Are Good Reasons to Give Preoperative Benzodiazepines to All Older Patients. One recent study showed that the majority of patients find anxiety emotionally distressing and desire the support of the anesthesiologist in coping with the anxiety.[1] Another study showed that the majority of patients desired anxiolytic medication whether or not they perceived themselves as anxious.[10] Our subsequent argument details the lack of evidence that a single, preoperative anxiolytic benzodiazepine when given judiciously causes harm. Untreated preoperative anxiety is associated with significant morbidity, including increased We explain in detail why appropriate treatment of high anxiety offers significant benefits and why alternative anxiolytic methods may not be suitable for all patients.

Connection Between Preoperative Benzodiazepine and Postoperative Ridiculousness in More established Grown-ups

The American Geriatrics Society (AGS) in 2015 and the European Culture of Anaesthesiology in 2017 gave the primary postoperative ridiculousness rules that suggested the act of benzodiazepine evasion in more established patients.[21,22] Nonetheless, the nature of the logical proof legitimizing such proposals was low.[21,22] These proposals depended on the consideration of benzodiazepines on the 2012 AGS Brews Models list because of proof of mischief found in patients who take benzodiazepines consistently and the relationship between long haul preoperative use and clinic delirium.[23,24] The Lagers Standards presents a rundown of possibly unseemly drugs regularly refreshed by the AGS, where the prescriptions may not be the most secure or most Con: Benzodiazepines should be avoided in older patients who are anxious before surgery. Doctors weigh the risks and benefits of a treatment with and for their patients based on scientific evidence. We acknowledge that anxiety before surgery is linked to actual and undesirable outcomes. Nevertheless, the current state of the literature does not support a benefit that outweighs the risk of administering benzodiazepines to older surgical patients for anxiety. Our contention underneath subtleties why the "supportive of benzodiazepine" studies might not have legitimacy for more established careful patients. Additionally, there is a dearth of evidence regarding the advantages of benzodiazepines for older adults. Postoperative delirium, on the other hand, is the most common complication in older surgical patients and is associated with severe outcomes such as an increase in morbidity and mortality, a prolonged stay in the hospital, functional decline, institutionalization, and a significant increase in cost. A recent study showed that postoperative delirium is associated with the development of posttraumatic stress disorder, another long-term consequence for older surgical patients.[51] Given the magnitude and severity of incident

postoperative delirium, combined with limited treatment options, we argue that anesthesia providers should do everything in their power to decrease the risk of delirium, including avoiding perioperative benzodiazepines for anxiolysis. Delirium is an independent risk factor for a worse cognitive trajectory after surgery, as well as We advise anesthesiologists to: (1) investigate the root of preoperative anxiety in elderly patients; and (2) employ one of the many safer methods for managing preoperative anxiety as needed.

Connection Between Preoperative Benzodiazepine and Postoperative Ridiculousness in More established Grown-ups

While the proof areas of strength for is midazolam increments ridiculousness in the ICU, we recognize that the proof in the postoperative wooziness writing isn't as clear. Midazolam's postoperative delirium rate was compared to that of other sedative and anesthetics in a recent network meta-analysis. 39 trials and 5991 participants made up the study; Participants who were at least 65 years old comprised more than half of the sample population. Midazolam was linked to a significantly higher risk of postoperative delirium than placebo (odds ratio [OR], 2.62; The 95 percent confidence interval (CI) is 1.07-6.43. Dexmedetomidine, on the other hand, had a lower risk of postoperative delirium compared to placebo (OR, 0.44; ketamine (OR, 0.30; 95 percent CI, 0.30-0.64), as well as 95% CI, 0.09–0.99).[52]A recent study examined a national dataset to determine the effect of midazolam on postoperative outcomes in order to address the lack of population-level data on perioperative midazolam use in orthopedic surgery. They discovered that more than 75% of the nearly 3 million total knee and hip arthroplasty patients received midazolam perioperatively. The adjusted OR for in-hospital falls increased when midazolam was administered. Additionally, the adjusted odds of postoperative delirium, naloxone use, and pulmonary complications were significantly increased when midazolam and gabapentinoids were used concurrently.[53] Although our colleagues' argument in favor of preoperative benzodiazepines will point to recent research that suggests no association between preoperative midazolam and postoperative delirium, the cited studies do have some limitations. The use of benzodiazepines during the immediate postoperative period was not examined by the European BioCog project. Instead, they looked at the connection between delirium and a group of long-term home medications, such as antibiotics, platelet inhibitors, and sedatives. Any reasonable person would agree that this study was not intended to assess the act of involving benzodiazepines for preoperative tension. We use the term "association" and note that the study was a secondary analysis of three pooled data sets. Another group specifically addressed midazolam used as a premedication and its association with postoperative delirium[20]. A single assessment from 9 a.m. to 12 p.m. on the first two postoperative days was all that was collected for delirium in this study. As a result, it is highly likely that cases of delirium were not diagnosed, which could have skewed the results of the study. We agree and eagerly await these crucial results as the authors acknowledge this limitation and refer us to a recently completed RCT to provide the definitive answer.[26]

A 2016 Cochrane review found no high-quality evidence that midazolam reduced preoperative anxiety compared to placebo.[54] Additionally, the majority of studies that examined the relationship between preoperative anxiety and postoperative outcomes excluded older adults. There is a lack of literature on the topic of the benefits of preoperative benzodiazepines for older adults. For instance, lorazepam premedication was found to have a shorter time to extubation and a lower rate of early cognitive recovery in two randomized, placebo-controlled clinical trials [55,56]. These studies also found that lorazepam premedication did not improve postoperative quality of recovery or patient satisfaction. However, these trials were conducted on patients younger than 70 years old, so their results may not apply to older patients. It is important to note that the studies cited by our pro-benzodiazepine colleagues only included patients under the age of 60.[3–9] When taking into account the diverse older surgical population, it is possible that patients with subtle preexisting conditions like frailty or cognitive impairment will be even more sensitive to benzodiazepines than younger patients. Although we did not find studies directly comparing preoperative anxiety in older versus younger populations, there are conditions where older surgical patients are known to have a different postoperative course than their younger counterparts, including postoperative pain and postoperative nausea and vomiting.[57,58] In a study of patients with an average age of 78 years undergoing cataract surgery with topical anesthesia, a double-blind controlled clinical trial found that intravenous midazolam did not significantly reduce pain or anxiety compared to topical an

There Are Safer Options for Older Surgical Patients with Preoperative Anxiety While preprocedure anxiety is said to be common, its severity varies. We recognize that a few more seasoned grown-ups may without a doubt be restless before medical procedure. Luckily, anesthesiologists have numerous nonpharmacologic and pharmacologic systems for anxiolysis past benzodiazepines. The first step is to build trust between the patient and the doctor. A recent meta-analysis found that psychological preparation for surgery, including providing information and relaxation techniques, may be beneficial for postoperative pain outcomes.[63] Several groups found that showing patients an informational video at the preoperative clinic visit effectively managed preoperative anxiety.[64] Patients who were shown a video had decreased anxiety at multiple time points, including immediately preoperatively and postoperatively.[65] Patients may need education about their condition, the procedure, and the anesthetics. We might discover that the most anxiety-inducing concern is simple to address if we ask the patient about it. For instance, a lot of patients want to know if they will be awake when the urinary catheter is inserted. According to one study, older patients were anxious the day before surgery due to a lack of anesthesia knowledge. As perioperative physicians, we are able to acknowledge the patient's anxiety, normalize their feelings, and provide the opportunity for questions. Their primary concerns were failure to awaken after surgery and postoperative pain[66].

In a recent systematic review, nonpharmacologic approaches to anxiety management prior to surgery were examined. An RCT compared intravenous midazolam with music through noise-canceling headphones

in patients undergoing preoperative nerve block placements. The authors came to the conclusion that music, aromatherapy, and acupuncture effectively reduced both preoperative anxiety and postoperative pain in women undergoing breast cancer surgery[67]. In a study evaluating the effects of chewing gum with fasting on preoperative anxiety, gynecologic surgery patients were randomized to either the conventional fasting group or the chewing gum with fasting group.[44] A clinical trial of patients (n = 159) scheduled for surgery with general anesthesia found that patients who received nature sounds or relaxation exercises the day before surgery had significantly decreased preoperative anxiety compared to the nonintervention control group.[68] The gum group had significantly less preoperative anxiety than the control group, and there was no significant difference between the groups in the gastric fluid analysis.[69] Anxiety that does not respond to nonpharmacologic treatment may need to be treated with medication. A 2020 Cochrane Survey observed that melatonin was more gainful for preoperative tension than placebo.[35] a similar audit found comparative anxiolysis when preoperative melatonin was contrasted with benzodiazepines. Ondansetron, acetaminophen, gabapentin, or opioids are common oral premedications, even though oral melatonin must be taken several hours before the procedure. Alpha-2 agonists like dexmedetomidine can also alleviate preoperative anxiety and can be continued during the procedure, as demonstrated in a recent study of patients undergoing carotid stent procedures.[72] Some cardiovascular effects, such as a decreased heart rate, may actually be beneficial, particularly for patients with coronary artery disease. Preoperative betablockers were found to have benefits for antianxiety in several earlier studies.[70,71]

We Suggest Evasion of Midazolam in More established Grown-ups

While we Anticipate Geriatric-explicit Examinations Regardless of numerous multidisciplinary clinical rules recommending they are avoided,[11-13] preoperative benzodiazepine use stays high in many practices in the US. When limiting the question to patients 65 and older, 57% received a benzodiazepine during anesthesia care in one large academic health system for patients 50 and older undergoing major, noncardiac surgery.[14] Midazolam was administered perioperatively to more than 38% of patients 65 and older undergoing inpatient, noncardiac surgery at another large academic hospital.[15] This is a significant amount of use of a drug with very serious potential risks. We agree that benzodiazepines like midazolam are appealing due to their short duration, rapid onset, and amnesic effects. Midazolam, on the other hand, does not outweigh the risks. We need evidence that (1) older adults are significantly anxious prior to surgery despite other strategies and (2) midazolam is not associated with delirium before we can recommend its use. Numerous different drugs and devices assist more established careful patients with managing tension. According to a recent study, 59% of patients with preoperative anxiety would appreciate support from an anesthesiologist in coping with their anxiety. [1] As perioperative physicians, anesthesiologists must consider our capacity to do more than just administer medications to comfort our older patients.

In conclusion, we are of the opinion that giving a benzodiazepine to an older adult prior to surgery is hardly necessary. The majority of patients receive little benefit, and there is a possibility of serious side effects. Let's say that treating a senior patient with a benzodiazepine is deemed necessary. In that case, it is essential to think about lowering the dose for this patient group in comparison to a younger group, just as we do for other anesthetics. At last, for the individuals who will keep controlling midazolam to more seasoned careful patients, we propose that patients ought to be taught about likely secondary effects, including mental disability, respiratory concealment, and amnesia. Like that, the patient can go with an educated choice and choose to renounce the medicine.

4. Conclusions

The continued use of perioperative benzodiazepines in elderly surgical patients, despite the growing number of published clinical guidelines recommending their avoidance, was the impetus for this pro- and conarticle. Anaesthesiologists must evaluate their patients and weigh the advantages and disadvantages of any anxiolytic strategy, as outlined in both arguments. The Table provides an outline of the arguments as a whole. Last but not least, all of the authors call for high-quality research into the most efficient method for administering anxiolysis to patients, enhancing outcomes and decreasing morbidity.

References

- Drews T, Franck M, Radtke FM, et al. Postoperative delirium is an independent risk factor for posttraumatic stress disorder in the elderly patient: a prospective observational study. Eur J Anaesthesiol. 2015;32:147–151.
- Cui Y, Li G, Cao R, Luan L, Kla KM. The effect of perioperative anesthetics for prevention of postoperative delirium on general anesthesia: a network meta-analysis. J Clin Anesth. 2020;59:89–98.
- Athanassoglou V, Cozowicz C, Zhong H, et al. Association of perioperative midazolam use and complications: a population-based analysis. Reg Anesth Pain Med. 2022;47:228.
- 4. Conway A, Rolley J, Sutherland JR. Midazolam for sedation before procedures. Cochrane Database Syst Rev. 2016;2016:Cd009491.
- Mijderwijk H, van Beek S, Klimek M, Duivenvoorden HJ, Grüne F, Stolker RJ. Lorazepam does not improve the quality of recovery in day-case surgery patients: a randomised placebo-controlled clinical trial. Eur J Anaesthesiol. 2013;30:743–751.
- Maurice-Szamburski A, Auquier P, Viarre-Oreal V, et al; PremedX Study Investigators. Effect of sedative premedication on patient experience after general anesthesia: a randomized clinical trial. JAMA. 2015;313:916–925.
- Tighe PJ, Le-Wendling LT, Patel A, Zou B, Fillingim RB. Clinically derived early postoperative pain trajectories differ by age, sex, and type of surgery. Pain. 2015;156:609–617.
- Gan TJ, Diemunsch P, Habib AS, et al; Society for Ambulatory Anesthesia. Consensus guidelines for the management of postoperative nausea and vomiting. AnesthAnalg. 2014;118:85–113.

- Habib NE, Mandour NM, Balmer HG. Effect of midazolam on anxiety level and pain perception in cataract surgery with topical anesthesia. J Cataract Refract Surg. 2004;30:437–443.
- Klopfenstein CE, Forster A, Van Gessel E. Anesthetic assessment in an outpatient consultation clinic reduces preoperative anxiety. Can J Anaesth. 2000;47:511–515.
- Kamau A, Mung'ayi V, Yonga G. The effect of a preanaesthesia clinic consultation on adult patient anxiety at a tertiary hospital in Kenya: a cohort study. Afr Health Sci. 2017;17:138–146.
- Raghavan G, Shyam V, Murdoch JAC. A survey of anesthetic preference and preoperative anxiety in hip and knee arthroplasty patients: the utility of the outpatient preoperative anesthesia appointment. J Anesth. 2019;33:250–256.
- Powell R, Scott NW, Manyande A, et al. Psychological preparation and postoperative outcomes for adults undergoing surgery under general anaesthesia. Cochrane Database Syst Rev. 2016;2016:Cd008646.
- Berger M, Schenning KJ, Brown CH, et al; Perioperative Neurotoxicity Working Group. Best practices for postoperative brain health: recommendations from the fifth international perioperative neurotoxicity working group. AnesthAnalg. 2018;127:1406–1413.
- Hughes CG, Boncyk CS, Culley DJ, et al; Perioperative Quality Initiative (POQI) 6 Workgroup. American Society for Enhanced Recovery and Perioperative Quality Initiative joint consensus statement on postoperative delirium prevention. AnesthAnalg. 2020;130:1572–1590.
- Inouye SK, Robinson T, Blaum C, et al. Postoperative delirium in older adults: best practice statement from the American Geriatrics Society. J Am Coll Surg. 2015;220:136–148e1.
- Lei VJ, Navathe AS, Seki SM, Neuman MD. Perioperative benzodiazepine administration among older surgical patients. Br J Anaesth. 2021;127:e69–e71.
- Burfeind KG, Zarnegarnia Y, Tekkali P, O'Glasser AY, Quinn JF, Schenning KJ. Potentially inappropriate medication administration is associated with adverse postoperative outcomes in older surgical patients: a retrospective cohort study. AnesthAnalg. 2022;135:1048– 1056.
- Fong TG, Inouye SK. The inter-relationship between delirium and dementia: the importance of delirium prevention. Nat Rev Neurol. 2022;18:579–596.
- Fong TG, Davis D, Growdon ME, Albuquerque A, Inouye SK. The interface between delirium and dementia in elderly adults. Lancet Neurol. 2015;14:823–832.
- 21. Inouye SK, Marcantonio ER, Kosar CM, et al. The shortterm and long-term relationship between delirium and cognitive trajectory in older surgical patients. Alzheimers Dement. 2016;12:766–775.
- Ahmed KJ, Pilling JD, Ahmed K, Buchan J. Effect of a patientinformation video on the preoperative anxiety levels of cataract surgery patients. J Cataract Refract Surg. 2019;45:475–479.
- Rajput SK, Tiwari T, Chaudhary AK. Effect of preoperative multimedia based video information on perioperative anxiety and hemodynamic stability in patients undergoing surgery under spinal anesthesia. J Family Med Prim Care. 2021;10:237–242.

- Lim S, Oh Y, Cho K, Kim MH, Moon S, Ki S. The question of preoperative anxiety and depression in older patients and family protectors. Anesth Pain Med (Seoul). 2020;15:217–225.
- Tola YO, Chow KM, Liang W. Effects of non-pharmacological interventions on preoperative anxiety and postoperative pain in patients undergoing breast cancer surgery: a systematic review. J Clin Nurs. 2021;30:3369–3384.
- 26. Ertuğ N, Ulusoylu O, Bal A, Özgür H. Comparison of the effectiveness of two different interventions to reduce preoperative anxiety: a randomized controlled study. Nurs Health Sci. 2017;19:250–256.
- Bang YJ, Lee JH, Kim CS, Lee YY, Min JJ. Anxiolytic effects of chewing gum during preoperative fasting and patient-centered outcome in female patients undergoing elective gynecologic surgery: randomized controlled study. Sci Rep. 2022;12:4165.
- Jakobsen CJ, Blom L, Brondbjerg M, Lenler-Petersen P. Effect of metoprolol and diazepam on pre-operative anxiety. Anaesthesia. 1990;45:40–43.
- 29. Dyck JB, Chung F. A comparison of propranolol and diazepam for preoperative anxiolysis. Can J Anaesth. 1991;38:704–709.
- Wu LP, Kang WQ. Effect of dexmedetomidine for sedation and cognitive function in patients with preoperative anxiety undergoing carotid artery stenting. J Int Med Res. 2020;48:300060520938959.
- Salzmann S, Rienmüller S, Kampmann S, Euteneuer F, Rüsch D. Preoperative anxiety and its association with patients' desire for support—an observational study in adults. BMC Anesthesiol. 2021;21:149.
- Eberhart L, Aust H, Schuster M, et al. Preoperative anxiety in adults—a cross-sectional study on specific fears and risk factors. BMC Psychiatry. 2020;20:140.
- Wada S, Inoguchi H, Sadahiro R, et al. Preoperative anxiety as a predictor of delirium in cancer patients: a prospective observational cohort study. World J Surg. 2019;43:134–142.
- Kain ZN, Sevarino F, Alexander GM, Pincus S, Mayes LC. Preoperative anxiety and postoperative pain in women undergoing hysterectomy. A repeated-measures design. J Psychosom Res. 2000;49:417–422.
- 35. Kain ZN, Sevarino FB, Rinder C, et al. Preoperative anxiolysis and postoperative recovery in women undergoing abdominal hysterectomy. Anesthesiology. 2001;94:415–422.
- 36. Doleman B, Leonardi-Bee J, Heinink TP, Bhattacharjee D, Lund JN, Williams JP. Pre-emptive and preventive opioids for postoperative pain in adults undergoing all types of surgery. Cochrane Database Syst Rev. 2018;12:CD012624.
- Sobol-Kwapinska M, Bąbel P, Plotek W, Stelcer B. Psychological correlates of acute postsurgical pain: a systematic review and metaanalysis. Eur J Pain. 2016;20:1573–1586.
- Williams JB, Alexander KP, Morin JF, et al. Preoperative anxiety as a predictor of mortality and major morbidity in patients aged >70 years undergoing cardiac surgery. Am J Cardiol. 2013;111:137–142.
- Pinto PR, McIntyre T, Almeida A, Araújo-Soares V. The mediating role of pain catastrophizing in the relationship between presurgical anxiety and acute postsurgical pain after hysterectomy. Pain.

2012;153:218-226.

- Euteneuer F, Kampmann S, Rienmüller S, Salzmann S, Rüsch D. Patients' desires for anxiolytic premedication - an observational study in adults undergoing elective surgery. BMC Psychiatry. 2022;22:193.
- Richardson SJ, Davis DHJ, Stephan BCM, et al. Recurrent delirium over 12 months predicts dementia: results of the Delirium and Cognitive Impact in Dementia (DECIDE) study. Age Ageing. 2021;50:914–920.
- 42. Wang ML, Min J, Sands LP, Leung JM, Group PMR. Midazolam premedication immediately before surgery is not associated with early postoperative delirium. AnesthAnalg. 2021;133:765–771.
- Adults AGSEPoPDiO. American Geriatrics Society abstracted clinical practice guideline for postoperative delirium in older adults. J Am Geriatr Soc. 2015;63:142–150.
- Aldecoa C, Bettelli G, Bilotta F, et al. European Society of Anaesthesiology evidence-based and consensus-based guideline on postoperative delirium. Eur J Anaesthesiol. 2017;34:192–214.
- 45. Panel BtAGSBCUE. American Geriatrics Society 2019 updated AGS Beers Criteria for potentially inappropriate medication use in older adults. J Am Geriatr Soc. 2019;67:674–694.
- Panel AGSBCUE. American Geriatrics Society updated Beers Criteria for potentially inappropriate medication use in older adults. J Am Geriatr Soc. 2012;60:616–631.
- Spence J, Belley-Côté E, Jacobsohn E, et al. Restricted versus liberal intraoperative benzodiazepine use in cardiac anaesthesia for reducing delirium (B-Free Pilot): a pilot, multicentre, randomised, cluster crossover trial. Br J Anaesth. 2020;125:38–46.
- Kowark A, Rossaint R, Keszei AP, et al; I-PROMOTE study group. Impact of PReOperative Midazolam on OuTcome of Elderly patients (I-PROMOTE): study protocol for a multicentre randomised controlled trial. Trials. 2019;20:430.
- Heinrich M, Nottbrock A, Borchers F, et al; BioCog Consortium. Preoperative medication use and development of postoperative delirium and cognitive dysfunction. Clin Transl Sci. 2021;14:1830– 1840.
- Evered L, Silbert B, Knopman DS, et al; Nomenclature Consensus Working Group. Recommendations for the nomenclature of cognitive change associated with anaesthesia and surgery-2018. AnesthAnalg. 2018;127:1189–1195.
- Li WX, Luo RY, Chen C, et al. Effects of propofol, dexmedetomidine, and midazolam on postoperative cognitive dysfunction in elderly patients: a randomized controlled preliminary trial. Chin Med J (Engl). 2019;132:437–445.
- 52. Mansouri N, Nasrollahi K, Shetabi H. Prevention of cognitive dysfunction after cataract surgery with intravenous administration of midazolam and dexmedetomidine in elderly patients undergoing cataract surgery. Adv Biomed Res. 2019;8:6.
- 53. Mahanna-Gabrielli E, Schenning KJ, Eriksson LI, et al. Stateof the clinical science of perioperative brain health: report from the American Society of Anesthesiologists brain healthinitiative summit 2018. Br J Anaesth. 2019;123:464–478.
- 54. Eckenhoff RG, Maze M, Xie Z, et al. Perioperative neurocognitive

disorder: state of the preclinical science. Anesthesiology. 2020;132:55-68.

- 55. Subramaniyan S, Terrando N. Neuroinflammation and perioperative neurocognitive disorders. AnesthAnalg. 2019;128:781–788.
- Safavynia SA, Goldstein PA. The role of neuroinflammation in postoperative cognitive dysfunction: moving from hypothesis to treatment. Front Psychiatry. 2018;9:752.
- Madsen BK, Zetner D, Møller AM, Rosenberg J. Melatonin for preoperative and postoperative anxiety in adults. Cochrane Database Syst Rev. 2020;12:CD009861.
- Johansen ME. Gabapentinoid use in the United States 2002 through 2015. JAMA Intern Med. 2018;178:292–294.
- Hong JSW, Atkinson LZ, Al-Juffali N, et al. Gabapentin and pregabalin in bipolar disorder, anxiety states, and insomnia: systematic review, meta-analysis, and rationale. Mol Psychiatry. 2022;27:1339–1349.
- Park CM, Inouye SK, Marcantonio ER, et al. Perioperative gabapentin use and in-hospital adverse clinical events among older adults after major surgery. JAMA Intern Med. 2022;182:1117–1127.
- Bongiovanni T, Anderson TS, Marcum ZA. Perioperative gabapentin use in older adults: revisiting multimodal pain management. JAMA Intern Med. 2022;182:1127–1128.
- Blessberger H, Lewis SR, Pritchard MW, et al. Perioperative betablockers for preventing surgery-related mortality and morbidity in adults undergoing cardiac surgery. Cochrane Database Syst Rev. 2019;9:CD013435.
- Blessberger H, Lewis SR, Pritchard MW, et al. Perioperative betablockers for preventing surgery-related mortality and morbidity in adults undergoing non-cardiac surgery. Cochrane Database Syst Rev. 2019;9:CD013438.
- Duncan D, Sankar A, Beattie WS, Wijeysundera DN. Alpha-2 adrenergic agonists for the prevention of cardiac complications among adults undergoing surgery. Cochrane Database Syst Rev. 2018;3:CD004126.
- 65. Weibel S, Jelting Y, Pace NL, et al. Continuous intravenous perioperative lidocaine infusion for postoperative pain and recovery in adults. Cochrane Database Syst Rev. 2018;6:CD009642.
- Graff V, Cai L, Badiola I, Elkassabany NM. Music versus midazolam during preoperative nerve block placements: a prospective randomized controlled study. Reg Anesth Pain Med. 2019;44:796– 799.
- Ganry L, Hersant B, Sidahmed-Mezi M, Dhonneur G, Meningaud JP. Using virtual reality to control preoperative anxiety in ambulatory surgery patients: a pilot study in maxillofacial and plastic surgery. J Stomatol Oral Maxillofac Surg. 2018;119:257–261.
- Sriramka B, Mallik D, Singh J, Khetan M. Effect of hand-holding and conversation alone or with midazolam premedication on preoperative anxiety in adult patients—a randomised controlled trial. Indian J Anaesth. 2021;65:128–132.
- Ciccozzi A, Marinangeli F, Colangeli A, et al. Anxiolysis and postoperative pain in patients undergoing spinal anesthesia for abdominal hysterectomy. Minerva Anestesiol. 2007;73:387–393.

- Kowark A, Berger M, Rossaint R, Schmid M, Coburn M, group P-S. Association between benzodiazepine premedication and 30day mortality rate: a propensity-score weighted analysis of the Peri-interventional Outcome Study in the Elderly (POSE). Eur J Anaesthesiol. 2022;39:210–218.
- 71. Walker KJ, Smith AF. Premedication for anxiety in adult day surgery. Cochrane Database Syst Rev. 2009;2009:CD002192.
- 72. Kim H, Park SS, Shim J. Differences in midazolam premedication effects on recovery after short-duration ambulatory anesthesia with propofol or sevoflurane for gynaecologic surgery in young patients: a randomized controlled trial. Medicine (Baltim). 2020;99:e23194.