



Multicenter Review of Practice Patterns Regarding Benzodiazepine Use in Cardiac Surgery

**Allison M Janda, MD
October 8, 2021**

Disclosures

Allison M Janda, M.D.

**Clinical Lecturer in Anesthesiology
University of Michigan, Ann Arbor, Michigan, USA**

- I have financial relationship(s) with:
 - **Grant / Research (current):**
 - *PCORI – Co-investigator, “Trajectories of Recovery after Intravenous Propofol vs. Inhaled Volatile Anesthesia (THRIVE)”*
 - **Grant / Research (current):**
 - *NIH R01 – Co-investigator, 1R01LM01389401 “A scalable service to improve health care quality through precision audit and feedback”*
 - **Grant / Research (past):**
 - *NIH T32 Research Fellowship Grant – 5T32GM103730-07 (past)*
 - **Grant / Research (past):**
 - *Becton Dickinson and Company (past)*
- My presentation does not include discussion of off-label or investigational use drugs or devices.

Introduction

There is little clarity on the impact of intraoperative benzodiazepines

- Benzodiazepine-sparing techniques are used due to the evidence in the ICU population relating benzodiazepine use to delirium¹⁻¹³
- Others routinely administer benzodiazepines for their amnestic properties for concern of intraoperative awareness in the higher risk cardiac surgery population^{1,4,14-16}

Practice patterns are dogmatic and debated

Clinical trials are underway studying outcomes related to benzodiazepine use,⁴ but little is known about benzodiazepine use patterns beyond survey data¹

Introduction

Aims

- Describe benzodiazepine use during cardiac surgery across MPOG centers
- Identify patient factors associated with benzodiazepine use
- Describe the prevalence and variation of benzodiazepine use for cardiac surgeries across patients, providers, and institutions
- Provide context for the future findings of clinical trials studying outcomes related to benzodiazepine administration

Hypothesis

- Patient, provider, and institutional factors were independently associated with benzodiazepine use during cardiac surgery, and patient factors are **not** the primary driver of variation in use

Methods

Study Cohort

- Adults who underwent elective or urgent cardiac surgical procedures from January 1, 2014 until August 1, 2019 at MPOG institutions

Exclusions

- ASA 5 or 6
- Lung transplants, heart transplants, mechanical circulatory support, aortic procedures, procedures requiring circulatory arrest or transcatheter procedures
- Case duration <120 minutes

Methods

Primary outcome: Exposure to benzodiazepines

- Defined as a bolus or infusion of midazolam, alprazolam, diazepam, clonazepam or lorazepam given within two hours of anesthesia start and anesthesia end

Methods

Covariates

- Patient level
 - Demographic data
 - Surgery type and duration, whether bypass was used
 - Comorbidities
 - Year of surgery
- Provider level
 - Case volume/year
- Institutional level
 - University affiliation
 - Case volume/year

Methods

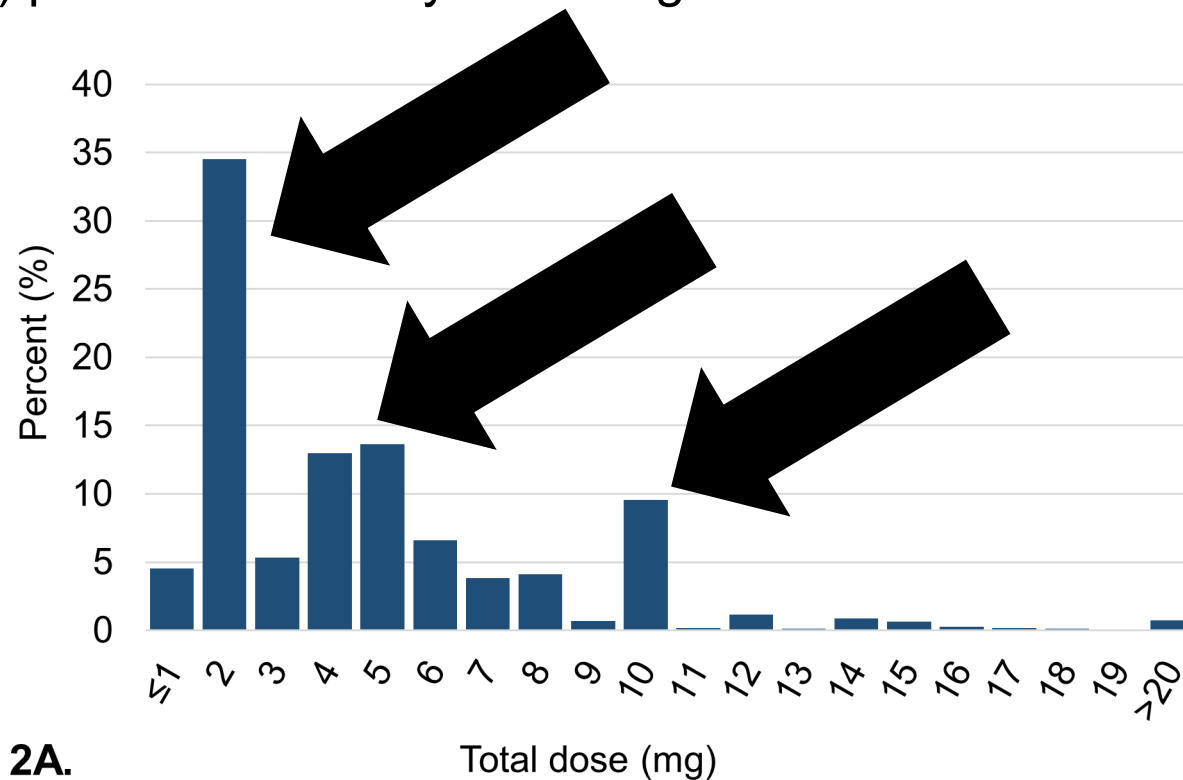
Statistical Analyses

- Mixed effects and multilevel models to assess random versus fixed effects of variation in benzodiazepine use
- Assessment of variance using variance estimates and median odds ratios (MORs)
- Goal is to inform us on the proportion of total variance in the outcome that is attributable to that factor

Results

65,508 patients across 33 MPOG institutions were included

- 58,004 (88.5%) patients received a benzodiazepine
- Median midazolam-equivalent dose of 4.0mg (IQR 2.0-6.0mg)
- 29,824 (84.4%) patients over 65 years of age received benzodiazepines



2A.

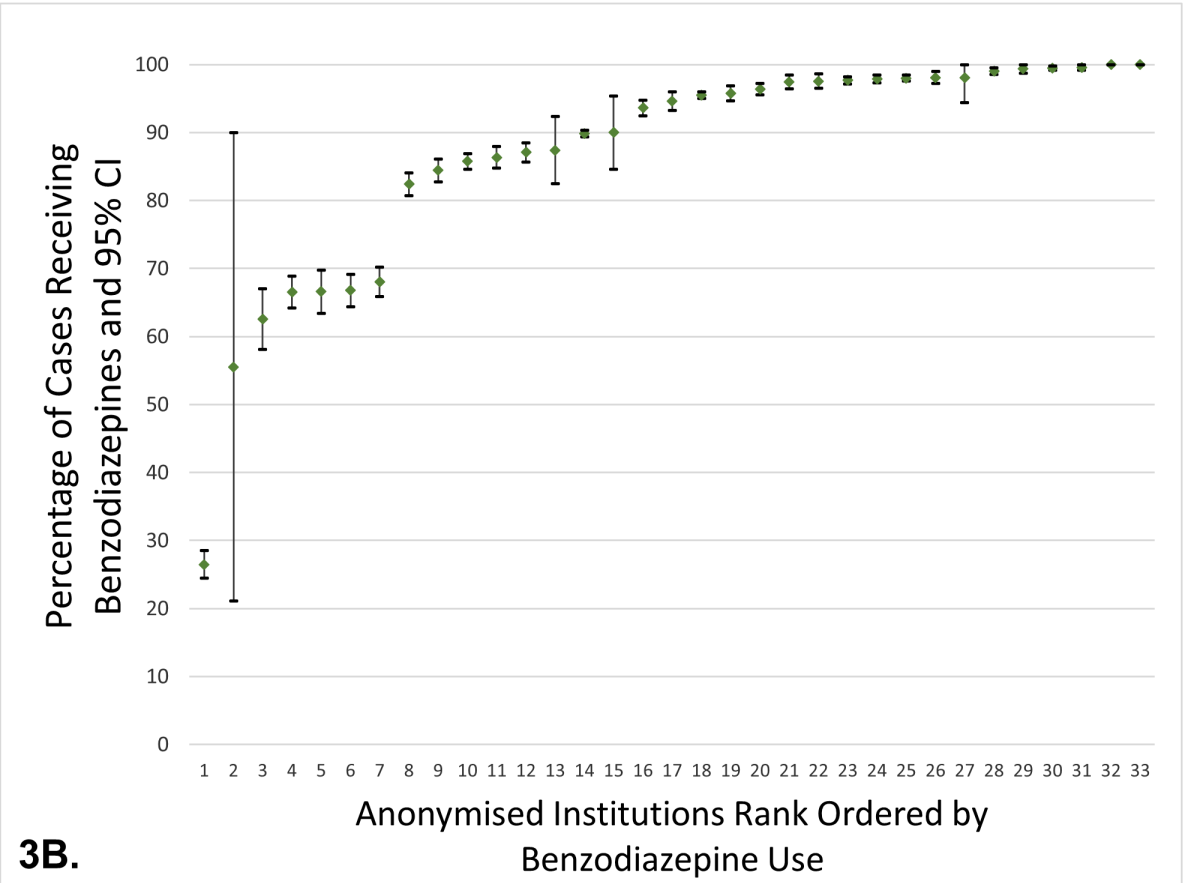
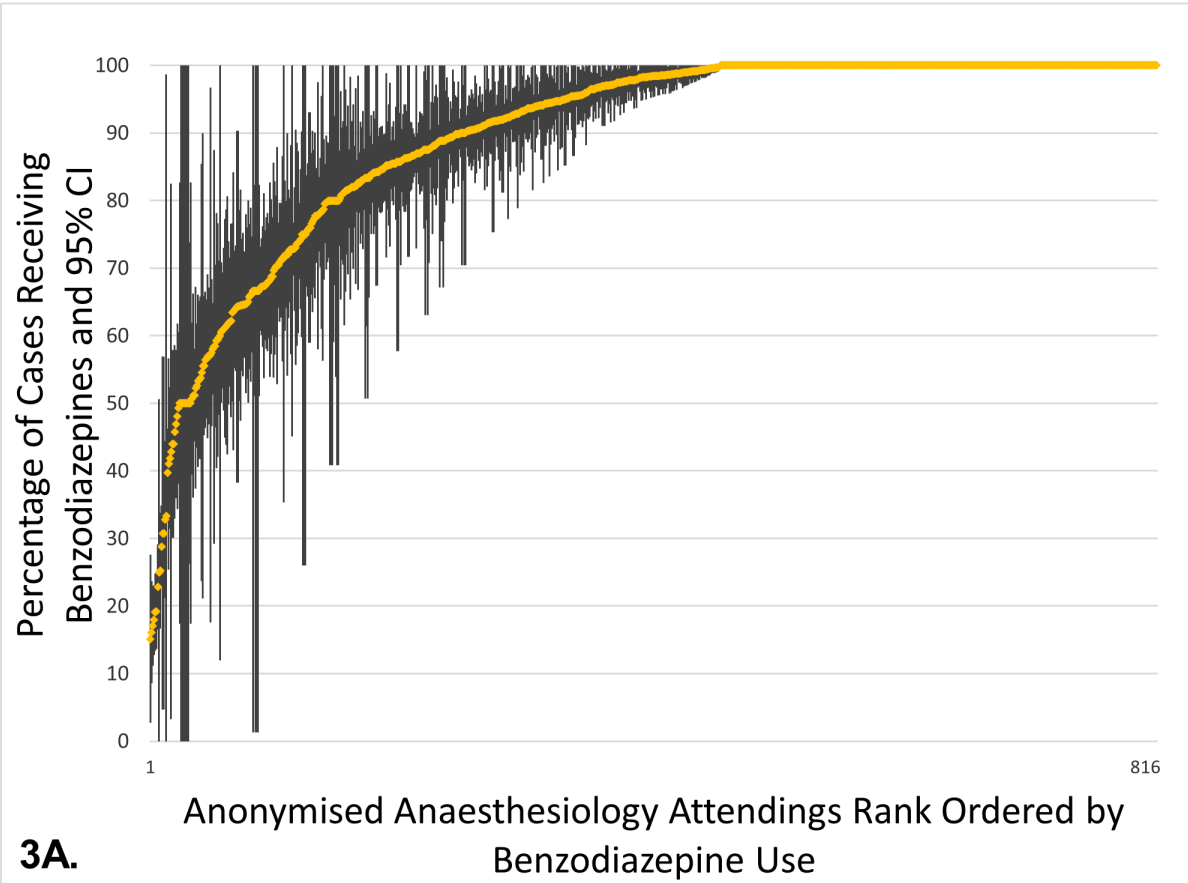
Results

Preliminary *unadjusted* generalized linear mixed models showed:

- 30.5% of the variation was explained by patient factors
- 14.7% by the primary provider
- 54.7% by the institution

Over **two-thirds** of the variation was explained by provider and institution

Results



Results

Adjusted multilevel models showed that factors strongly associated with a ***decreased*** likelihood of benzodiazepine administration included:

- Older age, >80 years versus ≤ 50 years, **aOR 0.04**, 95% CI 0.04-0.05, $p < 0.0001$
- Recent year of surgery, 2019 versus 2014, **aOR 0.42**, 95% CI 0.37-0.49, $p < 0.0001$
- University-affiliation, **aOR 0.08**, 95% CI 0.02-0.35, $p = 0.0007$
- Low provider case-volume, **aOR 0.44**, 95% CI 0.25-0.75, $p = 0.002$

Adjusted multilevel models showed that factors strongly associated with an ***increased*** likelihood of benzodiazepine administration included:

- History of drug abuse, **aOR 1.29**, 95% CI 1.02-1.65, $p = 0.04$
- Use of cardiopulmonary bypass, **aOR 2.26**, 95% CI 1.99-2.55, $p < 0.0001$

Results

Primary Analysis:

- Adjusted MORs for receiving a benzodiazepine:
 - **2.7** between randomly selected **providers**
 - **4.2** between randomly selected **institutions**

Low vs. High Dose Secondary Analysis:

- Adjusted MORs for receiving $<0.05\text{mg kg}^{-1}$ vs. $\geq 0.05\text{mg kg}^{-1}$
 - **3.1** between randomly selected **providers**
 - **6.9** between randomly selected **institutions**

Results

Increased Risk of Hemodynamic Instability Cohort *A Priori* Sensitivity Analysis:

- Adjusted MORs for receiving a benzodiazepine:
 - **2.1** between randomly selected **providers**
 - **3.5** between randomly selected **institutions**

Starting Provider *Post-hoc* Sensitivity Analysis:

- Adjusted MORs for receiving a benzodiazepine:
 - **2.9** between randomly selected **providers**
 - **4.3** between randomly selected **institutions**

Discussion

Conclusions

- **Institution and provider** as opposed to patient factors accounted for over **two-thirds** of the variation of benzodiazepine administration

Limitations

- MPOG does not include ALL institutions providing cardiac surgical care
- Most institutions (27/33) were university-affiliated

Future Directions

- These data may serve as a model for understanding cardiac anesthesiology practice variation provide context for the findings of randomized trials evaluating intraoperative benzodiazepine administration and outcomes

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Thank You

Huge thanks to my co-authors:

- Jessica Spence, MD PhD
- Timur Dubovoy, MD
- Emilie Belley-Côté, MD PhD
- Graciela Mentz, PhD
- Sachin Kheterpal, MD MBA
- Michael R. Mathis, MD

Questions?