

SYSTEMATIC REVIEW AND META-ANALYSES OF TRENDELENBURG AND
PRONE POSITION ON INTRAOCULAR PRESSURE IN ADULT PATIENTS
UNDERGOING SURGERY

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by
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*“Every good and perfect gift is from above,
coming down from the Father of the heavenly lights,
who does not change like shifting shadows.”*

(James 1:17 New International Version).

This dissertation is dedicated to my husband, James Hudson Van Wicklin, who is one of God's good and perfect gifts to me. My husband has been my constant support and encouragement as I have pursued this doctoral degree and throughout our 45 years of married life.

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LIST OF ABBREVIATIONS

ABBREVIATION	EXPLANATION
AORN	Association of periOperative Registered Nurses
ASA.....	American Society of Anesthesiologists
BMI.....	body mass index
CI.....	confidence interval
CNOR	Certified Nurse Operating Room
CPSN-R.....	Certified Plastic Surgical Nurse-Retired
CRNFA(E).....	Certified Registered Nurse First Assistant (Emeritus)
<i>d</i>	standard mean difference
EBL.....	estimated blood loss
F	fixed effect model
FAAN.....	Fellow of the American Academy of Nursing
<i>I</i> ²	I-squared
IOP	intraocular pressure
ISPAN-F	International Society of Plastic and Aesthetic Nurses-Fellow
<i>k</i>	comparisons
kg/m ²	kilograms per meter-squared
<i>LL</i>	lower limit
min	minutes
mL.....	milliliters
mmHg	millimeters of mercury
PI.....	principal investigator
PLNC	Professional Legal Nurse Consultant
Q.....	Cochrane's Q

R.....	random effects model
RN.....	registered nurse
<i>s</i>	studies
<i>SD</i>	standard deviation
SMD.....	standard mean difference
Std.....	standard
T.....	time points
<i>UL</i>	upper limit

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ABSTRACT

Background. Patients undergoing surgery in the Trendelenburg and prone positions may be at risk for postoperative vision loss associated with increased intraocular pressure. The purpose of this dissertation research is to estimate the magnitude of the increase in intraocular pressure at specific perioperative time points in adult patients undergoing surgery in the Trendelenburg and prone positions.

Methods. Comprehensive search strategies were used to identify eligible studies for two meta-analyses and to address the research questions. For each meta-analysis, standardized mean difference effect sizes were calculated for selected perioperative time points.

Results. Using a random effects model, the meta-analysis examining the effect of Trendelenburg position, showed that intraocular pressure decreased significantly after induction and before arousal. Intraocular pressure increased significantly after abdominal insufflation and during Trendelenburg position. The meta-analysis examining the effect of prone position, showed that intraocular pressure increased significantly between induction of anesthesia and up to 10 minutes of prone position and continued to increase significantly until the end of the prone position.

Conclusions. Intraocular pressure increases of the magnitude found in this research demonstrate the need for implementing interventions to reduce the risk for postoperative vision loss in patients undergoing surgery in the Trendelenburg and prone positions.

1. INTRODUCTION

This research project involved conducting two meta-analyses to estimate the overall magnitude and effect of the Trendelenburg and prone positions on intraocular pressure (IOP) in adult patients undergoing surgery. Patients 18 years or older were considered to be adult patients. The increase in IOP that occurs when the Trendelenburg and prone positions are used can potentially lead to postoperative vision loss and other ocular complications.

In the Trendelenburg position, the patient's feet are higher than the patient's head by 15 degrees to 30 degrees (MacDonald & Washington, 2012). Many surgeons use a steep Trendelenburg position of 30 degrees to 45 degrees, particularly during laparoscopic or robotic surgery, which is frequently used for prostatectomy, hysterectomy, colorectal surgery, and many other procedures. While providing enhanced visualization of the operative field, the Trendelenburg position also decreases venous return from the head, leading to venous pooling and increased IOP (Akhavan, Gainsburg, & Stock, 2010; Astuto, Minardi, Uva, & Gullo, 2011; Awad et al., 2009; Borahay et al., 2013; Cullen & Ferguson, 2012; Ghomi, 2012; Kan, Brown, & Gainsburg, 2015; Mondzelewski et al., 2015; Taketani et al., 2015). Increased IOP resulting from the use of the Trendelenburg position poses a risk for postoperative glaucoma, detached retina, or partial to complete vision loss (Astuto et al., 2011; Borahay et al., 2013; Emery et al., 2015; Ghomi, Kramer, Askari, Chavan, & Einarsson, 2012; Gkegkes, Karydis, Tyritzis, & Iavazzo, 2015; Gould, Cull, Wu, & Osmundsen, 2012; Grosso et al., 2013; Hoskikawa et al., 2014; Lee & Newman, 2018; Taketani et al., 2015).

Partial to complete postoperative vision loss is also a complication of surgery when the patient is positioned face-down on their abdomen in the prone position. This complication occurs with greater frequency after spine, head and neck, cardiac, and some orthopedic procedures (Emery et al., 2015; Lee & Newman, 2018). There have been reports of postoperative vision loss (Goni et al., 2012; Quraishi, Wolinsky, & Gokaslan, 2012; Reddy, Foroozan, Edmond, & Hinckley, 2008; Shifa, Abebe, Bekele, & Habte, 2016; Stang-Veldhouse, Yeu, Rothenberg, & Mizen, 2010) as well as subconjunctival hemorrhage (Akhaddar & Boucetta, 2012), subperiosteal orbital hemorrhage (Russell & Dutton, 2011), and Horner syndrome (Guillaume & Gowreesunker, 2013) after prolonged surgery in the prone position. When the anesthetized patient is in the prone position, IOP increases and the extent of this increase is related to the amount of time the patient remains in the prone position (Agah, Ghasemi, Roodneshin, Radpay, & Moradian, 2011; Eddama, 2013; Kamel & Barnette, 2014; Kendrick, 2012; Pinkney et al., 2012; Szmuk et al., 2013; Yoshimura, Hayashi, Tanaka, Nomura, & Kawaguchi, 2015). After only a few minutes in the prone position, IOP can increase significantly (Nuzzi & Tridico, 2015).

Researchers have independently investigated the quantitative increases of IOP that occur intraoperatively in adult surgical patients in the Trendelenburg (Astuto et al., 2011; Awad et al., 2009; Borahay et al., 2013; Grosso et al., 2013; Hoshikawa et al., 2014; Molloy, 2011; Mondzelewski et al., 2015; Taketani et al., 2015) and prone (Emery et al., 2015; Lee & Newman, 2018; Nuri Deniz et al., 2013; Quraishi et al., 2012; Shifa et al., 2016; Stang-Veldhouse et al., 2010; Yoshimura et al., 2015) positions. However, there is a need for systematic review and meta-analyses of these studies to demonstrate the magnitude of the increase in IOP and the overall effect sizes. Understanding the

degree to which IOP can increase intraoperatively can provide high-quality evidence supporting the need for implementing perioperative interventions to mitigate the increase in IOP and reduce the risk for postoperative vision loss in adult patients undergoing surgery in the Trendelenburg and prone positions. Meta-analysis research methods provide increased power compared to individual studies, improve estimates of effect size, and help resolve uncertainty when the results of individual studies disagree (Berlin, 1995). Meta-analysis research provides a high level of objectivity, precision, and generalizability because all of the evidence pertaining to a particular phenomenon is included in the analysis (Thompson, 1994). Patient outcomes can be optimized by reducing the risk for postoperative vision loss and other ocular complications resulting from increased IOP in adult surgical patients. Currently, there has been no quantitative meta-analytic synthesis of the existing studies examining the increase in IOP in surgical patients positioned in the Trendelenburg or prone positions.

The purpose of these systematic reviews and meta-analyses is to estimate the magnitude of the increase in IOP at relevant perioperative time points in adult patients undergoing any type of surgery in the Trendelenburg and prone position. The following chapter presents the research proposal for the systematic reviews and meta-analyses with a description of the approach used, an explanation of the significance and innovation of the project, and a discussion of the anticipated challenges. Chapter 3 is the manuscript to be submitted for publication titled, “Systematic Review and Meta-analysis of Trendelenburg Position on Intraocular Pressure in Adult Patients Undergoing Surgery” and Chapter 4 is the manuscript to be submitted for publication titled, “Systematic Review and Meta-analysis of Prone Position on Intraocular Pressure in Adult Patients

Undergoing Surgery.” Chapter 5 provides a summary of the results of the systematic reviews and meta-analyses as well as evidence-based recommendations for mitigating or reducing IOP when patients undergo surgery in the prone and Trendelenburg positions. These interventions may improve patient outcomes by preventing postoperative vision loss and other ocular complications.

2. RESEARCH PROPOSAL

This research project will involve conducting two systematic reviews and meta-analyses to estimate the overall magnitude and effect of the Trendelenburg and prone positions on intraocular pressure (IOP) in adult patients undergoing surgery. Patients 18 years or older are considered to be adult patients.

Significance

Understanding and mitigating the magnitude of the intraoperative increase in IOP can improve patient outcomes in adult patients undergoing surgery in the Trendelenburg and prone positions by reducing the risk for postoperative vision loss and other ocular complications.

Trendelenburg Position

In the Trendelenburg position, the patient's feet are higher than the patient's head by 15 degrees to 30 degrees (MacDonald & Washington, 2012). Many surgeons use a steep Trendelenburg position of 30 degrees to 45 degrees, particularly during laparoscopic and robotic surgery, which is frequently used for prostatectomy, hysterectomy, colorectal surgery, and various other procedures. This position moves the abdominal viscera cephalad to improve surgical access to the pelvic and abdominal organs. While providing enhanced visualization of the operative field, the Trendelenburg position also decreases venous return from the head, leading to venous pooling and increased IOP (Akhavan, Gainsburg, & Stock, 2010; Astuto, Minardi, Uva, & Gullo, 2011; Awad et al., 2009; Borahay et al., 2013; Cullen & Ferguson, 2012; Ghomi, 2012; Kan, Brown, & Gainsburg, 2015; Mondzelewski et al., 2015; Taketani et al., 2015).

Increased IOP resulting from the use of the Trendelenburg position poses a risk for postoperative glaucoma, detached retina, or partial to complete vision loss (Astuto et al., 2011; Borahay et al., 2013; Emery et al., 2015; Ghomi, Kramer, Askari, Chavan, & Einarsson, 2012; Gkegkes, Karydis, Tyritzis, & Iavazzo, 2015; Gould, Cull, Wu, & Osmundsen, 2012; Grosso et al., 2013; Hoshikawa et al., 2014; Lee & Newman, 2018; Taketani et al., 2015). Postoperative vision loss can be caused by an ischemic process that occurs as a result of decreased blood supply from the arteries of the optic nerve or by venous stasis that occurs as a result of decreased venous outflow (Gkegkes et al., 2015; Kan et al., 2015; Molloy, 2011). Some researchers have found the increased IOP associated with the Trendelenburg position poses a risk for postoperative vision loss, particularly in patients with pre-existing ocular disease (Astuto et al., 2011; Borahay et al., 2013; Grosso et al., 2017; Hoshikawa et al., 2014; Mondzelewski et al., 2015; Taketani et al., 2015). Older patients who have elevated baseline IOP are at greater risk for ischemic optic neuropathy (Freshcoln & Diehl, 2014; Ghomi, 2012; Grosso et al., 2013; Kan et al., 2015; Mondzelewski et al., 2015). Prolonged use of the steep Trendelenburg position may also lead to retinal detachment (Hewer, 1956). The increase in IOP and risk for postoperative vision loss is directly correlated with the amount of time the patient is in the Trendelenburg position (Mizumoto, Gosho, Iwaki, Zako, 2017; Nuzzi & Tridico, 2016; Pinkney et al., (2012); Stang-Veldhouse, Yeu, Rothenberg, & Mizen, 2010; Yoo et al., 2014).

Prone Position

In the prone position, the patient is positioned face-down on their abdomen. This position provides surgical access to the dorsal aspects of the patient's body. Partial to

complete postoperative vision loss is a complication of surgery in the prone position, and it occurs with greater frequency after spine, head and neck, cardiac, and some orthopedic procedures (Emery et al., 2015; Lee & Newman, 2018). There have been reports of postoperative vision loss (Goni et al., 2012; Quraishi, Wolinsky, & Gokaslan, 2012; Reddy, Foroozan, Edmond, & Hinckley, 2008; Shifa, Abebe, Bekele, & Habte, 2016; Stang-Veldhouse et al., 2010), as well as subconjunctival hemorrhage (Akhaddar & Boucetta, 2012), subperiosteal orbital hemorrhage (Russell & Dutton, 2011), and Horner syndrome (Guillaume & Gowreesunker, 2013) after prolonged surgery in the prone position.

Elevated IOP levels decrease perfusion pressure of the optic nerve and increase the patient's risk for postoperative vision loss. According to Hayreh (2001), perfusion pressure of the optic nerve is the difference between pressures of the ciliary arteries in the nerve and the venous drainage of the eye. This difference is approximated by the level of IOP. The "higher the intraocular pressure, the lower the perfusion pressure, and consequently, the lower the blood flow in the optic nerve head" (p. 608). The lower the blood flow in the optic nerve head, the greater the risk for postoperative vision loss.

When the anesthetized patient is in the prone position, IOP increases and the extent of this increase is related to the amount of time the patient is in the prone position (Agah, Ghasemi, Roodneshin, Radpay, & Moradian, 2011; Eddama, 2013; Kamel & Barnette, 2014; Kendrick, 2012; Pinkney et al., 2012; Szmuk et al., 2013; Yoshimura, Hayashi, Tanaka, Nomura, & Kawaguchi, 2015). After only a few minutes in the prone position, IOP can increase significantly (Nuzzi & Tridico, 2015). The most dramatic increases in IOP occur when the patient is in the jack-knife or Kraske position (i.e., a

prone position where the patient's hips are elevated and the patient's head is down; Kendrick, 2012; Pinkney et al., 2012; Nuzzi & Tridico, 2015). The jack-knife position is used for hemorrhoidectomy or other rectal surgery.

Yoshimura et al., (2015) suggested that measuring IOPs after one hour of surgery in the prone position could provide an opportunity for implementing interventions to prevent additional increases in IOP. Eddama (2013) also suggested that regular measurement of IOPs during prolonged surgeries could provide an opportunity for implementing a change in the patient's position when critical thresholds of IOP are reached.

Incidence

The exact incidence of postoperative vision loss in adult surgical patients is unknown because the data come largely from retrospective studies and case reports (Lee & Newman, 2018; Patil, Lad, Lad, Ho, & Boakye, 2008). In a 10-year prevalence study of postoperative vision loss in the United States, Shen, Drum, and Roth (2009) reviewed data from more than 5.6 million patients included in the Nationwide Inpatient Sample from 1995 to 2005. The researchers found the incidence of postoperative vision loss for patients undergoing laminectomy without fusion was 0.86 per 10,000. For patients undergoing spinal fusion, the incidence was 3.09 per 10,000. According to the American Academy of Orthopedic Surgeons (2014), there were 332,159 spinal fusion surgeries in the United States in 2005. Using the figures provided by Shen et al., this would equate to a total of 102.6 cases of postoperative vision loss. In 2011, the number of spinal fusions increased by 40% to 465,070 (American Academy of Orthopedic Surgeons (2014)), equating to a total of 143.7 cases of postoperative vision loss.

Although postoperative vision loss may be considered a rare occurrence, as the number of surgeries performed in the Trendelenburg and prone positions increases, the number of cases of postoperative vision loss can be anticipated to increase proportionally. Fingar, Stocks, Weiss, and Steiner (2014) found that in 2012, the most frequently performed surgical procedures in the United States included laminectomy (149.1 per 100,000 population) and spinal fusion (143.6 per 100,000 population). These procedures are typically performed in the prone position. Frequently performed procedures in 2012 also included abdominal or vaginal hysterectomy (99.4 per 100,000 population), unilateral or bilateral oophorectomy (71.3 per 100,000 population), appendectomy (97.4 per 100,000 population), and colorectal resection (97.4 per 100,000 population; Fingar et al., 2014). These procedures are often performed in the Trendelenburg position using minimally invasive techniques (e.g., laparoscopic, robotic). The World Health Organization (Weiser et al., 2016) estimated a total global surgical volume of 312.9 operations performed in 2012, a 38.2% increase from the estimated 226.4 million surgical procedures performed in 2004. In 2016, there were 27.2 million surgical procedures performed in the United States (American Hospital Association, 2019). In light of these large and increasing numbers of surgical procedures, the prevention of postoperative vision loss takes on even greater importance.

Meta-analysis

Although some researchers have studied the quantitative intraoperative increases of IOP in surgical patients in the Trendelenburg (Astuto et al., 2011; Awad et al., 2009; Borahay et al., 2013; Grosso et al., 2013; Hoshikawa et al., 2014; Molloy, 2011; Mondzelewski et al., 2015; Taketani et al., 2015), and prone position (Emery et al., 2015;

Lee & Newman, 2018; Nuri Deniz et al., 2013; Quraishi et al., 2012; Shifa et al., 2016; Stang-Veldhouse et al., 2010; Yoshimura et al., 2015), there is a need for systematic review and meta-analysis of these studies to demonstrate the magnitude and overall effect size of the increase in IOP in adult surgical patients. Meta-analysis research can provide high-quality evidence to support the need for implementing intraoperative interventions designed to mitigate or reduce the increase of IOP and thus decrease the patient's risk for postoperative vision loss or other ocular complications.

In the Trendelenburg position, these interventions may include reducing the degree of Trendelenburg position (Ghomie et al., 2012; Mathew et al., 2018; Ozcan et al., 2017; Raz et al., 2015), monitoring IOP (Hoshikawa et al., 2013; Lee, Dallas, Daniel, & Cotter, 2016; Vitish-Sharma et al., 2018), and implementing periodic intraoperative position changes (Blecha et al., 2017; Borahay et al., 2013; Freshcoln & Diehl, 2014; Gkegkes et al., 2015; Gould et al., 2012; Mizrahi, Hugkulstone, Vyakarnam, & Parker, 2011; Molloy & Watson, 2012). In the prone position, these interventions include using a head-elevated position (Carey, Shaw, Weber, & DeVine, 2014; Emery et al., 2015; Fukui, Ahmad, McHugh, Tempelhoff, & Cheng, 2004; Fukui Tempelhoff, & Cheng, 2005), monitoring IOP (Eddama, 2013; Yoshimura et al., 2015), or providing periodic position changes (Molloy & Watson, 2012).

Currently, there has been no quantitative meta-analytic synthesis of the existing studies in these expanding areas of research. There is a need for systematic review and meta-analyses of these studies to demonstrate the magnitude and overall effect size of the intraoperative increase in IOP in adult patients. This research can provide high-quality evidence supporting the need for implementing interventions to mitigate the increase of

IOP and reduce the risk for postoperative vision loss and other ocular complications in adult patients undergoing surgery in the Trendelenburg and prone positions. Meta-analysis research methods provide increased power compared to individual studies, improve estimates of effect size, and help resolve uncertainty when the results of individual studies disagree (Berlin, 1995). Meta-analysis research provides a high level of objectivity, precision, and generalizability because all of the evidence pertaining to a particular phenomenon is included in the analysis (Thompson, 1994).

Innovation

This research proposal is innovative in two ways. The first innovative concept is using meta-analysis research to determine the magnitude and effect of two specific surgical positions on a physiological response that increases the potential for ocular injury in adult patients undergoing surgery in these positions. Implementing meta-analysis research to improve the care and outcomes of surgical patients related to specific surgical positions has not been done previously. This meta-analysis research can help to reduce the incidence of postoperative vision loss and other ocular complications by providing high-quality evidence to support guidelines and recommendations for safe patient positioning of adults undergoing all types of surgery in the Trendelenburg and prone positions.

Using the Perioperative Patient Focused Model (Rothrock & Smith, 2000) as a conceptual framework for each of the proposed meta-analyses is the second innovative approach. The application of this Model to the proposed research is innovative because although the Model is applicable to nursing interventions implemented by perioperative registered nurses (RNs), it has not been previously applied to perioperative nursing

research. The Model is relevant to nursing interventions executed to prevent patient injury when positioning adult surgical patients in the Trendelenburg and prone positions.

As shown in Figure 2.1, the Model is patient-focused (Rothrock & Smith, 2000). A primary role of the perioperative RN is to provide effective perioperative patient care and serve as the patient's advocate. Secondary to being patient focused, the Perioperative Patient Focused Model is outcome focused (Rothrock & Smith, 2000). The care provided by perioperative RNs is directed toward achieving high quality patient outcomes, such as preventing postoperative vision loss associated with increased IOP. Providing optimal perioperative patient care, serving as the patient's advocate, and achieving high quality outcomes requires specific perioperative knowledge and skillful implementation of best practices for patient positioning.

Consistent with the Perioperative Patient Focused Model (Rothrock & Smith, 2000), the evidence-based practices implemented by perioperative RNs during patient positioning promote patient safety, and optimal physiological and behavioral responses. Providing evidence about the magnitude of the intraoperative increase in IOP resulting from the Trendelenburg and prone positions will support the implementation of nursing interventions specifically directed at preventing postoperative vision loss and other ocular complications in adult surgical patients.

Approach

The principal investigator (PI), addressed the research questions related to the effect of Trendelenburg and prone position on IOP in adult patients undergoing surgery with a separate meta-analysis for each position (i.e., one meta-analysis examined the

magnitude of the effect of Trendelenburg position on IOP, and one meta-analysis examined the magnitude of the effect of prone position on IOP).

Inclusion Criteria

For each meta-analysis, the PI searched the literature for studies that addressed the research question and met inclusion criteria. Included studies were those

- written in English;
- reported between January 1, 1990 and September 30, 2018 (for the meta-analysis examining the effect of Trendelenburg position on IOP only);
- that encompassed dissertation research, conference abstracts, and presentations;
- that used either a one-group, pretest posttest comparison or a multiple-group, pretest posttest design;
- where the minimum age of study participants was 18-years or older;
- where the participants IOP was measured using any type of tonometer;
- where the participants received any type of general or spinal anesthesia; and
- with sufficient data to calculate an effect size.

The year 1990 was selected as the initial searching date for the meta-analysis examining the effect of Trendelenburg position on IOP because the first laparoscopic cholecystectomy was performed in 1987 (Vecchio, MacFayden, & Palazzo, 2000). The use of the steep Trendelenburg position has increased dramatically with the introduction and use of laparoscopic and robotic surgery. There was no initial date restriction for the meta-analysis examining the effect of prone position on IOP. When reports did not include sufficient data to calculate an effect size, the PI contacted the researchers on at least two separate occasions two to three weeks apart to obtain missing data.

Literature Search

For each meta-analysis, the PI conducted a comprehensive and exhaustive literature search to avoid bias due to a narrow or limited search. The PI implemented search strategies that included

- online searching of PubMed, CINAHL, Scopus, and Cochrane Database of Systematic Reviews databases for published and unpublished literature;
- ancestry searches of references from relevant reports;
- author searches of individuals identified in the literature as experts in the field;
- hand searches of relevant journals; and a
- dissertation search of the ProQuest database.

The PI collaborated with an expert health sciences reference librarian from the University of Missouri libraries to identify the most appropriate search terms, search dates, and databases and to refine search strategies.

The PI reviewed report abstracts for eligibility and obtained potentially eligible reports using library resources or by contacting the author. As the literature was searched and eligible reports were obtained, the PI used the *EndNote* bibliographic software (Clarivate Analytics, 2018) to compile a separate reference library for each meta-analysis.

Evidence Appraisal

Relevant research was independently evaluated and critically appraised according to the strength and quality of the evidence using the Association of periOperative Registered Nurses (AORN) Research Evidence Appraisal Tool – Study (See Appendices A-1 and A-2) by the PI and a second experienced evidence appraiser. The PI and

evidence appraiser participated in conference calls until 100 percent consensus was achieved on the level of study strength and quality for each of the studies included in each meta-analysis.

Coding

The PI developed a separate codebook for each meta-analysis and performed detailed coding of the studies for criteria relevant to address each research question and statistically analyze the effects and methodologies of each of the reports. To ensure accuracy of outcome data, a second trained researcher performed independent coding.

For each meta-analysis, the PI used the *EndNote* bibliographic software (Clarivate Analytics, 2018) to track eligibility and coding status for each meta-analysis. Eligible studies from the EndNote database were exported to an Excel file. The PI searched the Excel file for overlapping reports, and evaluated the identified reports to ensure they were not entered in the analysis more than once.

The PI used the codebook for each meta-analysis to translate key information from the relevant reports into the quantitative values necessary to implement meta-analysis to address the research questions (e.g., study characteristics, participant attributes, study design, effect size data). The PI and second coder independently entered coded data onto Excel spreadsheets. The PI and second coder participated in conference calls until 100 percent consensus was achieved on effect size data for each of the studies included in each meta-analysis. The PI screened the data to identify improbable values, compared questionable values to primary reports, and cleaned the data to ensure accuracy of data entry.

Data Analysis

Data for each meta-analysis were analyzed using *Comprehensive Meta-Analysis*, a statistical software developed specifically for meta-analysis research (Borenstein, Hedges, Higgins, & Rothstein, 2018). The PI obtained education and training to use this software during a course on meta-analysis research methods provided by the University of Missouri Sinclair School of Nursing. The PI further completed an educational and training course provided by the software developers.

For each meta-analysis, the PI used outcome data from eligible studies to calculate an overall mean effect size across reports. A random effects model was selected a priori to synthesize effect sizes. The overall mean effect size is the mean of the treatment group minus the mean of the control group divided by the pooled standard deviation. The effect size values of the individual studies were weighted to account for sample size and adjusted for bias. This process created a unitless, standardized measure of effect size across studies. To facilitate interpretation of effect size findings, for each meta-analysis, the effect size information was converted to the metric used to measure IOP (i.e., millimeters of mercury [mmHg]). If applicable, the PI conducted moderator analyses for different time points. The extent of publication bias for each meta-analysis was assessed by constructing a funnel plot. Notably, a funnel plot may suggest publication bias, but does not eliminate the bias (Sutton, 2009). An Egger's test using linear regression was also conducted to measure symmetry of the funnel plot (Borenstein et al., 2018; Egger, Smith, Schneider, & Minder, 1997; Sterne, Egger, & Moher, 2011). When the Egger's test was significant, Orwin's *Fail-safe N* test was conducted to estimate the number of missing studies required to overturn the conclusions of the meta-analysis (Orwin, 1983; Sutton, 2009). Orwin's *Fail-safe N* assesses the impact of

publication bias to determine whether the overall observed effect is robust (Orwin, 1983; Sutton, 2009).

For each meta-analysis, the PI tested for homogeneity of variance among effect sizes using Cochrane's Q, which estimates statistical significance; Tau-squared (T^2), which estimates the absolute value of the true variance between studies, but not the proportion of the variance; and I^2 , which estimates the proportion of true variance, but not the absolute value of the variance, (Borenstein, Hedges, Higgins, & Rothstein, 2009). Heterogeneity among the studies for each meta-analysis was anticipated due to the diversity of sample characteristics among studies.

Challenges

When conducting the meta-analysis research, the PI anticipated and prepared for challenges related to sampling, study quality, missing data, and coding. Achieving a successful and valid meta-analysis is dependent upon obtaining as many relevant studies as possible, and because of the nature of the research topics, the PI suspected this would be difficult. Locating and obtaining unpublished literature on the meta-analyses topics was also expected to be challenging; however, this is necessary to minimize publication bias. The PI managed these challenges by collaborating with an expert reference librarian to assist with search strategies and by completing exhaustive literature searches that included thorough ancestry, author, journal, and dissertation searches.

Meta-analysis is a synthesis of the findings of multiple study reports. For this reason, the validity of a meta-analysis of studies can be affected by including poor quality studies (i.e., studies lacking methodological rigor and internal validity). Along with a second reviewer, the PI assessed the level of strength and quality of each of the studies

included in each meta-analysis using specific criteria (e.g., sample size, generalizability) defined by the AORN Research Evidence Appraisal Tool – Study (Appendices A-1 and A-2).

Coding errors can lead to errors in or inability to analyze data. The PI managed this challenge by developing a codebook that allowed for studies to be coded with minimal coder reasoning or interpretation and by using a second coder to independently code effect size data to assure all studies were coded correctly and accurately. Eligible studies that failed to include sufficient data to calculate an effect size led to challenges in coding and limitations of data analysis. The PI made every attempt possible to obtain missing data by contacting study researchers or statisticians. When the PI was unable to obtain the missing data, the study was excluded from the analysis.

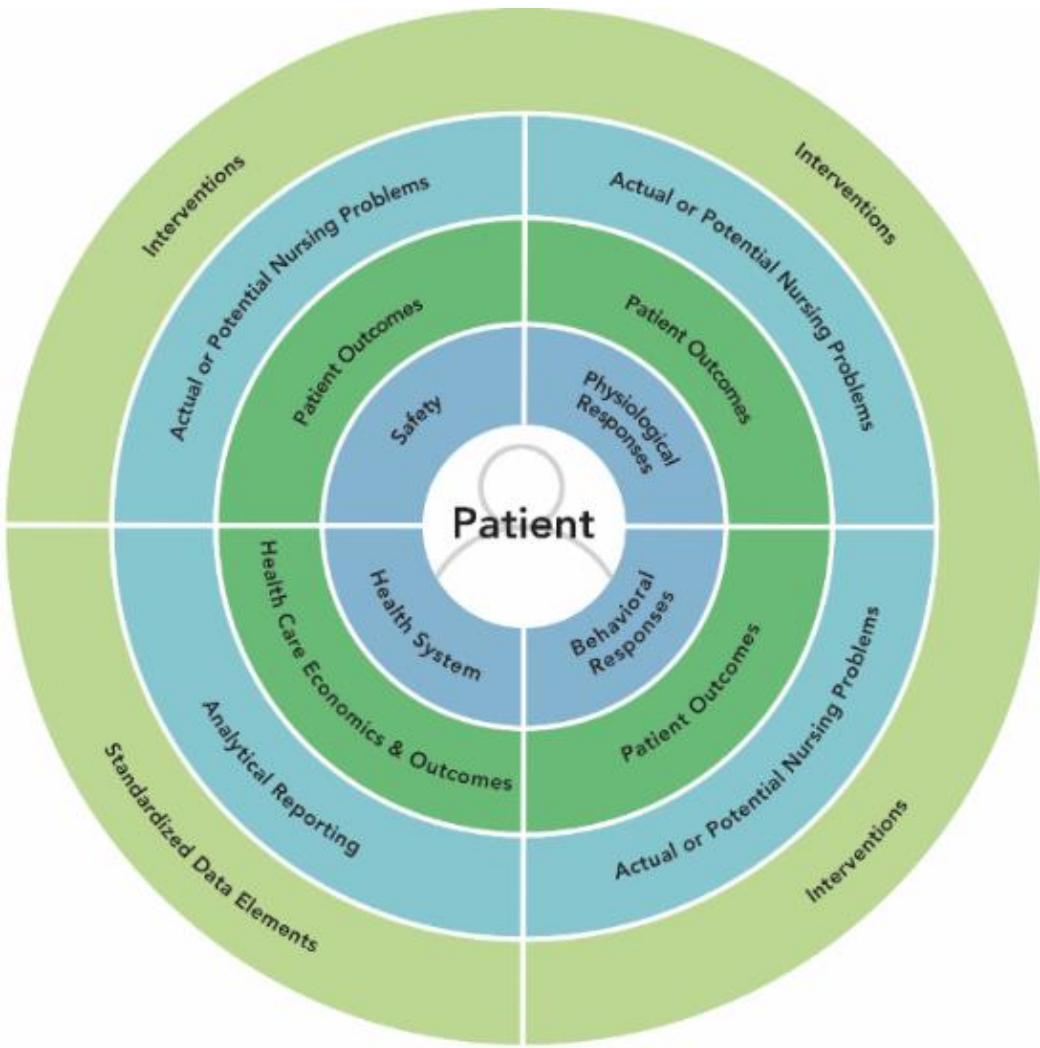


Figure 2.1. Perioperative Patient Focused Model (2017). Reprinted with permission from *Guidelines for Perioperative Practice*. Copyright © 2018, AORN, Inc, 2170 S. Parker Road, Suite 400, Denver, CO 80231. All rights reserved.

3. SYSTEMATIC REVIEW AND META-ANALYSIS OF TRENDENELBURG POSITION ON INTRAOCULAR PRESSURE IN ADULT PATIENTS UNDERGOING SURGERY

Van Wicklin, S. A. (2019). *Systematic review and meta-analysis of Trendelenburg position on intraocular pressure in adult patients undergoing surgery*. Manuscript in preparation.

ABSTRACT

Background. Patients undergoing surgery in the Trendelenburg position may be at risk for postoperative vision loss associated with increased intraocular pressure. The purpose of this systematic review and meta-analysis is to estimate the magnitude of the increase in intraocular pressure in adult patients.

Methods. Comprehensive search strategies were used to identify 18 eligible studies ($N = 762$). Standardized mean difference effect sizes were calculated for nine intraoperative time points (T).

Results. Using a random effects model, meta-analysis showed that intraocular pressure increased significantly after abdominal insufflation (T2: $d = 1.89, p = < 0.001$) and during Trendelenburg position (T3: $d = 1.34, p < 0.001$; T4: $d = 0.91, p < 0.001$; T6: $d = 0.30, p < 0.001$; T8: $d = 0.38, p < 0.001$).

Conclusions. Intraocular pressure increases of this magnitude demonstrate the need for implementing intraoperative interventions to reduce the risk for postoperative vision loss in patients undergoing surgery in the Trendelenburg position.

Keywords: intraocular pressure, Trendelenburg position, head-down tilt, pneumoperitoneum, Perioperative Patient Focused Model

In the Trendelenburg position, the patient's feet are higher than the patient's head by 15 degrees to 30 degrees (MacDonald & Washington, 2012). Many surgeons use a steep Trendelenburg position of 30 degrees to 45 degrees, particularly during laparoscopic and robotic surgery. The benefit of the Trendelenburg position is that it moves the abdominal viscera cephalad to improve visibility and surgical access to the abdominal and pelvic organs. However, there are potential harms associated with the Trendelenburg position. The Trendelenburg position increases intraocular pressure (IOP; Akhavan, Gainsburg, & Stock, 2010; Astuto, Minardi, Uva, & Gullo, 2011; Awad et al., 2009; Borahay et al., 2013; Cullen & Ferguson, 2012; Ghomi, 2012; Kan, Brown, & Gainsburg, 2015; Mondzelewski et al., 2015; Taketani et al., 2015). According to the American Academy of Ophthalmology (2018), normal IOP is 10 millimeters of mercury (mmHg) to 21 mmHg. Intraocular pressures higher than 21 mmHg pose a risk for glaucoma, detached retina, and postoperative vision loss (Astuto et al., 2011; Borahay et al., 2013; Emery et al., 2015; Ghomi, Kramer, Askari, Chavan, & Einarsson, 2012; Gkegkes, Karydis, Tyritzis, & Iavazzo, 2015; Gould, Cull, Wu, & Osmundsen, 2012; Grosso et al., 2013; Hoskikawa et al., 2014; Lee & Newman, 2018; Taketani et al., 2015).

Pathogenesis of Postoperative Vision Loss

The specific pathogenesis of postoperative vision loss associated with increased IOP is unclear; however, it is known that elevated IOP can lead to optic nerve injury and decreased ocular perfusion pressure resulting in ischemic optic neuropathy (Kan et al., 2015; Newman, 2008). Ischemic optic neuropathy is the most common cause of postoperative vision loss (Kan et al., 2015; Newman, 2008). The ischemic process can occur as a direct result of decreased blood supply from the arteries of the optic nerve or

by venous stasis that occurs as a result of decreased venous outflow (Gkegkes et al., 2015; Kan et al., 2015; Molloy, 2011). Periorbital swelling and venous congestion resulting from the Trendelenburg position can lead to a compartment syndrome in the orbital space that compromises blood flow to the eye, retina, and optic nerves (Molloy, 2011; Yoo et al., 2014). The amount of subsequent postoperative vision loss can range from temporary blurring to partial to complete blindness; however, once a loss of vision occurs, it is an irreversible complication (Emery et al., 2015; Lee & Newman, 2018).

Some researchers have found that increased IOP associated with the Trendelenburg position poses a greater risk for postoperative vision loss in patients who have existing ocular disease compared with patients who do not have ocular disease (Astuto et al., 2011; Borahay et al., 2013; Grosso et al., 2017; Hoshikawa et al., 2014; Mondzelewski et al., 2015; Taketani et al., 2015). Older patients with elevated baseline IOPs are also at greater risk for ischemic optic neuropathy than younger patients with normal baseline IOPs (Freshcoln & Diehl, 2014; Ghomi, 2012; Grosso et al., 2013; Kan et al., 2015; Mondzelewski et al., 2015). Likewise, patients with cardiovascular deficits may be at greater risk for postoperative vision loss than patients without cardiovascular deficits (Borahay et al., 2013). The increase in IOP and subsequent risk for postoperative vision loss is related to the amount of time the patient is in the Trendelenburg position (Mizumoto, Gosh, Iwaki, Zako, 2017; Nuzzi & Tridico, 2016; Pinkney et al., (2012); Stang-Veldhouse, Yeu, Rothenberg, & Mizen, 2010; Yoo et al., 2014).

Incidence of Postoperative Vision Loss

The incidence of postoperative vision loss following nonocular surgery has been estimated to be as low as 0.0002% and as high as 0.2% (Berg, Harrison, & Lee, 2010;

Newman, 2008); however, the incidence of postoperative vision loss in patients undergoing surgery in the Trendelenburg position remains unknown. The potential for serious ocular consequences associated with the use of the Trendelenburg position, including retinal detachment and postoperative vision loss, was first reported by Hewer (1956). To identify cases of ischemic optic neuropathy associated with prostatectomy procedures performed in the Trendelenburg position, Lee (2011) reviewed the American Society of Anesthesiologists Postoperative Vision Loss Registry, a database of 175 cases of postoperative vision loss occurring between 1987 and 2010, and found six cases. Case reports of postoperative vision loss following surgical procedures where the patient was in the Trendelenburg position have also been published.

Williams et al., (1999) reported a case of bilateral anterior ischemic optic neuropathy and branch retinal artery occlusion in a 50-year-old man undergoing laparoscopic radical prostatectomy in the Trendelenburg position. On the second postoperative day, the patient complained of bilateral vision loss. Weber, Colyer, Lesser, and Subramanian (2007) reported two cases of bilateral posterior ischemic optic neuropathy following radical prostatectomy in the Trendelenburg position. The first case involved a 62-year-old man undergoing a robotic-assisted laparoscopic procedure. On the first postoperative day, the patient complained of “purple vision” and loss of inferior visual fields in both eyes (p. 285). The second case involved a 64-year-old man undergoing a laparoscopic procedure (without robotic technology). After surgery, the patient complained of seeing a rainbow in his superior visual fields after which everything went black. Two months after surgery, the patient had no improvement in his vision.

Mizrahi, Hugkulstone, Vyakarnam, and Parker (2011) and Kumar and Vyakarnam (2013) reported a case of bilateral posterior optic neuropathy after laparoscopic colorectal surgery in a 58-year-old man with mild hypertension and obesity. The patient was positioned in a 45-degree Trendelenburg position for more than five hours. Approximately 14 hours postoperatively the patient complained of blurred vision. Six months after surgery, vision in his left eye was 6/7.5, with no improvement in his right eye. Molloy (2011) also described a case of posterior ischemic optic neuropathy resulting in bilateral blindness in a 63-year-old man following a laparoscopic prostatectomy in the steep Trendelenburg position.

Conceptual Framework

The Perioperative Patient Focused Model (Figure 3.1; Rothrock & Smith, 2000) provides a conceptual framework for this systematic review and meta-analyses. During operative procedures when the patient is anesthetized, the perioperative registered nurse (RN) fills an important role as the patient's advocate and also oversees the patient's perioperative care. Secondary to being patient focused, the Perioperative Patient Focused Model is outcome focused (Rothrock & Smith, 2000). The care provided by perioperative RNs is directed toward achieving high quality patient outcomes, such as preventing postoperative vision loss associated with increased IOP. Providing optimal perioperative patient care, serving as the patient's advocate, and achieving high quality outcomes requires specific perioperative knowledge and skillful implementation of best practices for patient positioning. Consistent with the Perioperative Patient Focused Model, the evidence-based practices implemented by perioperative RNs during patient positioning promote patient safety, and optimal physiological and behavioral responses in

perioperative patients. Providing evidence about the magnitude of the increase in IOP resulting from the intraoperative use of the Trendelenburg position will support the implementation of nursing interventions to help prevent postoperative vision loss and other ocular complications in adult patients undergoing surgery in the Trendelenburg position.

Purpose

Although some researchers have studied the quantitative increase of IOP in surgical patients in the Trendelenburg position (Astuto et al., 2011; Awad et al., 2009; Borahay et al., 2013; Grosso et al., 2013; Hoshikawa et al., 2014; Molloy, 2011; Mondzelewski et al., 2015; Taketani et al., 2015), there is a need for systematic review and meta-analysis of these studies to demonstrate the overall effect size and provide high-quality evidence supporting the implementation of intraoperative interventions designed to mitigate the increase of IOP and reduce the risk for postoperative vision loss. Meta-analysis research methods provide increased power compared to individual studies, improve estimates of effect size, and help resolve uncertainty when the results of individual studies disagree (Berlin, 1995). Because all of the evidence pertaining to a particular phenomenon is included in the analysis, meta-analysis research provides a high level of objectivity, precision, and generalizability (Thompson, 1994). Currently, there has been no quantitative meta-analytic synthesis of the existing studies examining the increase in IOP in adult patients undergoing surgery in the Trendelenburg position. The purpose of this systematic review and meta-analysis is to estimate the magnitude of the increase in IOP at selected perioperative time points in adult patients (i.e., individuals 18 years and older) undergoing any type of surgery in the Trendelenburg position.

Consequently, the research question to be addressed by this systematic review and meta-analysis is, “What is the magnitude of the increase in IOP at specific perioperative time points in adults undergoing surgery in the Trendelenburg position?”

Methods

To ensure rigorous and transparent presentation of the methods and results of this systematic review and meta-analysis, the Preferred Reporting Items for Systematic reviews and Meta-Analyses (PRISMA) guidelines have been followed (Moher, Liberati, Tetzlaff, Altman, & PRISMA Group, 2009).

Search Strategies

An expert health sciences reference librarian was consulted to identify the most appropriate search terms and databases and to assist with refining search strategies for an exhaustive and varied literature search. Keywords or medical subject headings (MeSH) included *intraocular pressure* or *ocular tension*, and *Trendelenburg position* or *head-down tilt*. Search strategies included

- online searching of the PubMed, CINAHL, and Scopus databases, and the Cochrane Database of Systematic Reviews for published and unpublished literature;
- ancestry searching of references from relevant reports to locate additional applicable references;
- author searching of individuals identified in the literature as experts in the field; and a
- dissertation search of the ProQuest database.

The author reviewed report abstracts for eligibility and obtained full-text copies of potentially eligible reports.

Inclusion and Exclusion Criteria

Criteria for inclusion in the meta-analysis were reports written in English; studies reported between January 1, 1990 and September 30, 2018; published or unpublished reports of primary studies that encompassed dissertations, conference abstracts, and presentations; studies that used either a one-group, pretest posttest design or a multiple-group, pretest posttest design; reports where the minimum age of the study participants was 18-years or older; studies that included a specific measured outcome of IOP using any type of tonometer; studies where the participants received any type of general anesthesia; and reports of studies that included sufficient data to calculate an effect size. The year 1990 was selected as the initial searching date because the first laparoscopic cholecystectomy was performed in 1987 (Vecchio, MacFayden, & Palazzo, 2000). The use of the steep Trendelenburg position has increased dramatically with the introduction and use of laparoscopic and robotic surgery. When reports did not include sufficient data to calculate an effect size, the author contacted the researchers on at least two separate occasions two to three weeks apart to obtain missing data.

Reports were excluded if the IOP was measured in adults not undergoing surgery. Reports were also excluded if data from only one time point of IOP measurement were provided. Participant groups were additionally excluded if they were receiving an intervention specifically intended to mitigate IOP; however, participant groups representing control arms receiving placebos or no interventions were included in the systematic review and meta-analysis.

Risk of Bias Within Individual Studies

To assess the risk of bias within individual studies, the Association of periOperative Registered Nurses (AORN) Research Evidence Appraisal Tool – Study,

available on the AORN website (<https://www.aorn.org/guidelines/about-aorn-guidelines/evidence-rating>) was independently used by the author and an experienced, evidence reviewer to evaluate and critically appraise each study for its level of strength and quality. When using the AORN appraisal tool, strong study designs (e.g., randomized controlled trials) are assigned the highest level of strength (i.e., experimental). Non-experimental designs (e.g., observational studies) are assigned the lowest level of strength and quasi-experimental designs are assigned a moderate level of strength. Measures such as sample size, generalizability, bias, reliability, and validity are assessed to determine whether the study quality is high, good, or low. The author and evidence reviewer participated in conference calls to discuss their independent appraisals until 100 percent consensus was achieved on study design and quality levels for each of the included studies.

Risk of Bias Across Studies

Risk of bias that may affect cumulative evidence was managed using several strategies (Valentine, 2009). To avoid bias due to a narrow or limited search (White, 2009), a comprehensive and diverse literature search was conducted. Only research studies were included in the systematic review and meta-analysis to ensure the included studies were of sufficient strength and quality. As well, studies that included objective measurements of IOP at less than two or more perioperative time points were excluded. An analysis of publication bias was also conducted to determine whether unpublished research was unintentionally excluded.

Coding and Data Extraction

An iterative process that included studying codebooks used by experienced meta-analysts for data extraction and coding of their research studies was used to develop the codebook. The relevant literature was also reviewed. After consultation with experienced meta-analysts and content analysis of studies specific to the effects of Trendelenburg position on IOP, the codebook was revised. The codebook was pilot-tested by the author using 10 randomly selected studies before being used to code and extract data from all eligible reports to identify missed coding categories and verify fit between coding categories and study characteristics (Wilson, 2009).

To extract data necessary to address the research question and statistically analyze the effects and methodologies of each of the reports, the author coded eligible studies using the developed codebook. Effect size data for each of the reports included in the systematic review and meta-analysis was independently coded by a trained researcher. The author and independent researcher discussed coding discrepancies until 100 percent consensus was achieved on effect size data for each of the eligible studies.

Data collected from each eligible study included study characteristics (i.e., authors, year of publication, publication status, geographic location, reported funding) and data related to study design (i.e., type of study, study quality, type of tonometer, inclusion of ophthalmologic exams by participants). When available, data related to participant and surgery characteristics (i.e., age; gender; American Society of Anesthesiologists [ASA; 2018] Physical Status Classification; body mass index [BMI]; comorbidities; type of surgery; degree of Trendelenburg; intra-abdominal pressure; type of anesthesia; duration of anesthesia, pneumoperitoneum, Trendelenburg position, and surgery; estimated blood loss [EBL]) were collected. Data necessary to calculate effect

sizes were extracted for the multiple time points recorded by the researchers during the perioperative phases of the procedures included in their studies.

Analyses

Meta-analyses were conducted using *Comprehensive Meta-Analysis Software*, Version 3, a statistical software developed specifically for meta-analysis research (Borenstein, Hedges, Higgins, & Rothstein, 2018). Time points for meta-analysis were selected from the time points recorded by the researchers to allow for similarity with the order of events as they occur during surgery and to achieve the greatest number of comparisons for analysis at each time point. Analyses were conducted for the nine time points (T) described in the Sidebar. Standardized mean difference effect sizes were calculated for each participant group and each measured time point (Cohen, 1992).

To account for sample size and adjust for bias, effect size values were weighted by the inverse of the variance. To account for between- and within-study variation, and because heterogeneity was observed among study designs, sample attributes, and outcome measures, a random effects model was selected a priori to synthesize effect sizes. Using a random effects model assumes that the true effect size varies from one study to the next (Borenstein, Hedges, Higgins, & Rothstein, 2009a, p. 77). A random effects model was used for seven analyses (T1, T2, T3, T5, T6, T7, T8). A fixed effect model was used to synthesize effect sizes for two analyses (T4, T9) because the number of included studies at these time points was limited. Using a fixed effect model assumes that the true effect size is the same for all studies (Borenstein et al., 2009a, p.77). Borenstein et al., (2009a, p. 84) suggest using a fixed effect analysis when the number of included studies is limited, even when heterogeneity among the studies is observed,

because when using a random effects analysis, the estimate of between-studies variance will have poor precision. Relative to interpretation of effect size, Cohen (1992) suggested that 0.2 be considered a small effect size, 0.5 be considered a medium effect size, and 0.8 or greater be considered a large effect size. Following the procedures described by Lipsey and Wilson (2001), the calculated effect sizes were converted to the metric used to measure IOP (i.e., mmHg) to facilitate interpretation of effect size findings.

The extent of publication bias for the meta-analysis was assessed by constructing a funnel plot. Notably, a funnel plot may suggest publication bias, but does not eliminate the bias (Sutton, 2009). In order to create a funnel plot, there must be a minimum of three studies (Borenstein et al., 2018). An Egger's test using linear regression was also conducted to measure asymmetry of the funnel plot (Borenstein et al., 2018; Egger, Smith, Schneider, & Minder, 1997; Sterne Egger, & Moher, 2011). Using an Egger's test is not advised when there are less than 10 studies included in the meta-analysis because the power of the test may be too low to distinguish true asymmetry from chance (Sterne et al., 2011). Therefore, when the analysis included 10 or more participant groups (T1, T2, T6), a funnel plot was constructed and an Egger's test was conducted.

The goal of a meta-analysis is three-fold: 1) to obtain a global effect size for the intervention or phenomenon being studied, 2) to determine whether the studies are homogeneous, and 3) if the studies are heterogeneous, to identify possible variables or characteristics moderating the meta-analysis results (Huedo-Medina, Sánchez-Meca, Marín-Martínez, & Botella, 2006). After deciding on the model and calculating effect sizes, the studies included in the meta-analysis were assessed for heterogeneity. According to the Cochrane Collaboration, heterogeneity is any kind of variability among

the studies included in a meta-analysis (Deeks, Higgins, & Altman, 2011). Heterogeneity testing explores the null hypothesis (i.e., that the same effect is being evaluated by all studies; Higgins, Thompson, Deeks, & Altman, 2003). Heterogeneity among the included studies in a meta-analysis is very common and should be anticipated, not regarded as the exception (Berlin, 1995).

Homogeneity of variance among effect sizes was tested using Cochrane's Q, which estimates statistical significance; Tau-squared (T^2), which estimates the absolute value of the true variance between studies, but not the proportion of the variance; and I^2 -squared (I^2), which estimates the proportion of true variance, but not the absolute value of the variance, (Borenstein, Hedges, Higgins, & Rothstein, 2009b). Higgins and Thompson (2002) recommend using an I^2 test for quantifying the impact of heterogeneity in a meta-analysis as this test quantifies the influence (as opposed to the amount) of heterogeneity and expresses the percentage of variability due to heterogeneity rather than chance (p. 1553). An I^2 value of 0% indicates there is no heterogeneity, values of 25% reflect low observed variation, values of 50% reflect moderate levels, and values of 75%, reflect high levels of observed variation (Higgins et al., 2003). Prediction intervals for each time point were also calculated to show the dispersion of true effect sizes around the mean (Borenstein, Hedges, Higgins, & Rothstein, 2009c).

Results

The flow of study selection is depicted in Figure 3.2. In total, 2693 records were identified for possible inclusion, and of these, 18 studies were included in the systematic review and meta-analysis. Four non-experimental studies had multiple participant groups (Agrawal, Dureja, Verma, & Kang, 2013; Kaur, Sharma, Kalra, Purohit, & Chauhan,

2018; Nishikawa, Watanabe, & Kurahashi, 2017; Yoo et al., 2015), resulting in a total of 24 participant groups and 762 participants for analysis. Table 3.1 contains a summary of the studies included in this review and meta-analysis.

Study Characteristics

All 18 studies included in the systematic review and meta-analysis were obtained from peer-reviewed journals. The researchers of six studies (33.3%) reported receiving some type of funding or donated supplies (Adisa, Onakpoya, Adenekan, & Awe, 2016; Blecha et al., 2017; Hirooka et al., 2018; Molloy & Cong, 2014; Molloy, Cong, & Watson, 2016; Yoo et al., 2015). Although the literature was searched from 1990 through 2018, studies included in the systematic review and meta-analysis were published between 2011 and 2018. Some earlier studies were located during the literature search (Awad et al., 2009; Lentschener, Benhamou, Niessen, Mercier, & Fernandez, 1996; Mowafi, Al-Ghamdi, & Rushood, 2003); however, these were excluded because of insufficient effect size data. The greatest number of studies (s) were published in 2015 ($s = 4$) and 2018 ($s = 4$), but the greatest number of participants occurred in 2013 ($n = 147$). The majority ($s = 14$) were conducted in Asia ($s = 7$) or North America ($s = 7$), with the majority of participants also from Asia ($n = 361$) or North America ($n = 292$).

Participant Characteristics

Participant and surgery characteristics are shown in Table 3.2. The mean age of the participants was 55.2 years (± 11.9) and ranged from 30.5 years to 66.9 years. The majority of the participants were men ($n = 415$; 67.0%). Participant race was only reported by two researchers (Adisa et al, 2016; Borahay et al, 2013). Socioeconomic status was not reported by any researchers. Participants were slightly overweight with a

mean BMI of 27.5 kg/m² (\pm 2.3; Centers for Disease Control and Prevention, 2016). The ASA physical classification status was reported by the researchers of 10 reports. Some researchers reported that all participants were ASA class I or II ($s = 2$; Agrawal et al., 2013; Taketani et al., 2015), other researchers reported the number or percentage of participants who were ASA class I, II, or III ($s = 8$; Adisa et al., 2016; Blecha et al., 2017; Grosso, et al., 2013; Kaur et al., 2018; Kitamura et al., 2018; Molloy, 2011; Molloy & Watson, 2012; Yoo et al., 2015). The majority of participants ($n = 432$; 95.4%) were ASA class I (healthy) or II (with mild systemic disease; American Society of Anesthesiologists, 2018). Notably, some researchers had exclusion criteria for participant age, BMI, and ASA classification (see Table 3.1). Researchers reported patient comorbidities for 191 participants ($s = 6$) that included asthma ($n = 2$ of 76; 2.6%), diabetes ($n = 8$ of 107; 7.5%), and hypertension ($n = 44$ of 117; 37.6%; Borahay et al., 2013; Kitamura et al., 2018; Molloy, 2011; Molloy & Watson, 2012; Raz et al., 2015; Yoo et al., 2015).

Surgery Characteristics

The majority of the participants underwent robotic-assisted laparoscopic radical prostatectomy ($n = 335$; 44.0%) or laparoscopic gynecologic surgery ($n = 151$; 19.8%). All participants ($N = 762$) received general anesthesia by either inhalation ($n = 298$; 39.1%), intravenous propofol ($n = 242$; 31.8%), or unspecified methods ($n = 222$; 29.1%). Intra-abdominal pressure for maintaining carbon dioxide pneumoperitoneum ranged between 12 mmHg and 15 mmHg with a mean of 13.6 mmHg (\pm 1.3). The mean degree of Trendelenburg was 28.4 (\pm 6.5) with a range of 17.5 to 45. The mean duration

of surgery was 197.9 minutes (\pm 64.4) with a range of 111 minutes to 318 minutes. The mean EBL was 252.4 milliliters (mL; \pm 113.3) with a range of 69.4 mL to 467 mL.

Study Design Characteristics

The 18 reports included in the systematic review and meta-analysis comprised four experimental ($n = 100$; Kitamura et al., 2018; Mathew et al., 2018; Molloy et al., 2016; Raz et al., 2015), two quasi-experimental ($n = 168$; Molloy & Cong, 2014; Molloy & Watson, 2012), and 12 non-experimental ($n = 494$; Adisa et al., 2016; Agrawal et al., 2013; Blecha et al., 2017; Borahay et al., 2013; Grosso et al., 2013; Hirooka et al., 2018; Kaur et al., 2018; Molloy, 2011; Mondzelewski et al., 2015; Nishikawa et al., 2017; Taketani et al., 2015; Yoo et al., 2015). Nine were high quality ($n = 461$; Blecha et al., 2017; Kaur et al., 2018; Kitamura et al., 2018; Mathew et al., 2018; Molloy, 2011; Molloy & Cong, 2014; Molloy et al., 2016; Molloy & Watson, 2012; Yoo et al., 2015) and nine were good quality ($n = 301$; Adisa et al., 2016; Agrawal et al., 2013; Borahay et al., 2013; Grosso et al., 2013; Hirooka et al., 2018; Mondzelewski et al., 2015; Nishikawa et al., 2017; Raz et al., 2015; Taketani et al., 2015). The researchers used five different types of tonometers to measure IOP, the Tono-Pen XL was used most frequently ($s = 9$; $n = 420$). The researchers reported having 195 participants undergo preoperative ophthalmologic examinations ($s = 7$; Adisa et al., 2016; Grosso et al., 2013; Hirooka et al., 2018; Kaur et al., 2018; Mathew et al., 2018; Mondzelewski et al., 2015; Taketani et al., 2015), and 100 participants undergo postoperative ophthalmologic examinations ($s = 4$; Hirooka et al., 2018; Mathew et al., 2018; Mondzelewski et al., 2015; Taketani et al., 2015). Notably, all participants who received postoperative ophthalmologic examinations also received preoperative examinations ($n = 100$).

Effect Sizes

Results of the synthesized effect sizes, prediction intervals, and meta-analyses for each time point of IOP measurement are shown in Table 3.3. A graphical representation of the magnitude of changes in IOP for T0 through T9 is shown in Figure 3.3. In total, between abdominal insufflation in supine position (T2) and 5 minutes (T3), 60 minutes (T4), 150 minutes (T6), and 240 minutes (T8) of Trendelenburg position, IOP increases significantly by 13.6 mmHg (i.e., 3.5 mmHg + 4.4 mmHg + 2.6 mmHg + 1.5 mmHg + 1.6 mmHg = 13.6 mmHg). Based on the upper limits of the prediction intervals (Figure 3.4), in 95% of all populations, IOP could increase by as much as 28.1 mmHg (i.e., 7.6 mmHg + 8.5 mmHg + 6.6 mmHg + 2.3 mmHg + 3.1 mmHg = 28.1 mmHg). The greatest increase in IOP occurs after the patient is placed into the Trendelenburg position (T3: +4.4 mmHg). The IOP continues to increase significantly while the patient is in Trendelenburg position, but to a lesser degree (T4: +2.6 mmHg; T6: +1.5 mmHg; T8: +1.6 mmHg). Intraocular pressure decreases significantly after induction of anesthesia in supine position (T1: -5.2 mmHg) and after a return to supine position for arousal from anesthesia (T5: -7.5 mmHg; T7: -8.2 mmHg; T9: -6.0 mmHg). The forest plot of effect sizes for each participant group included in the meta-analysis for T6 is shown in Figure 3.5. The funnel plot for publication bias for T6 is shown in Figure 3.6.

Discussion

The results of this systematic review and analysis have shown that IOP increases significantly for adult patients undergoing surgery in the Trendelenburg position. As shown in Figure 3.3, if an individual had a baseline IOP of 16.5 mmHg before induction of anesthesia (as indicated by the pooled mean calculated for T0), after 180 minutes to

240 minutes in the Trendelenburg position, the patient's IOP could increase to 24.9 mmHg ($16.5 \text{ mmHg} - 5.2 \text{ mmHg} + 3.5 \text{ mmHg} + 4.4 \text{ mmHg} + 2.6 \text{ mmHg} + 1.5 \text{ mmHg} + 1.6 \text{ mmHg} = 24.9 \text{ mmHg}$). Based on the upper limits of the prediction intervals (Figure 3.4), after 180 minutes to 240 minutes in the Trendelenburg position, IOP could increase to 35 mmHg ($16.5 \text{ mmHg} - 9.6 \text{ mmHg} + 7.6 \text{ mmHg} + 8.5 \text{ mmHg} + 6.6 \text{ mmHg} + 2.3 \text{ mmHg} + 3.1 \text{ mmHg} = 35 \text{ mmHg}$). An IOP of 24.9 mmHg to 35 mmHg is above the highest parameter of normal IOP (i.e., 21 mmHg). As shown in Table 3.2, the mean duration of Trendelenburg position for the studies included in this systematic review and meta-analysis is 104.8 minutes (± 58.2) with a range of 68 minutes to 207 minutes. The greatest increases in IOP occur during abdominal insufflation and within the first 60 minutes after Trendelenburg position; however, based on the collective range of 68 minutes to 207 minutes for duration of Trendelenburg position, a Trendelenburg time of 180 minutes to 240 minutes is not implausible. Another important consideration regarding the findings of this systematic review and meta-analysis is that the mean degree of Trendelenburg position was 28.4. Steep Trendelenburg is generally considered to be a head-down tilt of 30 degrees to 45 degrees (Demasi, Porpiglia, Tempia, & D'Amelio, 2017; Ghomi et al., 2012; Gould et al., 2012); thus, it is likely that Trendelenburg positions greater than 28.4 degrees would have produced even greater increases in IOP.

Implications for Practice

Increased IOP puts the patient at risk for glaucoma, detached retina, or partial to complete vision loss (Astuto et al., 2011; Borahay et al., 2013; Emery et al., 2015; Ghomi et al., 2012; Gkegkes et al., 2015; Gould et al., 2012; Grosso et al., 2013; Hoskikawa et al., 2014; Lee & Newman, 2018; Taketani et al., 2015). Intraocular pressure increases of

the magnitude found in this systematic review and meta-analysis clearly demonstrate the need for implementing intraoperative interventions to mitigate the increase in IOP and reduce the potential for serious ocular complications in patients undergoing surgery in the Trendelenburg position. These intraoperative interventions may include

- monitoring IOP at established intervals or continuously (Hoshikawa et al., 2013; Lee, Dallas, Daniel, & Cotter, 2016; Vitish-Sharma et al., 2018),
- reducing the degree of Trendelenburg position (Ghomie et al., 2012; Mathew et al., 2018; Ozcan et al., 2017; Raz et al., 2015),
- implementing a modified Trendelenburg position (Raz et al., 2015),
- providing periodic position changes or rest periods (Blecha et al., 2017; Borahay et al., 2013; Freshcoln & Diehl, 2014; Gkegkes et al., 2015; Gould et al., 2012; Mizrahi et al., 2011; Molloy & Watson, 2012), and
- administering specific medications or anesthetics (Agrawal et al., 2013; Hwang et al., 2013; Joo, Koh, Lee, & Lee, 2016; Joo, Kim, & Lee, 2017; Kan et al., 2015; Kaur et al., 2018; Kim et al., 2015; Mathew et al., 2018; Mowafi et al., 2003).

Because IOP increases during abdominal insufflation and Trendelenburg position, intraoperative monitoring of IOP either continuously or at established intervals or time points (e.g., after abdominal insufflation, after initiation of Trendelenburg position, after 60 minutes of Trendelenburg position, etc.) seems prudent. Elevated IOPs can be an indication of ocular venous congestion and decreased perfusion of the optic nerve (Yoo et al., 2014). Monitoring IOP can provide a baseline IOP and an objective measure that can help the surgical team maintain awareness of the patient's IOP, implement interventions

to reduce IOP as needed, and thus reduce the potential for ocular complications and postoperative vision loss (Hoshikawa et al., 2013; Lee et al., 2016).

Steeper degrees of Trendelenburg increase the risk for postoperative complications because they place greater physiologic stress on the patient's body (Ghomie et al., 2012; Gould et al., 2012; Kadono et al., 2013). Ghomi et al., (2012) found that robotic-assisted gynecologic surgery could be performed successfully with a modest head-down tilt of 16.4 degrees. In a study to determine the head-down tilt necessary to provide surgical access and visibility, Gould et al., (2012) found the mean head-down tilt most often selected by the surgeons was 28.1 degrees, which was much less than the 40-degree head-down tilt the surgeons were using.

Raz et al., (2015) found that modifying the Trendelenburg position so that the patient's head and shoulders remained level significantly decreased IOP and accelerated its return to baseline levels. Implementing periodic intraoperative position changes or rest periods in supine position (or positions where the ocular level is above the heart) can help to reduce IOP. In a quasi-experimental study, Molloy and Watson (2012) implemented a five-to-seven-minute level supine intervention after 60 minutes of 32-degree to 40-degree Trendelenburg position and found there was a significant decrease in IOP after 120 minutes of Trendelenburg position (Intervention: $18.7 \text{ mmHg} \pm 5.22$; Control: $35.7 \text{ mmHg} \pm 10.56$; $p < 0.001$). The dramatic and significant decrease in IOP that occurs before arousal from anesthesia found in this systematic review and meta-analysis (T5: -7.5 mmHg, $p < 0.001$; T7: -8.2 mmHg, $p < 0.001$; T9: -6.0 mmHg, $p < 0.001$) also supports the implementation of periodic intraoperative position changes or rest periods as a mechanism to help reduce IOP.

Administering specific medications or anesthetics may also be effective in reducing IOP or mitigating the intraoperative increase in IOP (Agrawal et al., 2013; Hwang et al., 2013; Joo et al., 2016; Joo et al., 2017; Lee et al., 2016; Kaur et al., 2018; Kim et al., 2015; Mathew et al., 2018; Mowafi et al., 2003). Agrawal et al., (2013) found that induction and maintenance of anesthesia with intravenous propofol was the most effective option for mitigating the increase in IOP in adult patients undergoing surgery in the Trendelenburg position. Likewise, Kaur et al., (2018) found that propofol-based total intravenous anesthesia was more effective than inhalational anesthesia with sevoflurane in mitigating the increase in IOP in patients undergoing laparoscopic surgery in the Trendelenburg position. Kitamura et al., (2018) found that continuous administration of dexmedetomidine in combination with propofol-based total intravenous anesthesia decreased IOP in patients undergoing robotic-assisted laparoscopic radical prostatectomy. Molloy and Cong (2014) found that intraoperative treatment with dorzolamide-timolol eyedrops significantly reduced elevated IOP in patients undergoing lengthy laparoscopic procedures in the Trendelenburg position, while Molloy et al., (2016) found that prophylactic therapy with dorzolamide-timolol eyedrops significantly reduced IOP in patients undergoing robotic-assisted laparoscopic prostate and gynecologic procedures.

Another important consideration for practice is the need to evaluate whether patients undergoing surgery in the Trendelenburg position should receive a preoperative ophthalmologic examination to reduce the risk for ocular injury (Borahay et al., 2013; Lee et al., 2016). Preoperative ophthalmologic examinations may be helpful in identifying patients at risk for postoperative vision loss. Increases in IOP may be more harmful in older patients or patients who are predisposed to developing diabetes or

glaucoma than in younger, healthier patients (Borahay et al., 2013; Grosso et al., 2013; Mondzelewski et al., 2015; Taketani et al., 2015).

The results of this systematic review and meta-analysis support the use of the Perioperative Patient Focused Model as the conceptual foundation for this perioperative research. Providing quantitative evidence about the magnitude of the intraoperative increase in IOP resulting from the Trendelenburg position supports the implementation of nursing interventions that are patient-focused and will improve patient outcomes by mitigating increases in IOP and reducing the risk for permanent postoperative vision loss and other ocular complications in adult surgical patients.

Implications for Future Research

Further research relative to the magnitude of IOP increases in patients undergoing surgery in the Trendelenburg position is warranted. Further research to provide validation and demonstrate reliability of the Perioperative Patient Focused Model is also warranted. To allow for consistent data collection, comparison, meta-analysis, and reporting, researchers of future studies should use standardized time points for measurement (i.e., before arousal, after arousal, after abdominal insufflation, after change to Trendelenburg position and every 30 minutes to 60 minutes thereafter, after return to supine position, before arousal, and postoperatively). Further, researchers should present data in a consistent format for each time point (i.e., sample size, mean, standard deviation). Additionally, to determine whether certain variables affect the strength of the relationship between Trendelenburg position and IOP, researchers should include patients of all ages (e.g., children, older adults), without restriction of BMI or comorbidities.

Limitations

This systematic review and meta-analyses has several limitations. The literature search yielded 107 potentially eligible studies. Studies were excluded for a variety of reasons (see Figure 3.2); however, 25 studies were excluded from the analyses because of a lack of data necessary to calculate an effect size. The researchers were contacted a minimum of two times to obtain missing data, but most did not respond. Some of the researchers excluded participants based on age, BMI, and comorbidities; therefore, the mean values for these variables may not fully reflect the true characteristics of all adult surgical patients. Because researchers measured IOP at different intraoperative time points, all studies could not be included at all time points examined in the meta-analysis. Likewise, there were not enough studies included at each time point to allow for moderator analyses. With the exception of T6 and T8, heterogeneity was significant, indicating that variation across studies was substantial, potentially limiting generalizability. The non-significant Egger's regression intercept ($\text{bias} = -0.05; p = 0.47$) is indicative of the absence of bias in the studies included in the meta-analysis for the T6 time point ($s = 12$); however, the Egger's test has low power for meta-analyses containing small to moderate numbers of studies (Sutton, 2009).

Conclusion

Intraocular pressure increases significantly between abdominal insufflation in supine position and 240 minutes of Trendelenburg position. The greatest increases in IOP occur after insufflation of the abdomen (while the patient is in the supine position) and within five minutes after the patient is placed into the Trendelenburg position. The IOP continues to increase significantly while the patient is in Trendelenburg position, but to a lesser degree. Intraocular pressure increases of the magnitude found in this systematic

review and meta-analysis clearly demonstrate the need for implementing intraoperative interventions to mitigate the increase in IOP and reduce the risk for postoperative vision loss and other ocular complications in patients undergoing surgery in the Trendelenburg position.

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Table 3.1.

Summary of Studies Included in the Systematic Review and Meta-analysis (s = 18).

First Author (Year) Country	Study Design (Quality)^a	Participant Groups and Observations/Interventions	IOP Measures (Tonometer)	Outcome Measures	Effect Sizes (Cohen's <i>d</i>)
Adisa (2016) Nigeria	Non-experimental (Good)	<ul style="list-style-type: none"> • 20 patients undergoing laparoscopic surgery in Trendelenburg position • 20 patients undergoing laparoscopic surgery in reverse Trendelenburg position^b <p><i>Note.</i> Patients with BMI ≥ 35 kg/m² excluded.</p>	Measurement of IOP at seven different time points (Perkins)	Differences in IOP levels at each time point	T1: -1.043 T2: 0.013 T3: 0.839
Agrawal (2013) India	Non-experimental (Good)	<ul style="list-style-type: none"> • 30 women undergoing laparoscopic gynecologic surgery in Trendelenburg position with anesthesia using propofol for induction and propofol for maintenance (A) • 30 women undergoing laparoscopic gynecologic procedures in Trendelenburg position with anesthesia using propofol for induction and 1% isoflurane for maintenance (B) • 30 women undergoing laparoscopic gynecologic procedures in Trendelenburg position with anesthesia using thiopentone for induction and propofol for maintenance (C) • 30 women undergoing laparoscopic gynecologic procedures in Trendelenburg position with anesthesia using thiopentone for induction and 	Measurement of IOP at six different time points (Schiotz)	Differences in IOP levels at each time point	T1(A): -5.047 T1(B): -2.533 T1(C): -2.412 T1(D): -2.027 T2(A): 1.231 T2(B): 1.131 T2(C): 0.859 T2(D): 1.559 T3(A): 1.327 T3(B): 1.523 T3(C): 1.671 T3(D): 2.419

First Author (Year) Country	Study Design (Quality)^a	Participant Groups and Observations/Interventions	IOP Measures (Tonometer)	Outcome Measures	Effect Sizes (Cohen's <i>d</i>)
Blecha (2017) Germany	Non-experimental (High)	<p>1% isoflurane for maintenance (D)</p> <p><i>Note.</i> Patients with weight > 70 kg excluded.</p> <ul style="list-style-type: none"> 51 men undergoing robotic-assisted laparoscopic prostatectomy in Trendelenburg position <p><i>Note.</i> Patients > 80 years, ASA Class > III, or BMI > 40 kg/m² excluded.</p>	Measurement of IOP at six different time points (Icare PRO)	Differences in IOP levels at each time point	T5: -1.650
Borahay (2013) United States	Non-experimental (Good)	<ul style="list-style-type: none"> 10 women undergoing elective robotic-assisted or laparoscopic-assisted hysterectomy procedures in Trendelenburg position 	Measurement of IOP at five different time points (Tono-Pen XL)	Differences in IOP levels at each time point	T6: 0.132
Grosso (2013) Italy	Non-experimental (Good)	<ul style="list-style-type: none"> 17 patients undergoing colorectal laparoscopic surgery in Trendelenburg position 12 patients undergoing colorectal laparoscopic surgery in supine position^b <p><i>Note.</i> Patients > 45 years, ASA Class > III, or BMI > 35 kg/m² excluded.</p>	Measurement of IOP at eight different time points (Icare PRO)	Differences in IOP levels at each time point	T2: 0.975 T3: 0.204 T4: 0.010
Hirooka (2018) Japan	Non-experimental (Good)	<ul style="list-style-type: none"> 40 men undergoing robotic-assisted laparoscopic radical prostatectomy in Trendelenburg position 	Measurement of IOP at six different time points (Tono-Pen XL)	Differences in IOP levels at each time point	T4: 0.964 T6: 0.091 T8: 0.539
Kaur (2018)	Non-experimental	<ul style="list-style-type: none"> 30 patients undergoing lower abdominal laparoscopic surgery 	Measurement of IOP at seven	Differences in IOP levels at each time point	T1(P): -4.087 T1(S): -4.134

First Author (Year) Country	Study Design (Quality)^a	Participant Groups and Observations/Interventions	IOP Measures (Tonometer)	Outcome Measures	Effect Sizes (Cohen's <i>d</i>)
India	(High)	<p>in Trendelenburg position with anesthesia using intravenous propofol (P)</p> <ul style="list-style-type: none"> 30 patients undergoing lower abdominal laparoscopic surgery in Trendelenburg position with anesthesia using inhaled sevoflurane (S) <p><i>Note.</i> Patients with extreme obesity excluded.</p>	different time points (Schiotz)		T2(P): 2.275 T2(S): 4.508
Kitamura (2018) Japan	Experimental (High)	<ul style="list-style-type: none"> 20 patients undergoing robotic-assisted laparoscopic radical prostatectomy in Trendelenburg position and receiving a placebo (saline) 20 patients undergoing robotic-assisted laparoscopic radical prostatectomy in Trendelenburg position and receiving dexmedetomidine^b <p><i>Note.</i> Patients with BMI > 35 kg/m² excluded.</p>	Measurement of IOP at eight different time points (Icare PRO)	Differences in IOP levels at each time point	T1: -1.466 T6: 0.165 T8: 0.611
Mathew (2018) Canada	Experimental (High)	<ul style="list-style-type: none"> 15 men undergoing robotic-assisted laparoscopic radical prostatectomy in Trendelenburg position and receiving a placebo (artificial tears) 11 men undergoing robotic-assisted laparoscopic radical prostatectomy in Trendelenburg position and receiving brimonidine tartrate 0.2%^b 	Measurements of IOP at six different time points (Tono-Pen AVIA)	Differences in IOP levels at each time point	T1: -1.654 T6: 0.068 T8: 0.282
Molloy (2011)	Non-experimental	<ul style="list-style-type: none"> 37 patients undergoing laparoscopic surgery in 	Measurements of IOP at six	Differences in IOP levels at each time point	T5: -1.589 T6: 0.029

First Author (Year) Country	Study Design (Quality)^a	Participant Groups and Observations/Interventions	IOP Measures (Tonometer)	Outcome Measures	Effect Sizes (Cohen's <i>d</i>)
United States	(High)	Trendelenburg position	different time points (Tono-Pen XL)		T7: -1.638
Molloy (2012)	Quasi-experimental	• 37 patients undergoing laparoscopic surgery in Trendelenburg position	Measurement of IOP at six different time points	Differences in IOP levels at each time point	T5: -1.497 T6: 0.330
United States	(High)	• 29 patients undergoing laparoscopic surgery in Trendelenburg position and receiving a level supine intervention ^b	(Tono-Pen XL)		T7: -1.855
Molloy (2014)	Quasi-experimental	• 131 patients undergoing robotic-assisted laparoscopic radical prostatectomy or pelvic gynecologic procedures in Trendelenburg position	Measurements of IOP at eight different time points (Tono-Pen XL)	Differences in IOP levels at each time point	T5: -0.793 T6: 0.352
United States	(High)	• 63 patients undergoing robotic-assisted laparoscopic prostatectomy or pelvic gynecologic procedures in Trendelenburg position receiving dorzolamide-timolol when IOP exceeds 40 mmHg ^b			T7: -1.082 T8: 0.188 T9: -1.165
Molloy (2016)	Experimental	• 44 patients undergoing laparoscopic procedures in Trendelenburg position and receiving an ophthalmic placebo (balanced salt solution) after induction of anesthesia	Measurements of IOP at eight different time points (Tono-Pen XL)	Differences in IOP levels at each time point	T5: -1.519 T6: 0.497
United States	(High)	• 63 patients undergoing laparoscopic procedures in Trendelenburg position receiving dorzolamide-timolol after anesthesia induction and when IOP exceeded 40 mmHg ^b			T7: -1.922 T8: 0.536 T9: -2.758

First Author (Year) Country	Study Design (Quality)^a	Participant Groups and Observations/Interventions	IOP Measures (Tonometer)	Outcome Measures	Effect Sizes (Cohen's <i>d</i>)
Mondelewski (2015) United States	Non-experimental (Good)	<ul style="list-style-type: none"> 18 patients undergoing robotic-assisted laparoscopic procedures in Trendelenburg position 9 patients undergoing laparoscopic procedures in supine position^b 12 patients undergoing open procedures in supine position^b 	Measurements of IOP at 11 different time points (Tono-Pen AVIA)	Differences in IOP levels at each time point	
Nishikawa (2017) Japan	Non-experimental (Good)	<ul style="list-style-type: none"> 15 men undergoing robotic-assisted laparoscopic radical prostatectomy in a 25° Trendelenburg position (25) 15 men undergoing robotic-assisted laparoscopic radical prostatectomy in a 30° Trendelenburg position (30) 	Measurement of IOP at six different time points (Tono-Pen XL)	Differences in IOP levels at each time point	T1(25): -1.934 T1(30): -1.076 T5(25): -2.103 T5(30): -2.584 T6(25): 0.555 T6(30): 0.895 T7(25): -2.887 T7(30): -3.744
Raz (2015) Australia	Experimental (Good)	<ul style="list-style-type: none"> 21 men undergoing robotic-assisted laparoscopic radical prostatectomy in Trendelenburg position 29 men undergoing robotic-assisted laparoscopic radical prostatectomy in a modified-Z Trendelenburg position^b 	Measurement of IOP at 18 different time points (Tono-Pen AVIA)	Differences in IOP levels at each time point	T4: 1.238 T5: -1.093 T6: 0.135 T7: -1.593
Taketani (2015) Japan	Non-experimental (Good)	<ul style="list-style-type: none"> 25 men undergoing robotic-assisted laparoscopic radical prostatectomy in Trendelenburg position. 	Measurement of IOP at eight different time points (Tono-Pen XL)	Differences in IOP levels at each time point	T6: 0.386 T8: 0.161
Yoo (2015) Korea	Non-experimental (High)	<ul style="list-style-type: none"> 32 patients undergoing robotic-assisted laparoscopic radical prostatectomy in the 	Measurement of IOP at eight different	Differences in IOP levels at each time point	T1(M): -2.952 T1(D): -3.954 T2(M): 3.262

First Author (Year) Country	Study Design (Quality)^a	Participant Groups and Observations/Interventions	IOP Measures (Tonometer)	Outcome Measures	Effect Sizes (Cohen's <i>d</i>)
		<p>Trendelenburg position with anesthesia using moderate neuromuscular blockade (M)</p> <ul style="list-style-type: none"> • 34 patients undergoing robotic-assisted laparoscopic radical prostatectomy in the Trendelenburg position with anesthesia using deep neuromuscular blockade (D) <p><i>Note.</i> Patients with BMI > 30 kg/m² excluded.</p>	timepoints (Tono-Pen XL)		T2(D): 3.389

Note. *s* = studies; IOP = Intraocular pressure; BMI = body mass index; kg/m² = kilograms/meter-squared; kg = kilograms; ASA = American Society of Anesthesiologists Physical Classification Status; mmHg = millimeters of mercury.

^aQuality ratings are based on cumulative scores obtained from the Association of periOperative Registered Nurses (AORN) Research Evidence Appraisal Tool – Study.

^bThis group was not eligible for inclusion in the meta-analysis.

Table 3.2

Study and Participant Characteristics (s = 18; N = 762)

Category	Number of Participants (Number of Studies)	Mean \pm SD (Range) or Percent
Age (years)	612 (16)	55.2 ± 11.9 (30.5 – 66.9)
Gender	624 (16)	33.0
Women	209	67.0
Men	415	
BMI (kg/m^2)	287 (12)	27.5 ± 2.3 (23.6 – 30.7)
ASA	308 (8)	
Class I	107	34.7
Class II	180	58.4
Class III	21	6.9
ASA	145 (2)	100
Comorbidities	300 (7)	
Asthma	76 (2)	2.6
Diabetes	107 (3)	7.5
Hypertension	117 (4)	37.6
Surgery type	762 (18)	
Laparoscopic	292	38.3
Colorectal	47	6.2
Gynecologic	151	19.8
Prostatectomy	14	1.8
Unspecified	80	10.8
Laparoscopic/Robotic	44	5.8
Unspecified	44	5.8
Robotic	426	55.9
Hysterectomy	8	1.0
Pelvic node	1	0.1
Prostatectomy	335	44.0
Vaginal repair	82	10.8
Anesthesia	762 (18)	
General-Inhalation	298	39.1
General-Propofol	242	31.8
General-Unspecified	222	29.1
Intra-abdominal pressure (mmHg)	536 (12)	13.6 ± 1.3 (12 – 15)
Trendelenburg degree	762 (18)	28.4 ± 6.5 (17.5 – 45)
Duration (min)		
Anesthesia	66 (1)	158.0 ± 2.8 (156 – 160)
Pneumoperitoneum	126 (2)	94.5 ± 15.1 (80 – 109.4)
Trendelenburg	159 (3)	104.8 ± 58.2 (68 – 207)
Surgery	484 (10)	197.9 ± 64.4 (111 – 318)
Estimated blood loss (mL)	444 (11)	252.4 ± 113.3 (69.4 – 467)

Note. s = studies; SD = standard deviation; BMI = body mass index; kg/m^2 = kilograms/meter-squared; ASA = American Society of Anesthesiologists Physical Classification Status; mmHg = millimeters of mercury; min = minutes; mL = milliliters.

Table 3.3

Effect Sizes and Magnitude of Change in IOP for T1 through T9 (Sidebar; s = 18; N = 762).

Time Points	s	k	n	Model	d	95% CI		Q	I^2	Mean change in IOP ^a	Prediction Interval ^b
						LL	UL				
T1	7	13	331	R	-2.45*	-1.70	-0.30	109.82*	89.07	-5.2 mmHg	-0.9 mmHg to -9.6 mmHg
T2	5	10	283	R	1.89*	1.14	2.63	121.56*	92.60	+3.5 mmHg from T1	-0.7 mmHg to +7.6 mmHg
T3	3	6	157	R	1.34*	0.78	1.90	24.90*	79.92	+4.4 mmHg from T2	+0.4 mmHg to +8.5 mmHg
T4	3	3	78	F	0.91*	0.57	1.25	12.96**	84.57	+2.6 mmHg from T3	-1.5 mmHg to +6.6 mmHg
T5	7	8	351	R	-1.54*	-1.93	-1.16	30.71*	77.20	-7.5 mmHg from T4	-3.1 mmHg to -12.0 mmHg
T6	11	12	410	R	0.30*	0.16	0.44	7.06	0.00	+1.5 mmHg from T4	+0.9 mmHg to +2.3 mmHg
T7	6	7	300	R	-1.94*	-2.47	-1.41	33.52*	82.10	-8.2 mmHg from T6	-3.1 mmHg to -13.3 mmHg
T8	6	6	275	R	0.38*	0.12	0.65	1.90	0.00	+1.6 mmHg from T6	+0.5 mmHg to +3.1 mmHg
T9	2	2	175	F	-1.58*	-2.06	-1.11	8.46**	88.18	-6.0 mmHg from T8	-13.9 mmHg to +1.7 mmHg

Note. s = studies; k = comparisons; d = standardized mean difference; CI = confidence interval; LL = lower limit; UL = upper limit; Q = Cochrane's Q; I^2 = heterogeneity statistic; IOP = intraocular pressure; T = time point; R = Random effects; mmHg = millimeters of mercury; F = Fixed effect.

^aMean effect sizes were converted to the metric used to measure IOP (i.e., mmHg) following the procedures described by Lipsey and Wilson (2001). Lipsey, M. W., & Wilson, D. B. (2001). Interpreting and using meta-analysis results. In *Practical meta-analysis: Applied social research* (Vol 49, pp. 146-168). Thousand Oaks, CA: Sage.

* $p < 0.001$. ** $p < 0.005$.

^bIn 95% of all populations, the true effect size will fall within this range.

Sidebar

Time Points Analyzed for Changes in IOP.

- | | |
|-------------|---|
| T1 — | Before induction of anesthesia to
0 minutes to 5 minutes after induction of anesthesia |
| T2 — | 0 minutes to 5 minutes after induction of anesthesia to
0 minutes to 5 minutes after abdominal insufflation |
| T3 — | 0 minutes to 5 minutes after abdominal insufflation to
0 minutes to 5 minutes after Trendelenburg position |
| T4 — | 0 minutes to 5 minutes after Trendelenburg position to
30 minutes to 60 minutes after Trendelenburg position |
| T5 — | 30 minutes to 60 minutes after Trendelenburg position to
before arousal from general anesthesia |
| T6 — | 30 minutes to 60 minutes after Trendelenburg position to
120 minutes to 150 minutes after Trendelenburg position |
| T7 — | 120 minutes to 150 minutes after Trendelenburg position to
before arousal from general anesthesia |
| T8 — | 120 minutes to 150 minutes after Trendelenburg position to
180 minutes to 240 minutes after Trendelenburg position |
| T9 — | 180 minutes to 240 minutes after Trendelenburg position to
before arousal from general anesthesia |

Note. IOP = intraocular pressure; T = time point.

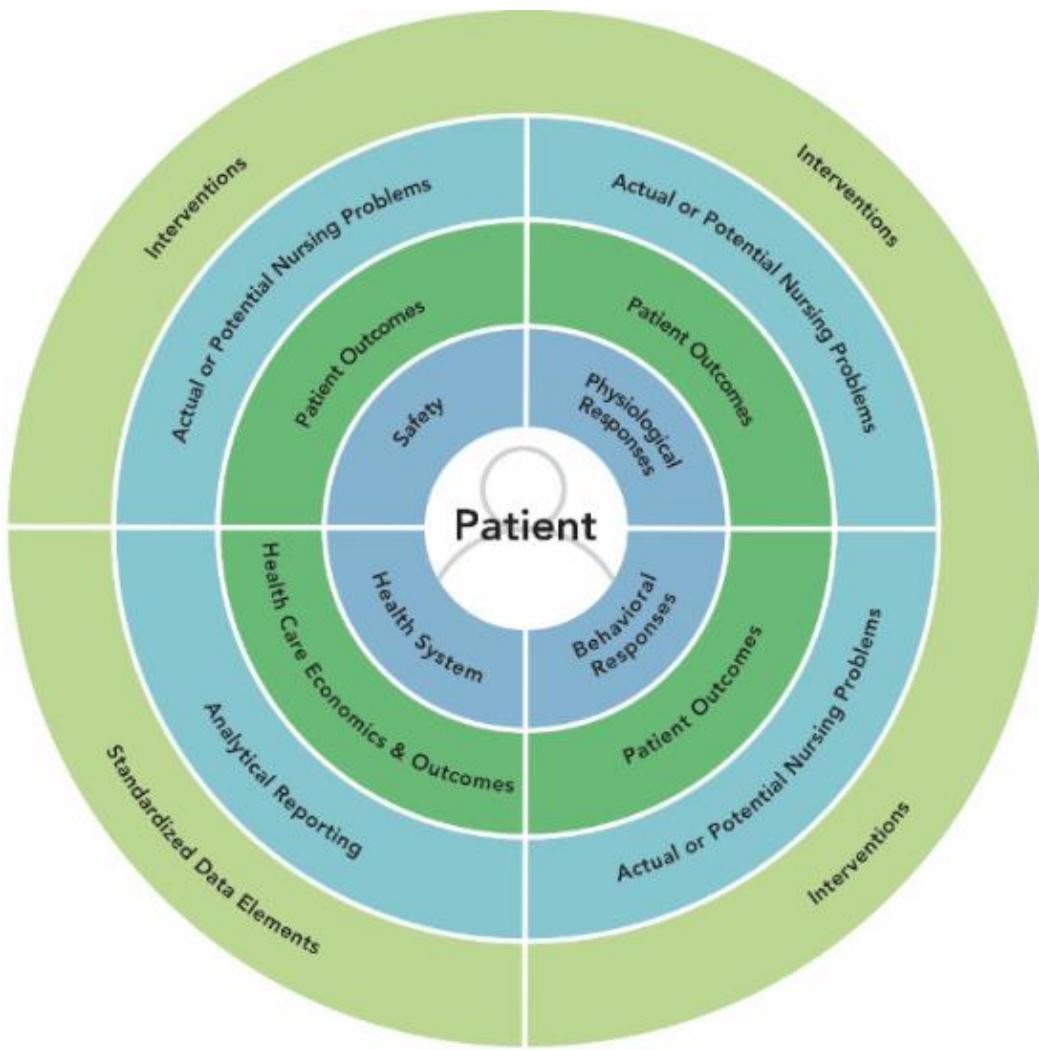


Figure 3.1. Perioperative Patient Focused Model (2017). Reprinted with permission from *Guidelines for Perioperative Practice*. Copyright © 2018, AORN, Inc, 2170 S. Parker Road, Suite 400, Denver, CO 80231. All rights reserved.

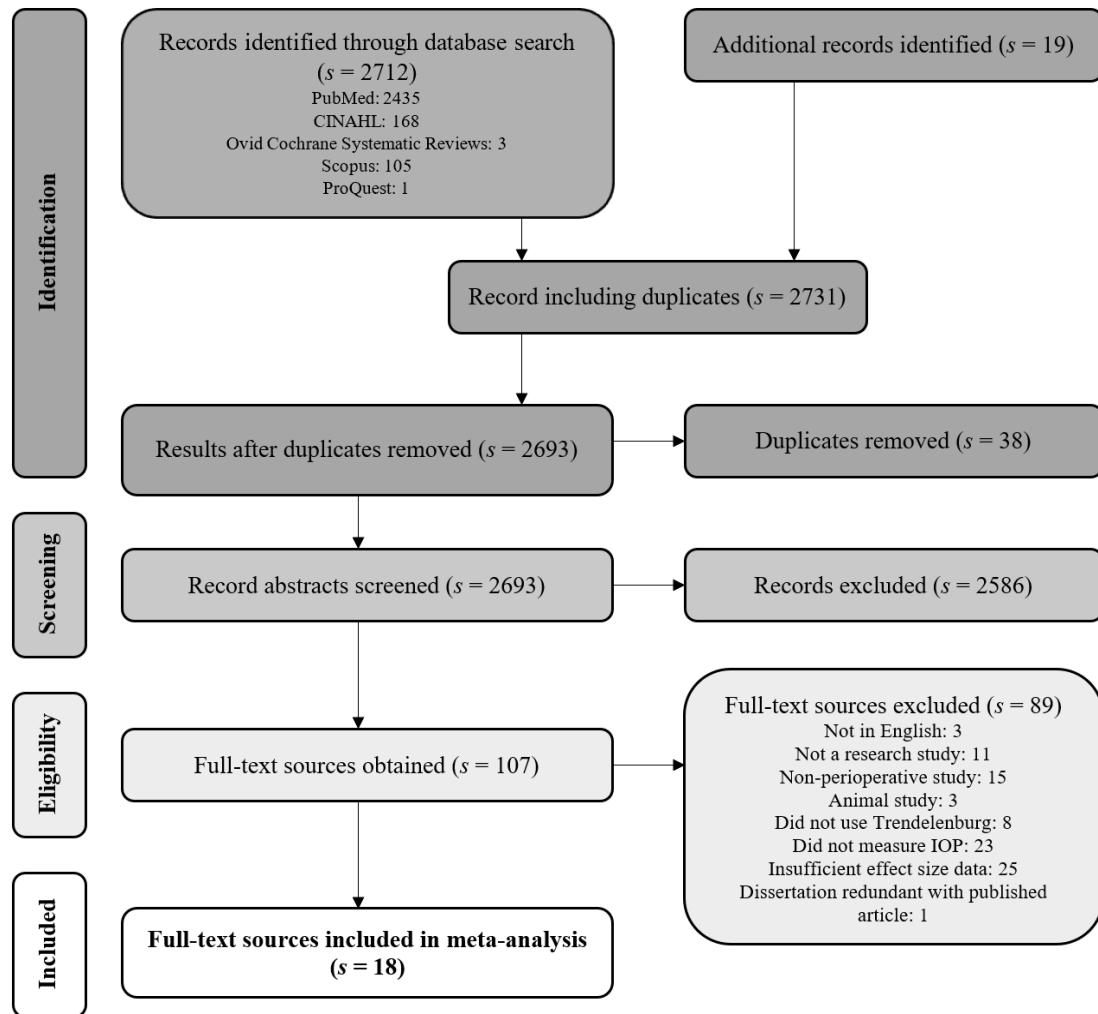


Figure 3.2. Flow diagram of meta-analysis study selection. Note. s = studies. Adapted from Moher D, Liberati A, Tetzlaff J, Atman DG, PRISMA Group (2009). Preferred Reporting Items for Systematic Reviews and Meta-Analyses: The PRISMA Statement.

PLoS Medicine, 6(6), e1000097. doi:10.1371/journal.pmed1000097

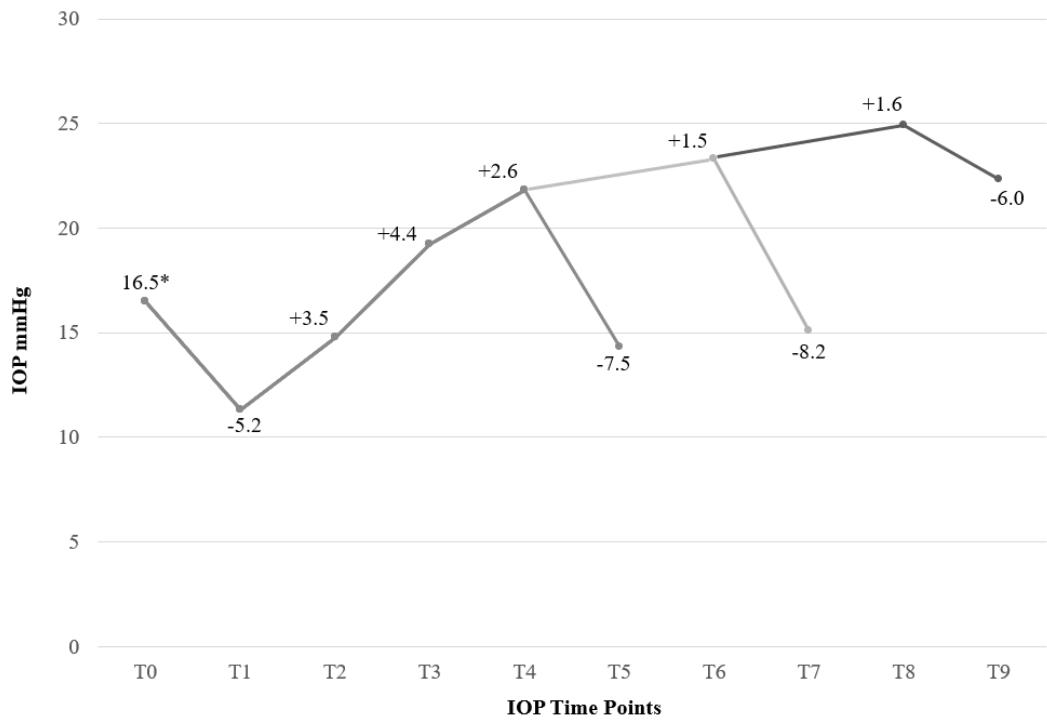
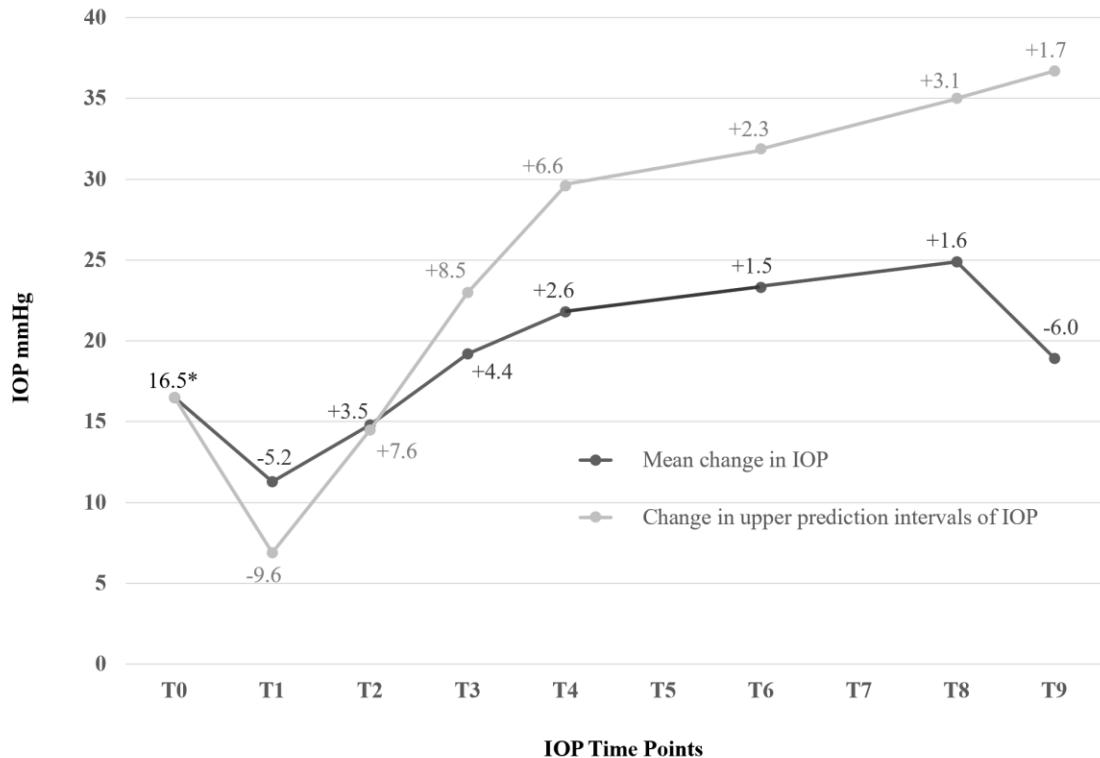


Figure 3.3. Magnitude of change in IOP for T0 through T9 (see Sidebar). Note. IOP = intraocular pressure; mmHg = millimeters of mercury; T = time point. IOP decreases significantly after induction of anesthesia (T1: -5.2 mmHg, $p < 0.001$) and before arousal from anesthesia (T5: -7.5 mmHg, $p < 0.001$; T7: -8.2 mmHg, $p < 0.001$; T9: -6.0 mmHg, $p < 0.001$) when the patient is in the supine position. IOP increases significantly after abdominal insufflation in the supine position (T2: +3.5 mmHg, $p < 0.001$), when the patient is placed in Trendelenburg position (T3: +4.4 mmHg, $p < 0.001$), and with extended time in the Trendelenburg position (T4: +2.6 mmHg, $p < 0.001$; T6: +1.5 mmHg, $p < 0.001$; T8: +1.6 mmHg, $p < 0.001$). *Pooled mean at T0—Before induction of anesthesia.



*Figure 3.4. Magnitude of change in IOP and upper prediction intervals of IOP for T0 through T9 (see Sidebar). Note. IOP = intraocular pressure; mmHg = millimeters of mercury; T = time point. The upper prediction interval shows that after 180 minutes to 240 minutes in the Trendelenburg position, in 95% of all populations, IOP could increase to 35 mmHg (16.5 mmHg - 9.6 mmHg + 7.6 mmHg + 8.5 mmHg + 6.6 mmHg + 2.3 mmHg + 3.1 mmHg = 35 mmHg). *Pooled mean at T0—Before induction of anesthesia.*

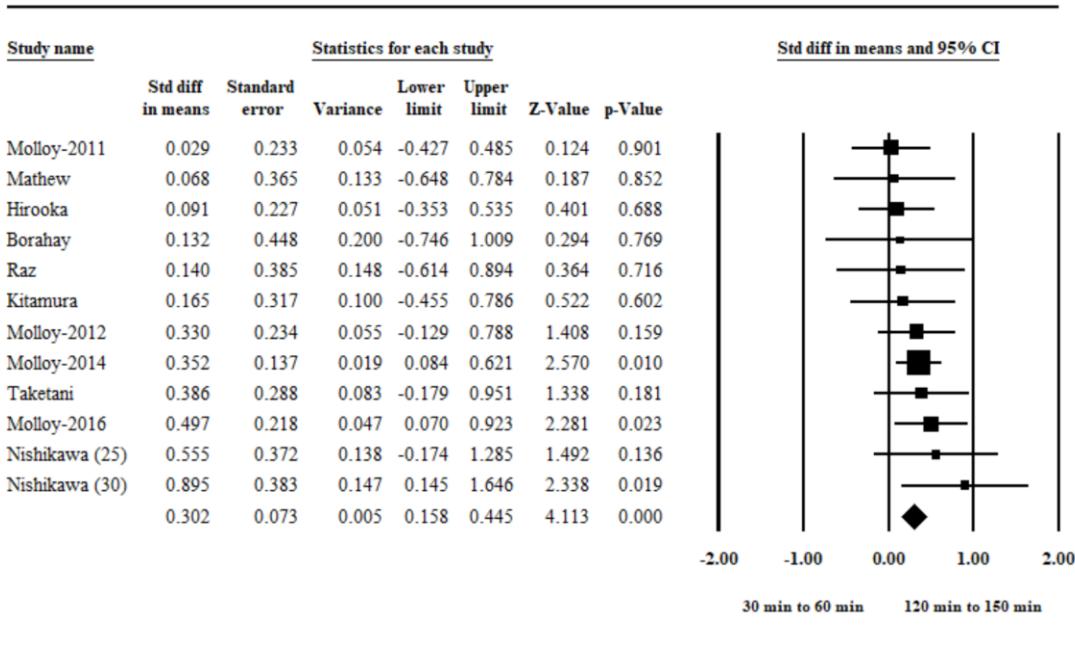


Figure 3.5. Forest plot of meta-analysis of T6. Note. T = time point; Std = Standard; min = minutes. This analysis included 11 studies representing 12 participant groups ($n = 410$). Effect sizes were calculated using a random effects model. The area of each square is proportional to study weight.

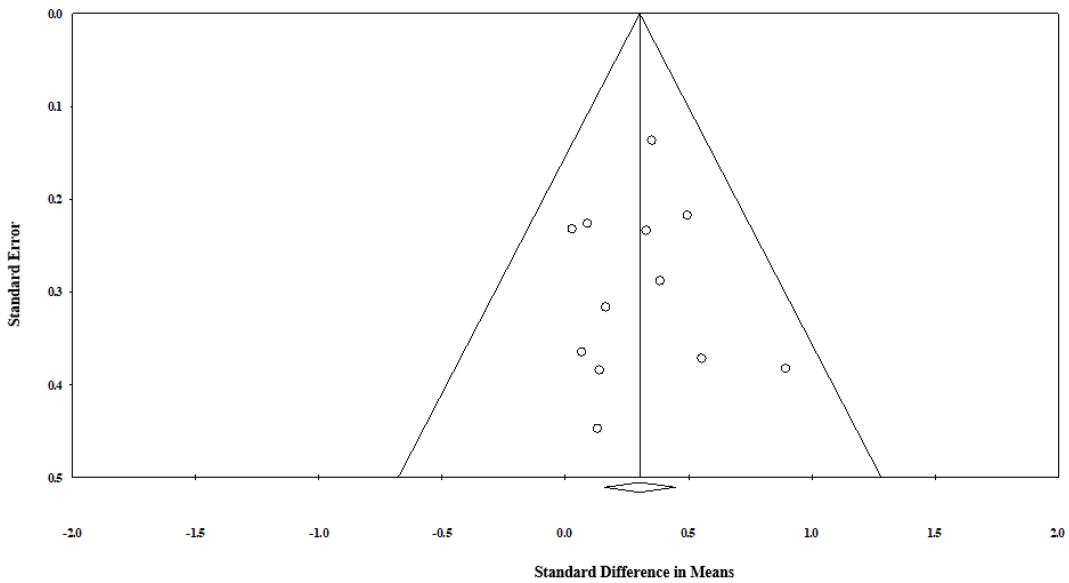


Figure 3.6. Funnel plot of publication bias for T6. Note. T = time point. Larger studies are shown at the top of the funnel. Positive smaller studies are shown at the right of the mean effect size (i.e., center line). The symmetrical distribution of studies (i.e., data points) around the mean effect size indicates a lack of publication bias. The non-significant Egger's regression intercept for this funnel plot (bias = -0.05; $p = 0.47$) is also indicative of the absence of bias in the studies included in the meta-analysis for this time point.

4. SYSTEMATIC REVIEW AND META-ANALYSIS OF PRONE POSITION ON INTRAOCULAR PRESSURE IN ADULT PATIENTS UNDERGOING SURGERY

Van Wicklin, S. A. (2019). *Systematic review and meta-analysis of prone position on intraocular pressure in adult patients undergoing surgery*. Manuscript in preparation.

ABSTRACT

Background. Patients undergoing surgery in the prone position may be at risk for postoperative vision loss associated with increased intraocular pressure. The purpose of this systematic review and meta-analysis is to estimate the magnitude of the increase in intraocular pressure at specific perioperative time points in adult patients.

Methods. Comprehensive search strategies were used to identify nine eligible studies ($N = 229$). Standardized mean difference effect sizes were calculated for two intraoperative time points (T).

Results. Meta-analysis showed that intraocular pressure increased significantly between induction of anesthesia and up to 10 minutes of prone position (T1: $d = 2.5$; $p < 0.001$) and continued to increase significantly until the end of the prone position (T2: $d = 3.4$; $p = 0.002$).

Conclusions. Intraocular pressure increases of this magnitude demonstrate the need for implementing interventions to reduce the risk for postoperative vision loss in patients undergoing surgery in the prone position.

Keywords: intraocular pressure, prone position, Perioperative Patient Focused Model, ischemic optic neuropathy, central retinal artery occlusion.

In the prone position, the patient is positioned face-down on their abdomen. This position provides surgical access to the dorsal aspects of the patient's body. There are ocular complications associated with the prone position. These complications include increased intraocular pressure (IOP; i.e., greater than normal amounts of "pressure exerted by the contents of the eye on its containing wall" [Kamel & Barnette, 2014, p. 432]), chemosis (i.e., conjunctival edema), ocular or orbital hemorrhage, orbital compartment syndrome, and postoperative vision loss (Amorim Correa & Acioly, 2018; Kwee, Ho, & Rozen, 2015; Leibovitch, Casson, Laforest, & Selva, 2006). Postoperative vision loss, which may be partial or complete and unilateral or bilateral, is a serious complication of surgery in the prone position, and it occurs with greater frequency after spine, head and neck, and some orthopedic procedures (Emery et al., 2015; Lee & Newman, 2018). Following surgery in the prone position, there have been reports of postoperative vision loss (Abraham, Sakhija, Sinha, & Rastogi, 2003; Asok, Aziz, Faisal, Tan, & Mallika, 2009; Bekar, Türeyen, & Aksoy, 1996; Dilger et al., 1998; Goni et al., 2012; Grossman & Ward, 1993; Hollenhorst, Svien & Benoit, 1954; Hoski, Eismont, & Green, 1993; Katz, Trobe, Cornblath, & Kline, 1994; Locastro, Novak, & Biglan, 1991; Manfredini, Ferrante, Gildone, & Massari, 2000; Quraishi, Wolinsky, & Gokaslan, 2012; Reddy, Foroozan, Edmond, & Hinckley, 2008; Shifa, Abebe, Bekele, & Habte, 2016; Stang-Veldhouse, Yeu, Rothenberg, & Mizen, 2010; West, Askin, Clarke, & Vernon, 1990; Wolfe, Lospinuso, & Burke, 1992), as well as reports of subconjunctival hemorrhage (Akhaddar & Boucetta, 2012), subperiosteal orbital hemorrhage (Russell & Dutton, 2011), orbital compartment syndrome (Amorim Correa & Acioly, 2018;

Leibovitch et al., 2006), acute angle-closure glaucoma (Singer & Salim, 2010; Stewart, Landy, & Lee, 2016), and Horner syndrome (Guillaume & Gowreesunker, 2013).

When the patient is in the prone position, IOP increases and the extent of this increase is related to the amount of time the patient is in the prone position (Agah, Ghasemi, Roodneshin, Radpay, & Moradian, 2011; Eddama, 2013; Kamel & Barnette, 2014; Kendrick, 2012; Pinkney et al., 2012; Szmuk et al., 2013; Yoshimura, Hayashi, Tanaka, Nomura, & Kawaguchi, 2015). After only a few minutes in the prone position, IOP can increase significantly (Nuzzi & Tridico, 2015). According to the American Academy of Ophthalmology (2018), normal IOP is 10 millimeters of mercury (mmHg) to 21 mmHg. Intraocular pressures higher than 21 mmHg pose a risk for glaucoma, detached retina, and postoperative vision loss (Emery et al., 2015; Lee & Newman, 2018; Weinreb & Khaw, 2004).

Pathogenesis of Postoperative Vision Loss

Postoperative vision loss in patients undergoing surgery in the prone position is generally related to one of two causes: ischemic optic neuropathy or central retinal artery occlusion (Kamel & Barnette, 2014; Stambough, Dolan, Werner, & Godfrey, 2007).

Ischemic Optic Neuropathy

Ischemic optic neuropathy is the most common cause of postoperative vision loss (Kamel & Barnette, 2014; Kan, Brown, & Gainsburg, 2015; Newman, 2008; Stambough et al., 2007). According to the American Academy of Ophthalmology (Boyd, 2018), ischemic optic neuropathy is caused by insufficient blood flow to the optic nerve. Ischemic optic neuropathy may present as anterior (involving ischemia and infarction of the intraocular optic nerve) or posterior (involving ischemia and infarction of the

intraorbital optic nerve; Kamel & Barnette, 2014). Although the phenomenon of ischemic optic neuropathy is not well understood, it is known that elevated IOP can lead to optic nerve injury and decreased ocular perfusion pressure (Kamel & Barnette, 2014; Kan et al., 2015; Newman, 2008). Perfusion pressure is the difference between pressures of the ciliary arteries in the optic nerve and the venous drainage of the eye (Hayreh, 2001). This difference is approximated by the level of IOP. The “higher the intraocular pressure, the lower the perfusion pressure, and consequently, the lower the blood flow in the optic nerve head” (Hayreh, 2001, p. 608). The lower the blood flow in the optic nerve head, the greater the risk for ischemic optic neuropathy and postoperative vision loss. The ischemic process can occur as a direct result of decreased blood supply from the arteries of the optic nerve or by venous stasis that occurs as a result of decreased venous outflow and a compartment syndrome of the optic nerve or optic canal (Gkegkes, Karydis, Tyritzis, & Iavazzo, 2015; Kamel & Barnette, 2014; Kan et al., 2015). The subsequent postoperative vision loss can range from temporary blurring to partial to complete blindness; however, once a loss of vision occurs, it is almost always an irreversible complication (Emery et al., 2015; Lee & Newman, 2018).

Central Retinal Artery Occlusion

Central retinal artery occlusion is most often caused by pressure on the eye from the prone position, and especially by positioning the patient’s head on a prominent headrest. Pressure on the eye increases IOP and decreases blood flow to the retina through the central retinal artery (Kamel & Barnette, 2014; Li, Swinney, Veeravagu, Bhatti, & Ratliff, 2015; Stambough et al., 2007). The increased IOP exceeds the profusion pressure of the central retinal artery, leading to ischemia of the retina

(Stambough et al., 2007). Most patients with central retinal artery occlusion are left with a unilateral, permanent blindness (Li et al., 2015; Stambough et al., 2007).

Risk Factors for Postoperative Vision Loss

Numerous risk factors for postoperative vision loss have been identified that include older patients with elevated baseline IOPs, patients with existing hypertension, diabetes, obesity, anemia, vascular disease, increased blood viscosity, and patients who smoke, as well as patients who experience intraoperative hypotension, blood transfusion, lower colloid use during fluid administration, or prolonged surgical times, and patients who are positioned on horseshoe-shaped headrests (Freshcoln & Diehl, 2014; Ghomi, 2012; Kamel & Barnette, 2014; Stambough et al., 2007). Patients who are predisposed to acute angle-closure glaucoma are also at high risk for ocular injury even during short procedures because the prone position can shift the lens-iris diaphragm forward so it obstructs aqueous humor outflow and increases IOP (Kwee et al., 2015).

Incidence of Postoperative Vision Loss

The exact incidence of postoperative vision loss is unknown because the data come largely from retrospective studies and case reports (Lee & Newman, 2018; Patil, Lad, Lad, Ho, & Boakye, 2008). The incidence of postoperative vision loss following nonocular surgery has been estimated to be as low as 0.0002% and as high as 0.2% (Berg, Harrison, & Lee, 2010; Newman, 2008). In a retrospective cohort study using the National Inpatient Sample, the largest inpatient database in the United States, Patil et al., (2008), examined the records of 4,728,815 patients who underwent spinal procedures between 1993 and 2002. The researchers found that 4134 patients (0.09%) developed postoperative visual impairment. An additional 271 patients (0.006%) had ischemic optic

neuropathy and an additional 47 patients (0.001%) had central retinal artery occlusion. In a 10-year prevalence study of postoperative vision loss in the United States, Shen, Drum, and Roth (2009) reviewed data from more than 5.6 million patients included in the Nationwide Inpatient Sample from 1995 to 2005. The researchers found the incidence of postoperative vision loss for patients undergoing laminectomy without fusion was 0.86 per 10,000 (0.009%). For patients undergoing spinal fusion, the incidence was 3.09 per 10,000 (0.03%) for all fusions, 0.66 per 10,000 (0.007%) fusions with anterior approach, and 5.50 per 10,000 (0.06%) fusions with posterior approach.

Conceptual Framework

The Perioperative Patient Focused Model (Figure 4.1; Rothrock & Smith, 2000) provides a conceptual framework for this systematic review and meta-analyses. During operative procedures when the patient is anesthetized, the perioperative registered nurse (RN) fills an important role as the patient's advocate and also oversees the patient's perioperative care. Secondary to being patient focused, the Perioperative Patient Focused Model is outcome focused (Rothrock & Smith, 2000). The care provided by perioperative RNs is directed toward achieving high quality patient outcomes, such as preventing postoperative vision loss associated with increased IOP. Providing optimal perioperative patient care, serving as the patient's advocate, and achieving high quality outcomes requires specific perioperative knowledge and skillful implementation of best practices for patient positioning. Consistent with the Perioperative Patient Focused Model, the evidence-based practices implemented by perioperative RNs during patient positioning promote patient safety, and optimal physiological and behavioral responses in perioperative patients. Providing evidence about the magnitude of the increase in IOP

resulting from the intraoperative use of the prone position will support the implementation of nursing interventions to help prevent postoperative vision loss and other ocular complications in adult patients undergoing surgery in the prone position.

Purpose

Although some researchers have studied the quantitative increase of IOP in surgical patients in the prone position (Emery et al., 2015; Lee & Newman, 2018; Nuri Deniz et al., 2013; Quraishi et al., 2012; Shifa et al., 2016; Stang-Veldhouse et al., 2010; Yoshimura et al., 2015), there is a need for systematic review and meta-analyses of these studies to demonstrate the overall effect size and provide high-quality evidence supporting the need for implementing interventions to mitigate the increase of IOP and reduce the risk for postoperative vision loss. Meta-analysis research methods provide increased power compared to individual studies, improve estimates of effect size, and help resolve uncertainty when the results of individual studies disagree (Berlin, 1995). Because all of the evidence pertaining to a particular phenomenon is included in the analysis, meta-analysis research provides a high level of objectivity, precision, and generalizability (Thompson, 1994).

Currently, there has been no quantitative meta-analytic synthesis of the existing studies examining the increase in IOP in adult patients undergoing surgery in the prone position. The purpose of this systematic review and meta-analysis is to estimate the magnitude of the increase in IOP at selected perioperative time points in adult patients (i.e., individuals 18 years and older) undergoing any type of surgery in the prone position. Consequently, the research question to be addressed by this systematic review and meta-

analysis is, “What is the magnitude of the increase in IOP at specific perioperative time points in adults undergoing surgery in the prone position?”

Methods

To ensure rigorous and transparent presentation of the methods and results of this systematic review and meta-analysis, the Preferred Reporting Items for Systematic reviews and Meta-Analyses (PRISMA) guidelines (Moher, Liberati, Tetzlaff, Altman, & PRISMA Group, 2009) have been followed.

Search Strategies

An expert health sciences reference librarian was consulted to identify the most appropriate search terms and databases for an exhaustive and varied literature search. Keywords or medical subject headings (MeSH) included *intraocular pressure* or *ocular tension*, and *prone position*. Search strategies included

- online searching of the PubMed, CINAHL, and Scopus databases, and the Cochrane Database of Systematic Reviews for published and unpublished literature;
- ancestry searching of reference lists from relevant reports to locate additional applicable references;
- author searching of individuals identified in the literature as experts in the field; and a
- dissertation search of the ProQuest database.

The author reviewed report abstracts for eligibility and obtained full-text copies of all potentially eligible reports.

Inclusion and Exclusion Criteria

Criteria for inclusion in the meta-analysis were reports written in English; published or unpublished reports of primary studies that encompassed dissertations,

conference abstracts, and presentations; studies that used either a one-group, pretest posttest design or a multiple-group, pretest posttest design; reports where the minimum age of the study participants was 18-years or older; studies that included a specific measured outcome of IOP using any type of tonometer; studies where the participants received any type of general or spinal anesthesia; and reports of studies that included sufficient data to calculate an effect size. When reports did not include sufficient data to calculate an effect size, the author contacted the researchers and statisticians on at least two separate occasions two to three weeks apart to obtain missing data.

Reports were excluded if the IOP was measured in adults not undergoing surgery. Reports were also excluded if data from only one time point of IOP measurement were provided. Participant groups were additionally excluded if they were receiving an intervention specifically intended to mitigate IOP; however, participant groups representing control arms receiving placebos or no interventions were included in the systematic review and meta-analysis.

Risk of Bias Within Individual Studies

Each report included in the systematic review and meta-analysis was independently appraised and critically evaluated for its level of strength and quality by the author and an experienced evidence reviewer to assess the risk of bias within individual studies. The Association of periOperative Registered Nurses (AORN) Research Evidence Appraisal Tool – Study, available on the AORN website (<https://www.aorn.org/guidelines/about-aorn-guidelines/evidence-rating>) was used for this purpose. When using the AORN appraisal tool, non-experimental designs (e.g., observational studies) are assigned the lowest level of strength, quasi-experimental

designs are assigned a moderate level of strength, and strong study designs (e.g., randomized controlled trials) are assigned the highest level of strength (i.e., experimental). An assessment of measures such as sample size, generalizability, bias, reliability, and validity is conducted to determine whether study quality is high, good, or low. The author and evidence reviewer achieved 100 percent consensus on study design and quality levels for each of the included studies by discussing their independent appraisals during conference calls.

Risk of Bias Across Studies

Several strategies were used to manage risk of bias that may affect cumulative evidence across studies (Valentine, 2009). To avoid bias due to a narrow or limited search, a comprehensive and diverse literature search was conducted (White, 2009). Only research studies were included in the systematic review and meta-analysis to ensure sufficient strength and quality of the included studies. An analysis of publication bias was conducted to determine whether unpublished research was unintentionally excluded. As well, only studies with objective measurements of IOP at two or more perioperative time points were included.

Coding and Data Extraction

The author used an iterative process that included reviewing the literature and studying codebooks used by experienced meta-analysts to develop a codebook for data extraction and coding of included studies. Based on content analysis of studies specific to the effects of prone position on IOP, the codebook was revised. The codebook was pilot-tested by the author using five randomly selected studies to identify missed coding

categories and verify fit between coding categories and study characteristics before it was used to code and extract data from all eligible reports (Wilson, 2009).

To address the research question and statistically analyze the effects and methodologies of each of the reports, the author coded eligible studies and extracted necessary data. Effect size data for each of the reports included in the systematic review and meta-analysis was independently coded by a trained researcher. Coding discrepancies on effect size data for each of the eligible studies were discussed by the author and independent researcher until 100 percent consensus was achieved.

Data related to study characteristics (i.e., authors, year of publication, publication status, geographic location, reported funding) and data related to study design (i.e., type of study, study quality, type of tonometer, type of head positioning device, inclusion of ophthalmologic exams by participants) were collected from each of the included studies. Data related to participant and surgery characteristics (i.e., age; gender; American Society of Anesthesiologists [ASA; 2018] physical status classification; body mass index [BMI]; type of surgery; type of anesthesia; duration of anesthesia, prone position, and surgery; estimated blood loss [EBL]) were also collected when available. The researchers extracted data necessary to calculate effect sizes for all time points recorded during the perioperative phases of the procedures.

Analyses

Meta-analyses were conducted using *Comprehensive Meta-Analysis Software*, Version 3 (Borenstein, Hedges, Higgins, & Rothstein, 2018). To allow for similarity with the order of events as they traditionally occur during surgery and to achieve the greatest number of comparisons for analysis at each time point, the time points for meta-analysis

were selected from the time points recorded by the researchers. Analyses were conducted for the following time points (T):

- T1 After induction of anesthesia to 0 minutes to 10 minutes of prone position, and
- T2 0 minutes to 10 minutes of prone position to the end of prone position.

Standardized mean difference effect sizes were calculated for each participant group and each measured time point (Cohen, 1992).

Effect size values were weighted by the inverse of the variance to account for sample size and adjust for bias. Because heterogeneity was observed among study designs, sample attributes, and outcome measures, and to account for between- and within-study variation, a random effects model was selected a priori to synthesize effect sizes. Using a random effects model assumes that the true effect size varies from one study to the next (Borenstein, Hedges, Higgins, & Rothstein, 2009a, p. 77). Effect sizes were interpreted as per Cohen (1992) with 0.2 considered a small effect size, 0.5 considered a medium effect size, and 0.8 or greater considered a large effect size. The calculated effect sizes were converted to the metric used to measure IOP (i.e., mmHg) using the procedures described by Lipsey and Wilson (2001) to facilitate interpretation of effect size findings.

Publication bias. A funnel plot was constructed to assess the extent of publication bias for the meta-analysis. Notably, a funnel plot may suggest publication bias, but does not eliminate the bias (Sutton, 2009). A minimum of three studies is necessary in order to create a funnel plot (Borenstein et al., 2018). Asymmetry of the funnel plot was measured by conducting an Egger's test (Borenstein et al., 2018; Egger, Smith, Schneider, & Minder, 1997; Sterne Egger, & Moher, 2011). When there are less

than 10 studies included in the meta-analysis an Egger's test is not advised because the power of the test may be too low to distinguish true asymmetry from chance (Sterne et al., 2011). For this reason, a funnel plot was constructed and an Egger's test was conducted only when the analysis included 10 or more participant groups (T1). Orwin's *Fail-safe N* test was conducted when the Egger's test was significant, to estimate the number of missing studies required to overturn the conclusions of the meta-analysis (Orwin, 1983; Sutton, 2009). Orwin's *Fail-safe N* assesses the impact of publication bias to determine whether the overall observed effect is robust (Orwin, 1983; Sutton, 2009).

Heterogeneity. The studies included in the meta-analysis were assessed for heterogeneity after deciding on the model and calculating effect sizes. Heterogeneity is any kind of variability among the studies included in a meta-analysis, according to the Cochrane Collaboration (Deeks, Higgins, & Altman, 2011). Heterogeneity testing explores whether the same effect is being evaluated by all studies (i.e., the null hypothesis); Higgins, Thompson, Deeks, & Altman, 2003). Cochrane's Q, which estimates statistical significance; Tau-squared (T^2), which estimates the absolute value of the true variance between studies, but not the proportion of the variance; and I^2 -squared (I^2), which estimates the proportion of true variance, but not the absolute value of the variance, were all used to test homogeneity of variance among effect sizes (Borenstein, Hedges, Higgins, & Rothstein, 2009b). Using an I^2 test for assessing the impact of heterogeneity in a meta-analysis is recommended by Higgins and Thompson (2002) as this test quantifies the influence (as opposed to the amount) of heterogeneity and expresses the percentage of variability due to heterogeneity rather than chance (p. 1553). An I^2 value of 0% indicates there is no heterogeneity, values of 25% reflect low observed

variation, values of 50% reflect moderate levels, and values of 75%, reflect high levels of observed variation (Higgins et al., 2003). To show the dispersion of true effect sizes around the mean, prediction intervals for each time point were also calculated (Borenstein, Hedges, Higgins, & Rothstein, 2009c).

Results

The flow of study selection is depicted in Figure 4.2. In total, 135 records were identified for possible inclusion, and of these, nine studies were included in the systematic review and meta-analysis. Four studies had multiple participant groups (Czorlick et al., 2018; Hunt et al., 2004; Pinar et al., 2018; Sugata et al., 2012), resulting in a total of 14 participant groups and 229 participants for analysis. Table 4.1 contains a summary of the studies included in this review and meta-analysis.

Study Characteristics

All of the studies included in the systematic review and meta-analysis were obtained from peer-reviewed journals. The researchers of three studies (33.3%) reported receiving some type of funding or donated supplies (Emery et al., 2015; Pinar et al., 2018; Sugata et al., 2012). Although the literature was searched without any date restriction, all of the studies included in the systematic review and meta-analysis were published between 2001 and 2018. The greatest number of studies (s) were published in 2018 ($s = 2$) with the greatest number of participants also occurring in 2018 ($n = 88$). The majority ($s = 7$) were conducted in Asia ($s = 4$) or North America ($s = 3$), with the majority of participants from Asia ($n = 107$).

Participant Characteristics

Participant and surgery characteristics are shown in Table 4.2. The mean age of the participants was 53.2 years (± 8.1) and ranged from 43.3 years to 69 years. The majority of the participants were men ($n = 122$; 55.0%). Participant race and socioeconomic status were not reported by any of the researchers. Participants ranged between having a healthy weight to being slightly overweight (22.5 kg/m^2 to 27.7 kg/m^2) with a mean BMI of 24.9 kg/m^2 (± 1.6 ; Centers for Disease Control and Prevention, 2016). Czorlich et al., (2018) excluded patients with a BMI greater than 30 kg/m^2 . The ASA (2018) physical classification status was reported cumulatively as class I (i.e., healthy), class II (i.e., with mild systemic disease), or class III (i.e., with severe systemic disease) by the researchers of four reports ($n = 130$; Agah et al., 2011; Cheng et al., 2001; Nuri Deniz et al., 2013; Pinar et al., 2018). Sugata et al., (2012) reported patient comorbidities for diabetes ($n = 5$ of 24; 20.8%), and hypertension ($n = 9$ of 24; 37.5%).

Surgery Characteristics

The majority of the participants underwent spine surgery ($n = 170$; 74.2%). Other participants underwent percutaneous nephrolithotomy ($n = 43$; 18.8%) or cranial surgery ($n = 16$; 7%). The vast majority of participants ($n = 222$; 97%) received general anesthesia by either inhalation ($n = 115$; 51.8%), intravenous propofol ($n = 60$; 27%), or unspecified methods ($n = 27$; 12.2%), or received spinal anesthesia ($n = 20$; 9%). As shown in Table 4.2, the mean duration of surgery was 156.0 minutes (± 24.2) with a range of 120 minutes to 181 minutes. The mean EBL was 330.1 milliliters (mL; ± 222.1) with a range of 120 mL to 615 mL.

Study Design Characteristics

The nine reports included in this systematic review and meta-analysis comprised four experimental ($n = 97$; Carey, Shaw, Weber, & DeVine, 2014; Emery et al., 2015; Nuri Deniz et al., 2013; Pinar et al., 2018), and five non-experimental ($n = 132$; Agah et al., 2011; Cheng et al., 2001; Czorlich et al., 2018; Hunt et al., 2004; Sugata et al., 2012). Five were high quality ($n = 146$; Carey et al., 2014; Czorlich et al., 2018; Emery et al., 2015; Pinar et al., 2018; Sugata et al., 2012) and four were good quality ($n = 83$; Agah et al., 2011; Cheng et al., 2001; Hunt et al., 2004; Nuri Deniz et al., 2013). The researchers used three different tonometers to measure IOP; however, the Tono-Pen XL was used most frequently ($s = 7$; $n = 166$). The most frequently used method for positioning the patient's head was skull pins or clamps ($n = 105$; 45.9%). Other methods included a horseshoe-shaped headrest ($n = 71$; 31%), a pillow or viscoelastic gel ring-shaped headrest ($n = 30$; 13.1%), or a silicone headrest ($n = 23$; 10%). Two researchers reported having 88 participants undergo preoperative ophthalmologic examinations (Czorlich et al., 2018; Pinar et al., 2018). One researcher reported having 48 participants undergo postoperative ophthalmologic examinations (Czorlich et al., 2018).

Effect Sizes

Results of the meta-analysis for each time point are shown in Table 4.3. A graphical representation of the magnitude of changes in IOP and upper prediction intervals for T0 through T2 is shown in Figure 4.3. In total, between induction of anesthesia and the end of prone position, IOP increases significantly by 17.6 mmHg (i.e., $7.5 \text{ mmHg} + 10.1 \text{ mmHg} = 17.6 \text{ mmHg}$). Based on the upper limits of the prediction intervals, in 95% of all populations, IOP could increase by as much as 57.8 mmHg (i.e., $19.8 \text{ mmHg} + 38.0 \text{ mmHg} = 57.8 \text{ mmHg}$). The IOP increases significantly after the

patient is placed into the prone position (T1: +7.5 mmHg, $p < 0.001$) and continues to increase significantly while the patient is in the prone position (T2: +10.1 mmHg, $p = 0.002$). The forest plots of effect sizes for each participant group included in the meta-analysis for T1 and T2 are shown in Figure 4.4 and Figure 4.5. The funnel plot for publication bias for T1 is shown in Figure 4.6.

Discussion

The results of this systematic review and analysis have shown that IOP increases significantly for adult patients undergoing surgery in the prone position. As shown in Figure 4.3, if an individual had a baseline IOP of 13.3 mmHg after induction of anesthesia (as indicated by the pooled mean calculated for T0), by the end of prone position, IOP could increase to 30.9 mmHg ($13.3\text{mmHg} + 7.5\text{ mmHg} + 10.1\text{ mmHg} = 30.9\text{ mmHg}$). Based on the upper limits of the prediction intervals, IOP could increase to 71.1 mmHg ($13.3\text{ mmHg} + 19.8\text{ mmHg} + 38.0\text{ mmHg} = 71.1\text{ mmHg}$). An IOP of 71.1 mmHg is more than three times the highest parameter of normal IOP (i.e., 21 mmHg).

The mean IOP for the general population has been reported as $15.5 (\pm 2.5)$ mmHg (Carey et al., 2014). An IOP of 23 mmHg is three standard deviations above the mean (i.e., $15.5\text{ mmHg} + 2.5\text{ mmHg} + 2.5\text{ mmHg} + 2.5\text{ mmHg} = 23\text{ mmHg}$) and has thus been considered as a marker of abnormally elevated IOP (Carey et al., 2014). Yoshimura et al., (2015) conducted a study to evaluate predictive factors associated with increased IOP during spine surgery in the prone position. The researchers found that an IOP of 23 mmHg or greater was predictive of an IOP of 30 mmHg or greater. Riva, Sinclair, and Grunwald (1981) found that the highest IOP at which the retina was able to maintain a constant blood flow was $29.6 (\pm 2)$ mmHg. Pillunat, Anderson, Knighton, Joos, and

Feuer (1997), found that blood flow to the optic nerve remained nearly constant until IOP reached 40 mmHg. The researchers noted; however, that some individuals do not exhibit autoregulation, and even a very modest increase in IOP can lead to a decline in blood flow to the optic nerve.

Implications for Practice

Increased IOP puts the patient at risk for glaucoma, detached retina, or partial to complete vision loss (Amorim Correa & Acioly, 2018; Kwee et al., 2015; Leibovitch et al., 2006). Intraocular pressure increases of the magnitude found in this systematic review and meta-analysis clearly demonstrate the need for implementing intraoperative interventions to mitigate the increase in IOP and reduce the potential for serious ocular complications in patients undergoing surgery in the prone position.

Positioning the patient in a 5-degree to 10-degree reverse Trendelenburg prone position may be a simple intervention to prevent some instances of postoperative vision loss. This position has been shown to decrease IOP in healthy volunteers (Ozcan et al., 2004; Walick, Kragh, Ward, & Crawford, 2007) and in patients undergoing spine surgery (Carey et al., 2014; Emery et al., 2015; Fukui, Ahmad, McHugh, Tempelhoff, & Cheng, 2004; Fukui, Tempelhoff, & Cheng, 2005). Positioning surgical patients with the head above the heart helps reduce venous congestion in the eye and orbit and decrease intraocular and intraorbital pressure (Bonnaig, Dailey, & Archdeacon, 2014; Carey et al., 2014; Grant et al., 2010; Kamel, & Barnette, 2014; Nickels, Manlapaz, & Farag, 2014). Reducing the length of time the patient is in the prone position may also help to mitigate the increase in IOP. The ASA Task Force on Perioperative Visual Loss, North American Neuro-Ophthalmology Society, and Society for Neuroscience in Anesthesiology and

Critical Care (2019) suggest staging procedures when patients will be in the prone position for prolonged periods of time. Utilizing a series of shorter procedures rather than one prolonged procedure may help reduce the patient's risk for postoperative vision loss; however, the risks associated with multiple surgeries may outweigh the benefits of staged procedures (Shifa et al., 2016).

Because IOP increases during prone position, intraoperative monitoring of IOP either continuously or at established intervals or time points (e.g., after initiation of prone position, after every 60 minutes of prone position) seems prudent. Yoshimura et al., (2015) suggested that measuring IOPs after one hour of surgery in the prone position could provide an opportunity for implementing interventions to prevent additional increases in IOP. Eddama (2013) also suggested that regular measurement of IOP during prolonged surgery provided an opportunity for implementing a change in the patient's position when critical thresholds are reached.

Implementing periodic intraoperative position changes or rest periods (where the ocular level is above the heart) can help to reduce IOP. In a quasi-experimental study, Molloy and Watson (2012) implemented a five-to-seven-minute level supine intervention after 60 minutes of steep Trendelenburg position and found there was a significant decrease in IOP.

When the patient is in the prone position, there is a risk for direct compression on the eye (Bonnaig et al., 2014). Yu, Chou, Yang, and Chang (2010) found that the prone position was a precipitating factor for eye injury. Preventing direct pressure and assessing and monitoring the patient's eyes at regular intervals throughout the procedure may help to reduce the incidence of postoperative vision loss (ASA Task Force on Perioperative

Visual Loss et al., 2019; Locastro et al., 1991; Nickels et al., 2014; Shifa et al., 2016). Avoiding specific headrests or positioning devices that may increase pressure on the eye (e.g., horseshoe-shaped, Wilson frame) or using skull pins or tongs to position the head may also help to prevent pressure on the orbits and reduce the risk for postoperative vision loss (Asok et al., 2009; Bekar et al., 1996; Grossman & Ward, 1993; Hollenhorst et al., 1954; Hoski et al., 1993; Quraishi et al., 2012; Wolfe et al., 1992). Direct compression from a horseshoe-shaped head positioner has been reported as a cause of postoperative vision loss when the patient is in the prone position (Abraham et al., 2003; Bekar et al., 1996; Grossman & Ward, 1993; Hollenhorst et al., 1954; Hoski et al., 1993; Locastro et al., 1991; Wolfe et al., 1992). In a case-controlled study of 80 patients with ischemic optic neuropathy compared with 315 matched control patients, the ASA Postoperative Visual Loss Study Group (2012) found that Wilson frame use was an independent risk factor for postoperative vision loss.

Administering specific medications or anesthetics may also be effective in reducing IOP or mitigating the intraoperative increase in IOP (Farag et al., 2012; Pinar et al., 2018; Sugata et al., 2012). Pinar et al., (2018) found that the increase in IOP was significantly less in patients undergoing lumbar disc surgery in the prone position under spinal anesthesia compared with patients receiving general anesthesia. Sugata et al., (2012) found that IOPs were higher in patients undergoing prone spine surgery with general anesthesia maintained with sevoflurane compared with patients receiving general anesthesia maintained with intravenous propofol. Farag et al., (2012) found that the administration of topical brimonidine 2% helped reduce intraoperative IOP.

Another important consideration for practice is the need to evaluate whether patients undergoing surgery in the prone position should receive a preoperative ophthalmologic examination to reduce the risk for ocular injury (Akhaddar & Boucetta, 2012; ASA Task Force on Perioperative Visual Loss et al., 2019; Singer & Salim, 2010; Stang-Veldhouse et al., 2010; Stewart et al., 2016). Preoperative ophthalmologic examinations may be helpful in identifying patients at risk for postoperative vision loss. Increases in IOP may be more harmful in older patients, patients with risk factors for postoperative vision loss, or in patients who are predisposed to developing diabetes or glaucoma than in younger, healthier patients (Akhaddar & Boucetta, 2012; ASA Task Force on Perioperative Visual Loss et al., 2019; Singer & Salim, 2010; Stang-Veldhouse et al., 2010; Stewart et al., 2016). Patients at risk for acute angle-closure glaucoma associated with the prone position may benefit from preoperative laser iridotomy (Singer & Salim, 2010; Stewart et al., 2016). The ASA Task Force on Perioperative Visual Loss et al., (2019) recommend evaluating the need for preoperative ophthalmologic examination on a case-by-case basis.

The results of this systematic review and meta-analysis support the use of the Perioperative Patient Focused Model as the conceptual foundation for this perioperative research. Providing quantitative evidence about the magnitude of the intraoperative increase in IOP resulting from the prone position supports the implementation of nursing interventions that are patient-focused and will improve patient outcomes by mitigating increases in IOP and reducing the risk for permanent postoperative vision loss and other ocular complications in adult surgical patients.

Implications for Future Research

Further research relative to the magnitude of IOP increases in patients undergoing surgery in the prone position is warranted. Further research to provide validation and demonstrate reliability of the Perioperative Patient Focused Model is also warranted. To allow for consistent data collection, comparison, meta-analysis, and reporting, researchers of future studies should use standardized time points for measurement (i.e., before induction, after induction, after change to prone position and every 30 minutes to 60 minutes thereafter, after return to supine position, before arousal, and postoperatively). Further, researchers should present data in a consistent format for each time point (i.e., sample size, mean, standard deviation). To determine whether certain variables affect the strength of the relationship between prone position and IOP, researchers should study patients of all ages and ethnicities, without restriction of BMI or comorbidities.

Limitations

This systematic review and meta-analyses has several limitations. The number of included studies and participants is small ($s = 9$; $N = 229$). As shown in Figure 4.2, studies were excluded for a variety of reasons; however, five studies were excluded from the analyses solely because of a lack of data necessary to calculate an effect size. The researchers were contacted a minimum of two times to obtain missing data, but none responded. One researcher excluded participants based on BMI (Czorlich et al., 2018); therefore, the mean value for this variable may not fully reflect the true characteristics of all adult surgical patients.

The meta-analysis examined only two intraoperative time points. The analysis for T2 reflects IOP measurements from patients in the prone position for varying lengths of time. Having data for specific time points would be preferable. This lack of data

collection by researchers is likely due to the difficulty of measuring IOP with the patient in prone position. Because researchers measured IOP at varying intraoperative time points, all studies could not be included at all time points examined in the meta-analysis. Additionally, there were not enough studies included at each time point to allow for moderator analyses.

Heterogeneity was significant at both time points (T1: $I^2 = 91.5, p < 0.001$; T2: $I^2 = 95.4, p < 0.001$), indicating that variation across studies was substantial, potentially limiting generalizability. Notably, heterogeneity among the studies included in a meta-analysis is very common and should be anticipated, not regarded as the exception (Berlin, 1995). The significant Egger's regression intercept for the funnel plot (bias = 9.2; $p = 0.00019$) may also be indicative of potential publication bias in the studies included in the meta-analysis. The Egger's test has low power for meta-analyses containing small to moderate numbers of studies (Sutton, 2009). However, Orwin's *Fail-safe N* is 177 ($SMD = 1.71$), suggesting a need for 177 additional studies with an effect size of 0 before the cumulative effect would become trivial (defined as a Cohen's d of 0.1). With such high numbers of studies required to overturn the conclusions of the meta-analysis, the overall observed effect size can be considered robust (Orwin, 1983; Sutton, 2009).

Conclusion

Intraocular pressure increases significantly while the patient is in the prone position. The greatest increase in IOP occurs within 10 minutes after the patient is placed into the prone position. The IOP continues to increase significantly while the patient is in prone position, but to a lesser degree. Intraocular pressure increases of the magnitude found in this systematic review and meta-analysis clearly demonstrate the need for

implementing interventions to mitigate the increase in IOP and reduce the risk for postoperative vision loss and other ocular complications in patients undergoing surgery in the prone position.

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Table 4.1.

Summary of Studies Included in the Systematic Review and Meta-analysis (s = 9; N = 229).

First Author (Year) Country	Study Design (Quality)^a	Participant Groups and Observations/Interventions	IOP Measures (Tonometer) [Head Positioning]	Outcome Measures	Effect Sizes (Cohen's <i>d</i>)
Agah (2011) Iran	Non-experimental (Good)	• 20 patients undergoing PCNL in prone position	Measurement of IOP at five different time points (Tono-Pen XL) [Ring-shaped headrest]	Differences in IOP levels at each time point	T1: 11.34 T2: 10.5
Carey (2014) United States	Experimental (High)	• 7 patients undergoing spine surgery in neutral prone position • 6 patients undergoing spine surgery in 5° reverse Trendelenburg position ^b • 6 patients undergoing spine surgery in 10° reverse Trendelenburg position ^b	Measurement of IOP at five different time points (Tono-Pen XL) [Horseshoe-shaped headrest]	Differences in IOP levels at each time point	
Cheng (2001) United States	Non-experimental (Good)	• 20 men undergoing spine surgery in prone position	Measurement of IOP at five different time points (Tono-Pen XL) [Skull pins]	Differences in IOP levels at each time point	
Czorlich (2018) Germany	Non-experimental (High)	• 16 patients undergoing spine surgery and dural opening in prone position (A) • 16 patients undergoing cranial surgery and dural opening in the prone position (B) • 16 patients undergoing cranial surgery and dural opening in the lateral position (C) ^b	Measurement of IOP at seven different time points (Tono-Pen XL) [Skull clamp]	Differences in IOP levels at each time point	T1A: 2.82 T1B: 3.44 T1D: 2.32

First Author (Year) Country	Study Design (Quality)^a	Participant Groups and Observations/Interventions	IOP Measures (Tonometer) [Head Positioning]	Outcome Measures	Effect Sizes (Cohen's <i>d</i>)
Emery (2015) United States	Experimental (High)	<ul style="list-style-type: none"> 16 patients undergoing spine surgery in prone position (D) <p><i>Note.</i> Patients with BMI > 30 kg/m² excluded.</p>			
Hunt (2004) United Kingdom	Non-experimental (Good)	<ul style="list-style-type: none"> 27 patients undergoing spine surgery in prone position 25 patients undergoing spine surgery in 10° reverse Trendelenburg position^b 10 patients undergoing spine surgery in prone position 10 patients undergoing spine surgery in prone position 	Measurement of IOP at six to 19 different time points (Tono-Pen XL) [Tong traction]	Differences in IOP levels at each time point	T1: 1.97
Nuri Deniz (2013) Turkey	Experimental (Good)	<ul style="list-style-type: none"> 23 patients undergoing PCNL in prone position 22 patients undergoing PCNL in prone position with 45° right lateral head rotation^b 	Measurement of IOP at three different time points (Tono-Pen XL) [PL: Pillow or Ring-shaped headrest PN: Skull pins]	Differences in IOP levels at each time point	T1PL: 1.28 T1PN: 1.08 T2PL: 0.58 T2PN: 0.5
Pinar (2018) Turkey	Experimental (High)	<ul style="list-style-type: none"> 20 patients undergoing lumbar disc surgery in prone position with general anesthesia (G) 20 patients undergoing lumbar disc surgery in prone position with spinal anesthesia (S) 	Measurements of IOP at four different time points (Tono-Pen AVIA) [Horseshoe-shaped headrest]	Differences in IOP levels at each time point	T1G: 0.52 T1S: 0.39 T2G: 3.9 T2S: 2.98

First Author (Year) Country	Study Design (Quality)^a	Participant Groups and Observations/Interventions	IOP Measures (Tonometer) [Head Positioning]	Outcome Measures	Effect Sizes (Cohen's <i>d</i>)
Suguta (2012) Japan	Non-experimental (High)	<ul style="list-style-type: none"> • 12 patients undergoing spine surgery in prone position with propofol anesthesia (P) • 12 patients undergoing spine surgery in prone position with sevoflurane anesthesia (S) 	Measurements of IOP at five different time points (Tono-Pen XL) [Horseshoe-shaped headrest]	Differences in IOP levels at each time point	T1P: 3.01 T1S: 3.61

Note. *s* = studies; IOP = intraocular pressure; PCNL = percutaneous nephrolithotomy; BMI = body mass index; kg/m² = kilograms/meter-squared.

^aQuality ratings are based on cumulative scores obtained from the Association of periOperative Registered Nurses (AORN) Research Evidence Appraisal Tool – Study.

^bThis group was not eligible for inclusion in the meta-analysis.

Table 4.2

Study and Participant Characteristics (s = 9; N = 229)

Category	Number of Participants (Number of Studies)	Mean \pm SD (Range) or Percent
Age (years)	222 (8)	53.2 ± 8.1 (43.3 - 69)
Gender	222 (8)	
Women	100	45.0
Men	122	55.0
BMI (kg/m^2)	155 (5)	24.9 ± 1.6 (22.5 – 27.7)
ASA Class I, II, or III	130 (5)	100
Surgery type	229 (9)	
Cranial	16	7.0
PCNL	43	18.8
Spine	170	74.2
Anesthesia	222 (8)	
General-Inhalation	115	51.8
General-Propofol	60	27.0
General-Unspecified	27	12.2
Spinal	20	9.0
Duration (min)		
Anesthesia	24 (1)	247.0 ± 9.9 (240 - 254)
Prone position	44 (2)	161.6 ± 70.9 (80 - 203)
Surgery	74 (4)	156.0 ± 24.2 (120 - 181)
Estimated blood loss (mL)	95 (3)	330.1 ± 222.1 (120 – 615)

Note. s = studies; SD = standard deviation; BMI = body mass index; kg/m^2 = kilograms/meter-squared; ASA = American Society of Anesthesiologists Physical Classification Status; PCNL = percutaneous nephrolithotomy; min = minutes; mL = milliliters.

Table 4.3

Effect Sizes and Magnitude of Change in IOP for T1 through T2 (s = 9; N = 229).

Time Points	s	k	n	Model	d	95% CI		Q	I^2	Mean change in IOP ^a	Prediction Interval ^b
						LL	UL				
T1	6	11	179	R	2.55*	1.61	3.5	117.8*	91.5	+7.5 mmHg	+2.6 mmHg to +19.8 mmHg
T2	3	5	80	R	3.44**	1.25	5.64	86.11*	95.3	+10.1 mmHg from T1	+10.5 mmHg to +38.0 mmHg

Note. s = studies; k = comparisons; d = standardized mean difference; CI = confidence interval; LL lower limit; UL = upper limit; Q = Cochrane's Q; I^2 = heterogeneity statistic; IOP = intraocular pressure; T = time point; R = random effects; mmHg = millimeters of mercury.

^aMean effect sizes were converted to the metric used to measure IOP (i.e., mmHg) following the procedures described by Lipsey and Wilson (2001).

Lipsey, M. W., & Wilson, D. B. (2001). Interpreting and using meta-analysis results. In *Practical meta-analysis: Applied social research* (Vol 49, pp. 146-168). Thousand Oaks, CA: Sage.

* $p \leq 0.001$; ** $p = 0.002$.

^bIn 95% of all populations, the true effect size will fall within this range.

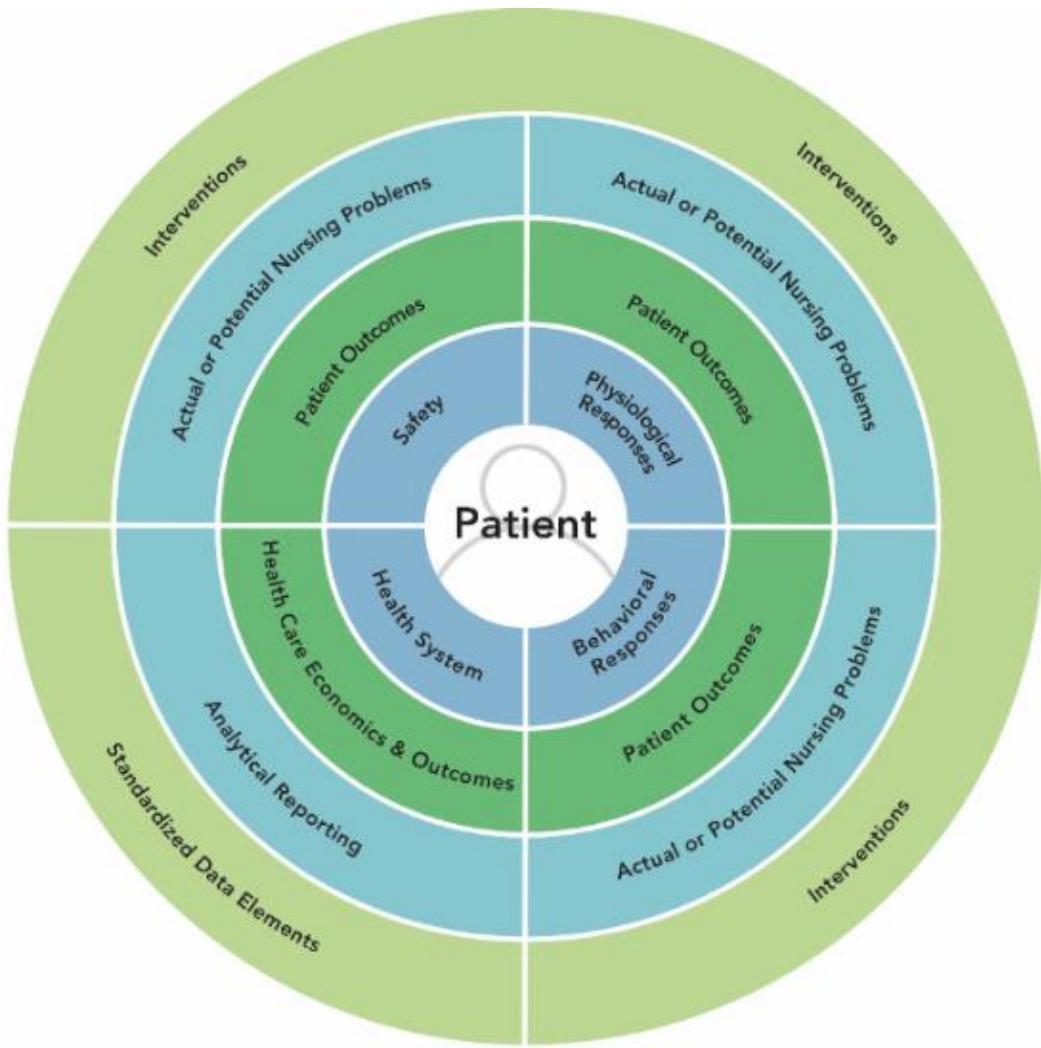


Figure 4.1. Perioperative Patient Focused Model (2017). Reprinted with permission from Guidelines for Perioperative Practice. Copyright © 2018, AORN, Inc, 2170 S. Parker Road, Suite 400, Denver, CO 80231. All rights reserved.

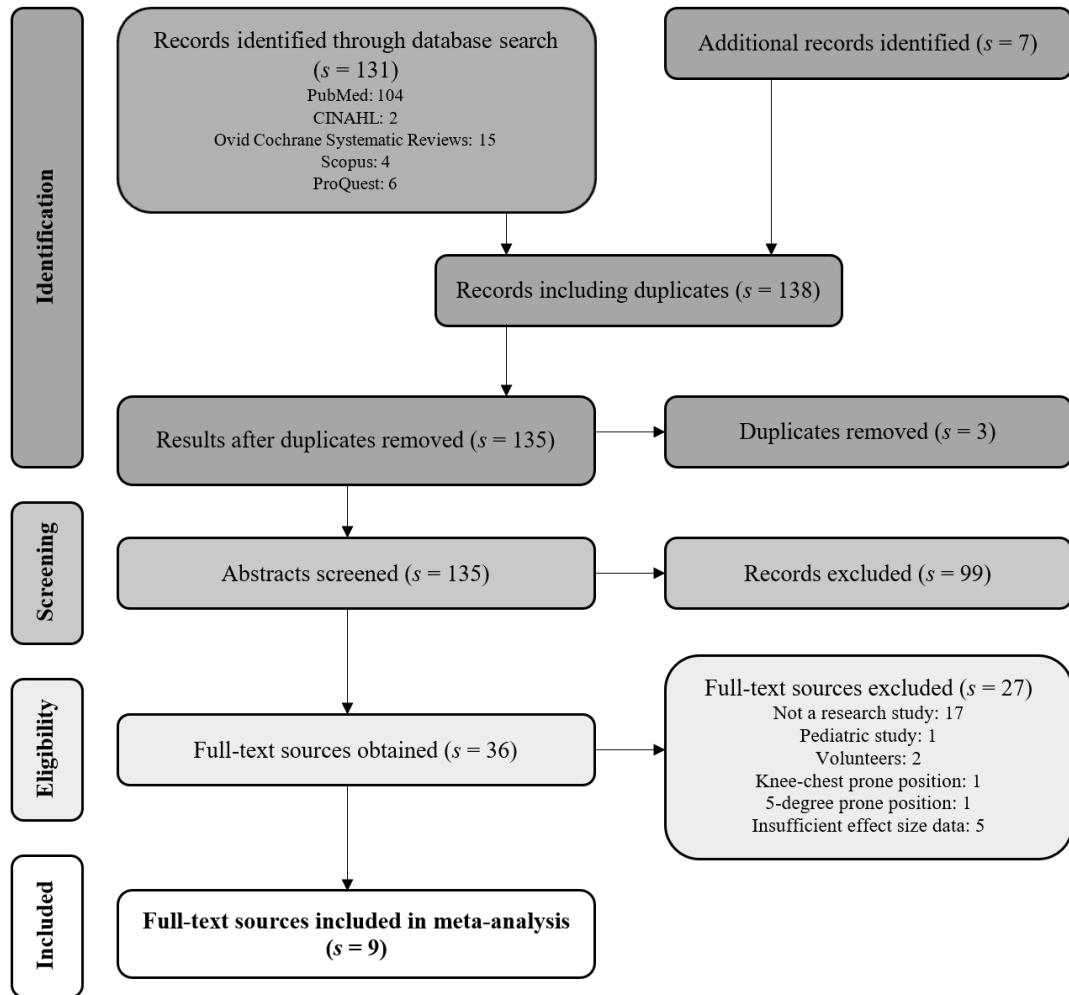


Figure 4.2. Flow diagram of meta-analysis study selection. Note. s = studies. Adapted from Moher D, Liberati A, Tetzlaff J, Atman DG, PRISMA Group (2009). Preferred Reporting Items for Systematic Reviews and Meta-Analyses: The PRISMA Statement. *PLoS Medicine*, 6(6), e1000097. doi:10.1371/journal.pmed1000097

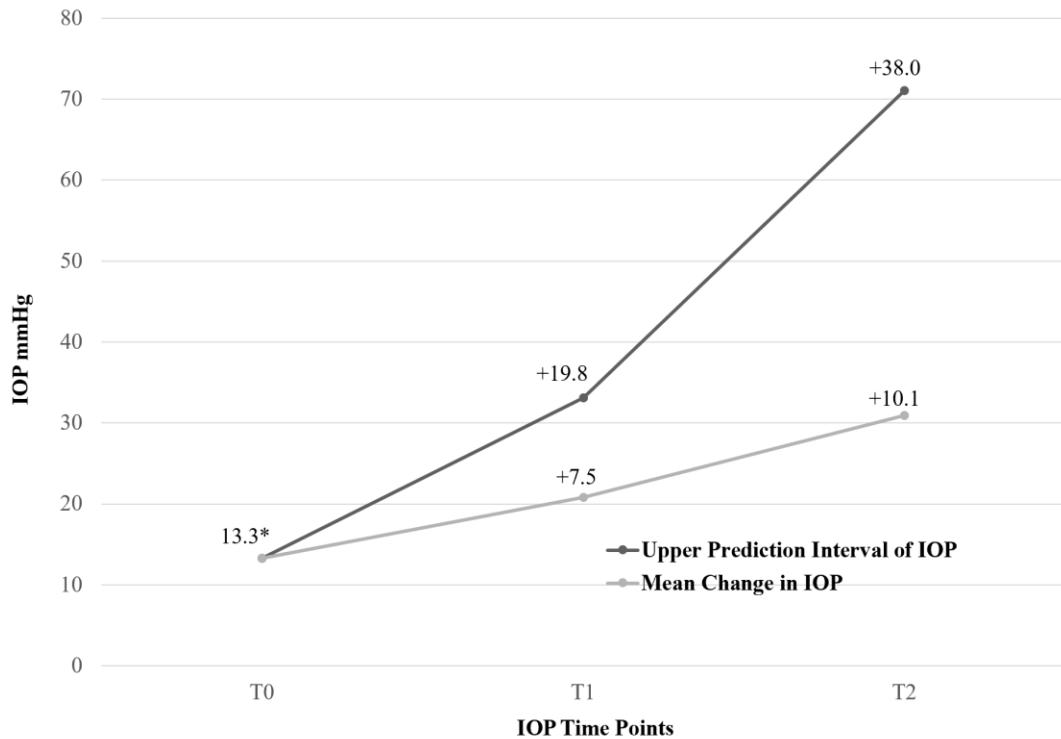


Figure 4.3. Magnitude of change in IOP and upper prediction intervals of IOP for T0 through T2. *Note.* IOP = intraocular pressure; mmHg = millimeters of mercury; T = time point. Intraocular pressure increases significantly when the patient is placed in prone position (T1: +7.5 mmHg; $p < 0.001$), and continues to increase significantly until the end of prone position (T2: +10.1 mmHg; $p = 0.002$). The upper prediction interval shows that in 95% of all populations, at the end of prone position, IOP could increase to 71.1 mmHg ($13.3 \text{ mmHg} + 19.8 \text{ mmHg} + 38.0 \text{ mmHg} = 71.1 \text{ mmHg}$). *Pooled mean at T0—After induction of anesthesia.

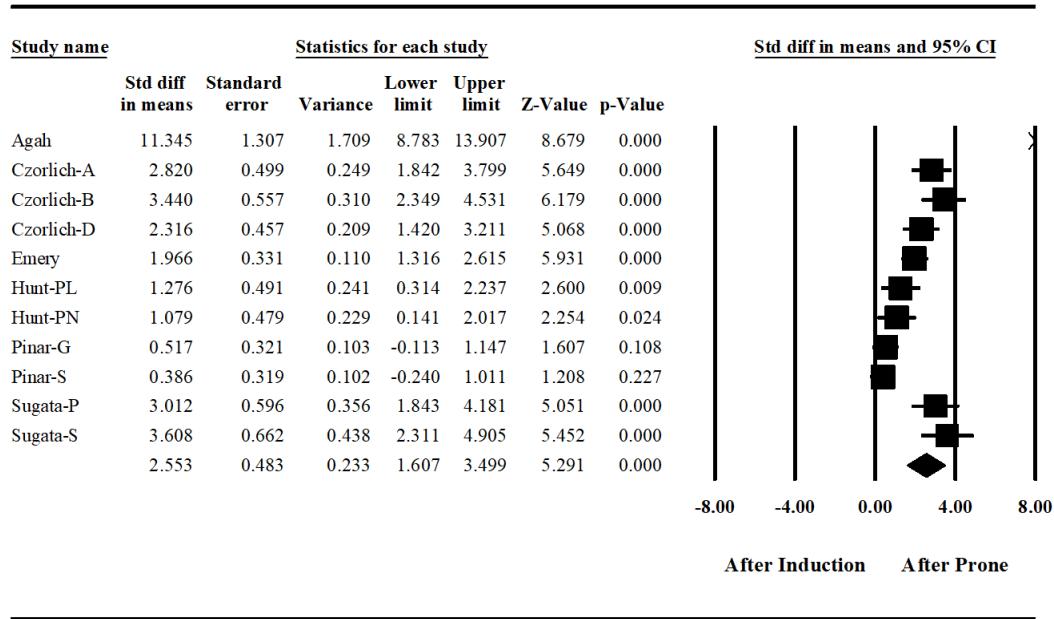


Figure 4.4. Forest plot of meta-analysis of T1. Note. T = time point; Std = Standard; min = minutes. This analysis included six studies representing 11 participant groups ($n = 179$). Effect sizes were calculated using a random effects model. The area of each square is proportional to study weight.

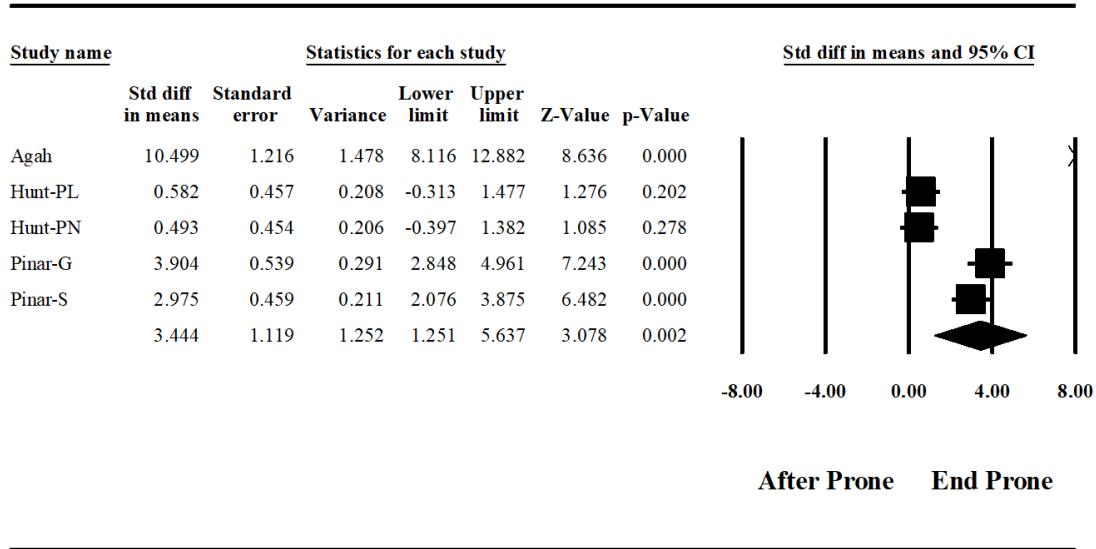


Figure 4.5. Forest plot of meta-analysis of T2. Note. T = time point; Std = Standard; min = minutes. This analysis included three studies representing five participant groups ($n = 80$). Effect sizes were calculated using a random effects model. The area of each square is proportional to study weight.

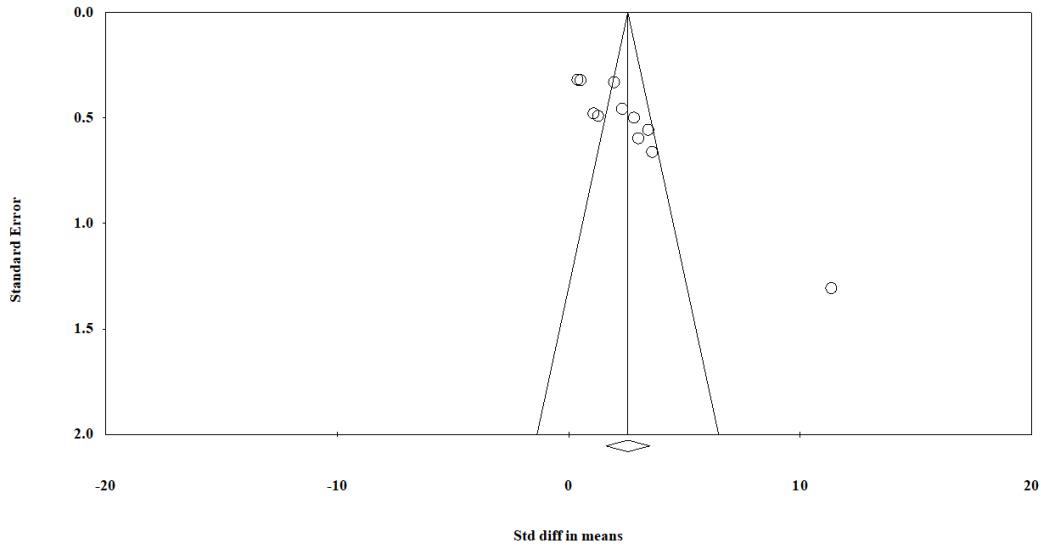


Figure 4.6. Funnel plot of publication bias for T1. Note. T = time point; Std = standard. Larger studies are shown at the top of the funnel. Positive smaller studies should appear at the right of the mean effect size (i.e., center line). The significant Egger's regression intercept for this funnel plot (bias = 9.2; $p = 0.00019$) may be indicative of potential publication bias in the studies included in the meta-analysis for this time point. However, Orwin's *Fail-safe N* is 177 ($SMD = 1.71$), suggesting a need for 177 additional studies with an effect size of 0 before the cumulative effect would become trivial (defined as a Cohen's d of 0.1).

5. SUMMARY

The results of the systematic reviews and meta-analyses have shown that intraocular pressure (IOP) increases significantly for adult patients undergoing surgery in the Trendelenburg and prone positions. Increased IOP puts the patient at risk for glaucoma, detached retina, or partial to complete vision loss (Amorim Correa & Acioly, 2018; Astuto, Minardi, Uva, & Gullo, 2011; Borahay et al., 2013; Emery et al., 2015; Ghomi, Kramer, Askari, Chavan, & Einarsson, 2012; Gkegkes, Karydis, Tyritzis, & Iavazzo, 2015; Gould, Cull, Wu, & Osmundsen, 2012; Grosso et al., 2013; Hoshikawa et al., 2014; Kwee, Ho, & Rozen, 2015; Lee & Newman, 2018; Leibovitch, Casson, Laforest, & Selva, 2006; Tacketti et al., 2015). Intraocular pressure increases of the magnitude found in these systematic reviews and meta-analyses demonstrate the need for implementing intraoperative interventions to mitigate the increase in IOP and reduce the potential for postoperative vision loss and other ocular complications in patients undergoing surgery in the Trendelenburg and prone position.

Recommendations for Patients Undergoing Surgery in the Trendelenburg Position

Interventions that may be helpful for mitigating or reducing IOP when the patient is undergoing surgery in the Trendelenburg position include

- monitoring IOP at established intervals or continuously (Hoshikawa et al., 2013; Lee, Dallas, Daniel, & Cotter, 2016; Vitish-Sharma et al., 2018),
- reducing the degree of Trendelenburg position (Ghomie et al., 2012; Mathew et al., 2018; Ozcan et al., 2017; Raz et al., 2015),
- implementing a modified Trendelenburg position (Raz et al., 2015),

- providing periodic position changes or rest periods (Blecha et al., 2017; Borahay et al., 2013; Freshcoln & Diehl, 2014; Gkegkes et al., 2015; Gould et al., 2012; Mizrahi, Hugkulstone, Vyakarnam, & Parker, 2011; Molloy & Watson, 2012), and
- administering specific medications or anesthetics (Agrawal, Dureja, Verma, & Kang, 2013; Hwang et al., 2013; Joo, Koh, Lee, & Lee, 2016; Joo, Kim, & Lee, 2017; Kan, Brown, & Gainsburg, 2015; Kaur, Sharma, Kalra, Purohit, & Chauhan, 2018; Kim et al., 2015; Mathew et al., 2018; Molloy, Cong, & Watson, 2016; Mowafi, Al-Ghamdi, & Rushood, 2003).

Because IOP increases during abdominal insufflation and Trendelenburg position, intraoperative monitoring of IOP either continuously or at established intervals or time points (e.g., after abdominal insufflation, after initiation of Trendelenburg position, after 60 minutes of Trendelenburg position, etc.) seems prudent. Elevated IOPs can be an indication of ocular venous congestion and decreased perfusion of the optic nerve (Yoo et al., 2014). Monitoring IOP can provide a baseline IOP and an objective measure that can help the surgical team maintain awareness of the patient's IOP, implement interventions to reduce IOP as needed, and thus reduce the potential for ocular complications and postoperative vision loss (Hoshikawa et al., 2013; Lee et al., 2016).

Steeper degrees of Trendelenburg increase the risk for postoperative complications because they place greater physiologic stress on the patient's body (Ghomie et al., 2012; Gould et al., 2012; Kadono et al., 2013). Ghomi et al., (2012) found that robotic-assisted gynecologic surgery could be performed successfully with a modest head-down tilt of 16.4 degrees. In a study to determine the head-down tilt necessary to provide surgical access and visibility, Gould et al., (2012) found the mean head-down tilt

most often selected by the surgeons was 28.1 degrees, which was much less than the 40-degree head-down tilt the surgeons were using.

Raz et al., (2015) found that modifying the Trendelenburg position so that the patient's head and shoulders remained level significantly decreased IOP and accelerated its return to baseline levels. Implementing periodic intraoperative position changes or rest periods in supine position (or positions where the ocular level is above the heart) can help to reduce IOP. In a quasi-experimental study, Molloy and Watson (2012) implemented a five-to-seven-minute level supine intervention after 60 minutes of 32-degree to 40-degree Trendelenburg position and found there was a significant decrease in IOP after 120 minutes of Trendelenburg position. The dramatic and significant decrease in IOP that occurs before arousal from anesthesia found in this systematic review and meta-analysis (T5: -7.5 mmHg, $p < 0.001$; T7: -8.2 mmHg, $p < 0.001$; T9: -6.0 mmHg, $p < 0.001$) also supports the implementation of periodic intraoperative position changes or rest periods as a mechanism to help reduce IOP.

Administering specific medications or anesthetics may also be effective in reducing IOP or mitigating the intraoperative increase in IOP (Agrawal et al., 2013; Hwang et al., 2013; Joo et al., 2016; Joo et al., 2017; Lee et al., 2016; Kaur et al., 2018; Kim et al., 2015; Mathew et al., 2018; Molloy et al., 2016; Mowafi et al., 2003). Agrawal et al., (2013) found that induction and maintenance of anesthesia with intravenous propofol was the most effective option for mitigating the increase in IOP in adult patients undergoing surgery in the Trendelenburg position. Likewise, Kaur et al., (2018) found that propofol-based total intravenous anesthesia was more effective than inhalational anesthesia with sevoflurane in mitigating the increase in IOP in patients undergoing

laparoscopic surgery in the Trendelenburg position. Kitamura et al., (2018) found that continuous administration of dexmedetomidine in combination with propofol-based total intravenous anesthesia decreased IOP in patients undergoing robotic-assisted laparoscopic radical prostatectomy. Molloy and Cong (2014) found that intraoperative treatment with dorzolamide-timolol eyedrops significantly reduced elevated IOP in patients undergoing lengthy laparoscopic procedures in the Trendelenburg position, while Molloy et al., (2016) found that prophylactic therapy with dorzolamide-timolol eyedrops significantly reduced IOP in patients undergoing robotic-assisted laparoscopic prostate and gynecologic procedures.

Recommendations for Patients Undergoing Surgery in the Prone Position

Interventions that may be helpful for mitigating or reducing IOP when the patient is undergoing surgery in the prone position include

- positioning the patient in a 5-degree to 10-degree reverse Trendelenburg prone position (Carey, Shaw, Weber, & DeVine, 2014; Emery et al., 2015; Fukui, Ahmad, McHugh, Tempelhoff, & Cheng, 2004; Fukui, Tempelhoff, & Cheng, 2005; Ozcan et al., 2004; Walick, Kragh, Ward, & Crawford, 2007),
- monitoring IOP at established intervals or continuously (Eddama, 2013; Yoshimura, Hayashi, Tanake, Nomura, & Kawaguchi, 2015),
- reducing the length of time the patient is in the prone position,
- staging procedures when patients will be in the prone position for prolonged periods of time (ASA Task Force on Perioperative Visual Loss, North American Neuro-Ophthalmology Society, & Society for Neuroscience in Anesthesiology and Critical Care, 2019),

- providing periodic position changes or rest periods (Molloy & Watson, 2012),
- preventing direct pressure on the patient's eyes and assessing and monitoring the eyes at regular intervals during the procedures (ASA Task Force on Perioperative Visual Loss et al, 2019; Locastro, Novak, & Biglan, 1991; Nickels, Manlapaz, & Farag, 2014; Shifa, Abebe, Bekele, & Habte, 2016),
- avoiding specific headrests or positioning devices that may increase pressure on the eye or using skull pins or tongs to position the head (Asok, Aziz, Faisal, Tan, & Mallika, 2009; Bekar, Türeyen, & Aksoy, 1996; Grossman & Ward, 1993; Hollenhorst, Svien, & Benoit, 1954; Hoski, Eismont, & Green, 1993; Quraishi, Wolinsky, & Gokaslan, 2012; Wolfe, Lospinuso, & Burke, 1992), and
- administering specific medications or anesthetics (Farag et al., 2012; Pinar et al., 2018; Sugata et al., 2012).

Positioning the patient in a 5-degree to 10-degree reverse Trendelenburg prone position may be a simple intervention to prevent some instances of postoperative vision loss. This position has been shown to decrease IOP in healthy volunteers (Ozcan et al, 2004; Walick et al., 2007) and in patients undergoing spine surgery (Carey et al., 2014; Emery et al., 2015; Fukui et al., 2004; Fukui et al., 2005). Positioning surgical patients with the head above the heart helps reduce venous congestion in the eye and orbit and decrease intraocular and intraorbital pressure (Bonnaig, Dailey, & Archdeacon, 2014; Carey et al., 2014; Grant et al., 2010; Kamel, & Barnette, 2014; Nickels et al., 2014).

Reducing the length of time the patient is in the prone position may also help to mitigate the increase in IOP. The ASA Task Force on Perioperative Visual Loss et al., (2019) suggests staging procedures when patients will be in the prone position for

prolonged periods of time. Utilizing a series of shorter procedures rather than one prolonged procedure may help reduce the patient's risk for postoperative vision loss; however, the risks associated with multiple surgeries may outweigh the benefits of staged procedures (Shifa et al., 2016).

Because IOP increases during prone position, intraoperative monitoring of IOP either continuously or at established intervals or time points (e.g., after initiation of prone position, after every 60 minutes of prone position) seems prudent. Yoshimura et al., (2015) suggested that measuring IOPs after one hour of surgery in the prone position could provide an opportunity for implementing interventions to prevent additional increases in IOP. Eddama (2013) also suggested that regular measurement of IOP during prolonged surgery provided an opportunity for implementing a change in the patient's position when critical thresholds are reached. Implementing periodic intraoperative position changes or rest periods (where the ocular level is above the heart) can help to reduce IOP. In a quasi-experimental study, Molloy and Watson (2012) implemented a five-to-seven-minute level supine intervention after 60 minutes of steep Trendelenburg position and found there was a significant decrease in IOP.

When the patient is in the prone position, there is a risk for direct compression on the eye (Bonnaig et al., 2014). Yu, Chou, Yang, and Chang (2010) found that the prone position was a precipitating factor for eye injury. Preventing direct pressure and assessing and monitoring the patient's eyes at regular intervals throughout the procedure may help to reduce the incidence of postoperative vision loss (ASA Task Force on Perioperative Visual Loss et al., 2019; Locastro et al., 1991; Nickels et al., 2014; Shifa et al., 2016). Avoiding specific headrests or positioning devices that may increase pressure on the eye

(e.g., horseshoe-shaped, Wilson frame), or using skull pins or tongs to position the head may also help to prevent pressure on the orbits and reduce the risk for postoperative vision loss and other ocular complications (Asok et al., 2009; Bekar et al., 1996; Grossman & Ward, 1993; Hollenhorst et al., 1954; Hoski et al., 1993; Quraishi et al., 2012; Wolfe et al., 1992). Direct compression from a horseshoe-shaped head positioner has been reported as a cause of postoperative vision loss when the patient is in the prone position (Abraham, Sakhija, Sinha, & Rastogi, 2003; Bekar et al., 1996; Grossman & Ward, 1993; Hollenhorst et al., 1954; Hoski et al., 1993; Locastro et al., 1991; Wolfe et al., 1992). In a case-controlled study of 80 patients with ischemic optic neuropathy compared with 315 matched control patients, the ASA Postoperative Visual Loss Study Group (2012) found that Wilson frame use was an independent risk factor for postoperative vision loss.

Administering specific medications or anesthetics may also be effective in reducing IOP or mitigating the intraoperative increase in IOP (Farag et al., 2012; Pinar et al., 2018; Sugata et al., 2012). Pinar et al., (2018) found that the increase in IOP was significantly less in patients undergoing lumbar disc surgery in the prone position under spinal anesthesia compared with patients receiving general anesthesia. Sugata et al., (2012) found that IOPs were higher in patients undergoing prone spine surgery with general anesthesia maintained with sevoflurane compared with patients receiving general anesthesia maintained with intravenous propofol. Farag et al., (2012) found that the administration of topical brimonidine 2% helped reduce intraoperative IOP.

Recommendations for Preoperative Ophthalmologic Examinations

Another important consideration is the need to determine whether patients undergoing surgery in the Trendelenburg or prone positions should receive a preoperative ophthalmologic examination to reduce the risk for ocular injury (Akhaddar & Boucetta, 2012; ASA Task Force on Perioperative Visual Loss et al., 2019; Borahay et al., 2013; Lee et al., 2016; Singer & Salim, 2010; Stang-Veldhouse, Yeu, Rothenberg, & Mizen, 2010; Stewart, Landy, & Lee, 2016). Preoperative ophthalmologic examinations may be helpful in identifying patients at risk for postoperative vision loss. Increases in IOP may be more harmful in older patients or patients who are predisposed to developing diabetes or glaucoma than in younger, healthier patients (Akhaddar & Boucetta, 2012; ASA Task Force on Perioperative Visual Loss et al., 2019; Borahay et al., 2013; Grosso et al., 2013; Mondzelewski et al., 2015; Singer & Salim, 2010; Stang-Veldhouse et al., 2010; Stewart et al., 2016; Taketani et al., 2015). Patients at risk for acute angle-closure glaucoma associated with the prone position may benefit from preoperative laser iridotomy (Singer & Salim, 2010; Stewart et al., 2016). The ASA Task Force on Perioperative Visual Loss et al., (2019) recommend evaluating the need for preoperative ophthalmologic examination on a case-by-case basis.

Perioperative Patient Focused Model

The results of this systematic review and meta-analysis support the use of the Perioperative Patient Focused Model as the conceptual foundation for this perioperative research. Providing quantitative evidence about the magnitude of the intraoperative increase in IOP resulting from the Trendelenburg and prone positions supports the implementation of nursing interventions that are patient-focused and will improve patient

outcomes by mitigating increases in IOP and reducing the risk for permanent postoperative vision loss and other ocular complications in adult surgical patients.

Conclusion

Intraocular pressure increases significantly between abdominal insufflation in supine position and 240 minutes of Trendelenburg position. The greatest increases in IOP occur after insufflation of the abdomen while the patient is in the supine position and within five minutes after the patient is placed into the Trendelenburg position. The IOP continues to increase significantly while the patient is in Trendelenburg position, but to a lesser degree. Intraocular pressure increases significantly while the patient is in the prone position. The greatest increase in IOP occurs within 10 minutes after the patient is placed into the prone position. The IOP continues to increase significantly while the patient is in prone position, but to a lesser degree. Intraocular pressure increases of the magnitude found in these systematic reviews and meta-analyses clearly demonstrate the need for implementing interventions to mitigate or lessen the increase in IOP and reduce the risk for postoperative vision loss and other ocular complications in patients undergoing surgery in the Trendelenburg and prone positions.

Appendix A-1

AORN

AORN RESEARCH EVIDENCE
APPRaisal TOOL - STUDY

DATE _____
REVIEWER _____
APPRaisal SCORE _____

RW#	CITATION
<p>Does this evidence address the perioperative practice question? <input type="checkbox"/> Yes <input type="checkbox"/> No - Do not proceed with evidence appraisal.</p> <p>Does this evidence have a major flaw? <input type="checkbox"/> No <input type="checkbox"/> Yes - Determine level of evidence and score quality as C. Provide explanation of flaw in comments.</p>	
LEVEL OF EVIDENCE	
<p>Is this a report of a single research study? <input type="checkbox"/> Yes <input type="checkbox"/> No (If No, go to the AORN Research Evidence Appraisal Tool - Summary)</p>	
<p>INTERVENTION/MANIPULATION <input type="checkbox"/> Yes <input type="checkbox"/> No The researcher performed an intervention with at least some of the participants (ie, there was some type of treatment being tested).</p>	
<p>CONTROL/COMPARISON GROUP <input type="checkbox"/> Yes <input type="checkbox"/> No The researcher provided standard care or a comparison intervention that was different from the experimental intervention.</p>	
<p>RANDOM ASSIGNMENT <input type="checkbox"/> Yes <input type="checkbox"/> No The researcher assigned participants to a control or treatment group on a random basis (ie, in a manner determined by chance).</p>	
YES to Intervention/Manipulation, Control/Comparison Group, and Random Assignment	<input type="checkbox"/> LEVEL I Randomized Controlled Trial (RCT)
YES to Intervention/Manipulation or YES to Intervention/Manipulation, and Control/Comparison Group	<input type="checkbox"/> LEVEL II Quasi-Experimental (eg, controlled trial, controlled trial without randomization, pre-test/post-test, time series)
NO to Intervention/Manipulation	<input type="checkbox"/> LEVEL III Non-Experimental (eg, descriptive, comparative, observational, correlational, case-control, retrospective, cross-sectional) <input type="checkbox"/> LEVEL III Qualitative (eg, interviews, surveys, focus groups)

ADDITIONAL COMMENTS:

Appendix A-2

QUALITY OF EVIDENCE	A HIGH	B GOOD	C LOW	NA
PURPOSE/BACKGROUND				
• Was the purpose of the systematic review clearly defined?				
• Was the research question clear?				
• Did the researcher(s) identify what is known and not known about the research question and how the systematic review would address any gaps in knowledge?				
SEARCH				
• Was the search strategy reproducible?				
• Were the key search terms stated?				
• Were multiple databases searched and identified?				
• Was the inclusion/exclusion criteria described?				
• Was both published and unpublished literature identified and retrieved where possible?				
• Are the types of studies to be included in the review described?				
EVIDENCE REVIEW				
• Was there an explanation of the number of studies eliminated at each level of review?				
• Were the details of the included studies presented (design, sample, methods, results, outcomes, strengths, limitations)?				
• Were methods for appraising the strength of evidence (level and quality) rigorous?				
• Was the evidence reviewed and appraised by at least two members of the research team?				
• Were the supporting references the most current available?				
• Were the supporting references relevant to the research question?				
DATA COLLECTION				
• Were methods of statistical analysis described?				
• Were methods of retrieving data from the individual studies described?				
• Was the data extracted by at least two members of the research team?				
RESULTS/CONCLUSIONS				
• Were the conclusions of the researcher(s) consistent with the results of the studies and the overall strength of the evidence?				
• Was the strength of the phenomenon being studied quantified in a summary statistic (ie, effect size) that can be compared across the studies?				
LIMITATIONS/FUTURE RESEARCH				
• Were limitations of the review discussed?				
FINAL QUALITY SCORE				

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VITA

Sharon Ann Van Wicklin has more than 45 years of experience as a perioperative registered nurse (RN). She has worked in all facets of the perioperative environment from scrub person to director. Sharon received her Associate of Science in Nursing from Ventura College, Ventura, California. She received her Bachelor of Science in Nursing and Master of Science in Nursing from Middle Tennessee State University, Murfreesboro, Tennessee. Sharon is currently a Doctor of Philosophy in Nursing candidate at Missouri State University, Columbia, Missouri. She is a member of the Sigma Theta Tau Honor Society of Nursing and the Honor Society of Phi Kappa Phi. Sharon has earned certification in operating room nursing (CNOR), as an emeritus RN first assistant (CRNFA[E]), as a retired plastic and reconstructive surgical nurse (CPSN-R), and as a legal nurse consultant (PLNC).

Sharon was recognized by the Association of periOperative Registered Nurses (AORN) as a recipient of the Outstanding Achievement in the Application of Perioperative Clinical Research Award in 2005. This award recognizes a perioperative RN whose application of perioperative clinical research reflects the goal of excellence in patient care. In 2017, Sharon received the honor of being inducted as a Fellow in the American Academy of Nursing (FAAN), and in 2018, she was inducted as a Fellow in the International Society of Plastic and Aesthetic Nurses (ISPAN-F).

In a previous position as a perioperative educator, Sharon was responsible for the creation and coordination of educational projects and programs designed to improve hospital processes for orientation and ongoing educational development for more than 125 personnel in nine perioperative departments. As a Senior Perioperative Practice Specialist for AORN, Sharon provided consultative services, authored various AORN publications including the

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In her current position as a perioperative consultant and editor-in-chief of *Plastic Surgical Nursing*, Sharon serves as a clinical consultant, peer reviewer, clinical editor, author, and presenter for a variety of perioperative educational products, seminars, workshops, and publications. Her work as a legal expert witness involves reading and reviewing medical records and testifying as to the standard of perioperative nursing care.