



## General anesthesia for cesarean section: are we doing it well?

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Korea has a higher rate of cesarean sections under general anesthesia than in other countries. Neuraxial anesthesia is the gold standard for a cesarean section, but there are some cases in which general anesthesia is inevitable. Therefore, obstetric anesthesiologists should be familiar in performing general anesthesia for cesarean section. Rapid-sequence induction and intubation with cricoid pressure using thiopental-succinylcholine have been the standard for cesarean section under general anesthesia for a long time. Recently, with the introduction of new drugs (propofol, rocuronium, and sugammadex) and equipments (videolaryngoscopy and supraglottic airways), anesthesia methods have also gradually changed. Pursuing the safety of obstetric patients and anesthesiologists at the same time, this review will help update the knowledge or training in performing general anesthesia for cesarean section.

**Keywords:** Cesarean section; General anesthesia; Obstetrics; Rapid sequence induction.

## INTRODUCTION

Approximately one-third of all new-borns in Korea are delivered via cesarean section [1]. World Health Organization (WHO) considers the ideal rate for cesarean sections to be between 10 to 15% of total childbirths. However, there has been an explosive rise in the rate of cesarean sections worldwide. In some countries, the current cesarean section rate exceeds 50% [2].

Anesthesiologists should choose between general or regional anesthesia depending on the individual patient's condition and clinical situation. The use of appropriate and effective anesthesia for cesarean section is important not only to reduce the incidence of maternal and fetal morbidities but also to reduce the incidence of intraoperative maternal awareness.

Neuraxial anesthesia is currently recommended as the

gold standard rather than general anesthesia for most patients undergoing cesarean section [3,4], since the mortality rate of cesarean section under general anesthesia is 16.7 times higher than that under regional anesthesia [5].

There is no consensus on the ideal proportion of general anesthesia for cesarean sections, but approximately 5–6% are conducted under general anesthesia, which can be further reduced by obstetric anesthesiologist teams or obstetric specialized anesthesiologists [6,7]. However, in patients with emergent conditions (e.g., placental abruption, cord prolapse, antenatal placental bleeding, and non-reassuring fetal tracing), the rate of general anesthesia has been reported to be up to 20% [8]. Except for emergencies, induction of general anesthesia will continue in situations deemed “unavoidable and necessary,” including patient refusals or contraindications to neuraxial anesthesia [9].

As the overall safety of general anesthesia has significantly

improved over the past two decades with newly developed drugs, devices, and monitors, general anesthesia no longer has an impact on anesthesia-related maternal mortality rates [10,11]. In addition, general anesthesia is valuable in clinical obstetric conditions that require hemodynamic stability or rapid induction of anesthesia [12]. General anesthesia for cesarean sections was not associated with overall neurodevelopmental delay at two years of age [13].

The choice of general anesthesia varies across countries or hospitals. In Korea, unlike previous data from the United Kingdom or the United States, the rate of general anesthesia use is steadily decreasing with time, but it is still over 20% [14]. In some hospitals with a high volume of emergency cases or high-risk parturients, the rate of general anesthesia is close to 90% [15]. Racial, ethnic, and socioeconomic disparities also contribute to the choice of general anesthesia [11].

Opportunities for training in obstetric airway management have declined over the past four decades. A retrospective audit conducted by a single British institution reported that the use of general anesthesia for cesarean section decreased from 76% in 1982 to 7.7% in 1998 and 4.9% in 2006 [16,17]. With the worldwide declining trend of general anesthesia for cesarean sections, it is estimated that many residents/trainees will graduate without experience in inducing general anesthesia on pregnant women [18]. Reduced or biased cases of general anesthesia can deprive them of training experience and eventually affect patient safety. Continuous education and training are essential to ensure safe anesthesia. Unfortunately, most anesthesiologists base their management to previous experiences and partly outdated approaches. It is necessary to update our knowledge to provide safe anesthesia for cesarean section, especially in Korea, which has low fertility rates.

For anesthesiologists in varied settings, this review will help to update the knowledge or training in general anesthesia for cesarean section.

## INDUCTION

Rapid-sequence induction and intubation (RSII) with cricoid pressure using thiopental and succinylcholine has been the standard of general anesthesia for cesarean sections for a long time.

### Intravenous induction agents

In the past, a single dose of thiopental was recommended

as the induction agent of choice for general anesthesia in cesarean section; however, many textbooks or guidelines also recommend propofol. In fact, this would have been affected more by the current state that thiopental is no longer available in the United States. There is more evidence of a shift in propofol use [19], which is probably similar worldwide. However, it is highly likely to be used off-label because it is not licensed in many countries except the United States [20,21]. It has also not been approved for obstetric anesthesia in Korea. This needs to be corrected in the future. Except for propofol, most drugs used in obstetric anesthesia are permitted to be administered only if the therapeutic benefits exceed the risk.

The recommended dose of thiopental varies depending on the textbooks or guidelines, but it is approximately 4–6 mg/kg and has little effect on neonates up to 6 mg/kg [22,23]. Thiopental 7 mg/kg is superior to 5 mg/kg in creating a deeper hypnotic state in the parturient. However, it negatively affects Apgar scores and neonatal neurobehavioral tests [24].

Induction agents administered to the parturient are transferred to the fetus through the placenta. When a neonatologist is not present at delivery, it would be prudent to reduce the doses of induction agents to the lowest possible and to shorten the duration from the administration of anesthetics to delivery of a baby. However, initiating surgery without providing sufficient anesthesia increases the risk of the mother becoming aware of and developing tachycardia and hypertension.

During general anesthesia for cesarean section, anesthesiologists should pay attention to intraoperative maternal awareness, which shows a relatively high incidence of awareness compared to other surgeries [25]. The use of thiopental is one of the causes of awareness during cesarean sections under general anesthesia [25]. If intubated using small doses of thiopental, additional inhalational anesthetics may be required before the baby is delivered.

In terms of anesthetic depth, propofol seems to be better than thiopental [26]. The recommended dose of propofol is approximately 2.0–2.8 mg/kg. However, propofol 2.5 mg/kg is a sufficient dose for induction to prevent maternal awareness; it causes worse baby outcomes and higher reduction in maternal blood pressure than thiopental does [27,28]. Furthermore, propofol 2 mg/kg compared with thiopental 4 mg/kg tends to have a higher incidence of Apgar scores of 7 or less [29]. Although it depends on the dose of propofol used, the prevailing opinion is that propofol is associated

with a worse neonatal profile. Therefore, propofol should be used with caution in limited cases until the supply of thiopental is terminated, for example, in the presence of a medical staff to take care of the neonate entirely, in case of a hypertensive parturient or reduced dose usage.

In the presence of hemodynamic instability, ketamine (1–1.5 mg/kg), or etomidate (0.3–0.5 mg/kg) may be used in the presence of hemodynamic instability. The addition of a low dose of ketamine to thiopental is associated with better sedation and lower analgesic requirements, without side effects [30,31].

### Neuromuscular blocking agents

Until recently, succinylcholine 1–1.5 mg/kg was the standard treatment used for RSII because of its rapid onset. The only reason succinylcholine has been used since a long time is probably because it is relatively safe.

In fact, for cesarean section, there is a high risk of complications related to airway management (aspiration pneumonia, hypoxia, etc.) and a high incidence of difficult or failed intubation [19]. Therefore, succinylcholine remains the first choice for cesarean section, although its use is being discontinued because of rare but fatal side effects such as malignant hyperthermia and hyperkalemia. In addition, owing to the short duration of action of succinylcholine, spontaneous breathing can be quickly resumed if intubation is difficult or even fails.

Recently, rocuronium has replaced succinylcholine as the first choice with the introduction of sugammadex, which can be immediately reversed because the onset time is similar to that of succinylcholine [32]. However, the recommended dose of rocuronium remains controversial. Pühringer et al. [33] reported that rocuronium 0.6 mg/kg provides clinically acceptable intubating conditions because of the higher cardiac output in parturients. In contrast, McGuigan et al. [32] suggested a higher dose of 1 mg/kg of rocuronium to achieve faster and better intubation conditions without increasing the dose of hypnotics and consequently without compromising hemodynamic stability.

Due to the short duration of a cesarean section, the duration of action of rocuronium can be prolonged, and sugammadex may eventually be used. If the neuromuscular monitoring is performed and the appropriate dose of sugammadex is given adequately, rocuronium 1.0 mg/kg is considered as an appropriate dosage for RSII. However, anesthesiologists should pay attention to not administer a large amount

of rocuronium because its fetal/maternal plasma drug concentration ratio is about 0.16 [23] compared to succinylcholine, which is quickly metabolized and is not detected in the fetal vein in about 5–10 min [34].

### Cricoid pressure

Cricoid pressure, also known as the Sellick maneuver, to prevent pulmonary aspiration, is widely used in RSII [20]. Some textbooks also recommend a more accurate (10 N while awake; increased to 30 N after loss of consciousness) cricoid pressure [35,36]. However, some studies have indicated that cricoid pressure is difficult to apply because it is difficult to compress the cricoid cartilage efficiently and causes discomfort in conscious patients [37]. In addition, questions have been raised regarding the effectiveness of cricoid pressure in preventing aspiration [38]. It is necessary to apply cricoid pressure in patients at a high risk of aspiration so that it can be appropriately provided with sufficient force. Ideally, pressure should be applied on the cricoid cartilage towards the body of C6 directed at 90° to the tilted table.

### Videolaryngoscopy and supraglottic airways

The most important recent changes in difficult airway management are the introduction of videolaryngoscopy (VL) and supraglottic airways (SGA).

The usefulness of VL has already been demonstrated in adults who require intubation [39] and especially in obese patients [40]. This supports its increased adoption in obstetrics, where VL, rather than direct laryngoscopy, is recommended as the first attempt at intubation for all obstetric patients [36,41].

SGAs play an important role in the airway management. Unlike VL, SGAs enable ventilation even in patients with difficult facemask ventilation and simultaneous use as a conduit for tracheal intubation [42]. Therefore, the use of SGAs is now widely recommended in many guidelines for difficult airway algorithms [43–45]. Although SGAs are reasonable alternatives to endotracheal intubation, they are not recommended as the first line for elective cesarean section [36], but are recommended as a device for rescue ventilation. For pregnant patients, the use of second-generation SGAs is recommended rather than first-generation SGAs when used for rescue ventilation after difficult airway management [46]. The risk of regurgitation and pulmonary aspiration may be reduced by aspirating the gastric tube passing through the

SGA and minimizing the fundal pressure at delivery [46].

## MAINTENANCE

The goals for anesthetic maintenance include (1) appropriate depth of anesthesia to prevent awareness and recall, (2) minimal adverse effects on the neonate, and (3) minimal effects on uterine contractions after delivery. These goals can be accomplished using inhalational anesthesia or, less commonly, total intravenous anesthesia (TIVA).

To minimize neonatal depression and its effect on uterine tone, an end-tidal minimum alveolar concentration (MAC) of inhalational anesthetics (0.5) has been traditionally used for cesarean section under general anesthesia. However, the use of a higher MAC is not necessarily associated with increased neonatal depression [47]. For an appropriate depth of anesthesia, the gap between the intravenous induction agent and inhalational anesthetics should be reduced. Therefore, to minimize this gap, adequate MAC should be achieved as soon as possible, for example, by using a high initial concentration of volatile agent combined with high fresh gas flows [48] or the additional use of nitrous oxide as a carrier gas. Nitrous oxide can reduce inhalational anesthetic requirements and does not decrease uterine tone. Nitrous oxide is rapidly transferred across the placenta, where fetal tissue uptake reduces the fetal arterial concentration for the first 20 min [49]. Theoretically, there is a risk of diffusion hypoxia; therefore, if it takes time from incision to delivery, lowering the concentration of nitrous oxide used or administering 100% oxygen should also be considered. Inhalational anesthetics produce dose-dependent uterine relaxation [50], which can result in uterine atony and hemorrhage. When high concentrations of inhalational anesthetics are used, uterotonic agents should be used to maintain the uterine tone.

After delivery, once the fetal transfer of medication is no longer a concern, a short-acting benzodiazepine (e.g., midazolam), a short-acting opioid (e.g., fentanyl, alfentanil, or remifentanyl), and/or nitrous oxide 50–70% can be added to allow a reduced dose of inhalational anesthetics to 0.5–0.75 MAC. Conversely, TIVA can replace inhalational anesthetics. Currently, propofol is the only drug that can be used for TIVA. Propofol can relax the uterus less than inhalational anesthetics [51].

All opioids, particularly those with high lipid solubility (e.g., remifentanyl, fentanyl, and sufentanil), readily pass through the placenta to the fetus. Consequently, opioid ad-

ministration is usually avoided until after delivery to reduce the risk of neonatal depression.

Dexmedetomidine has recently been reported to enhance oxytocin-induced contractions and is expected to be used as a sedative after fetal delivery in the future [51].

## RECOVERY

Extubation should be performed with the patient fully awake while maintaining airway reflexes because anesthesia-related deaths from airway obstruction or hypoventilation occur during emergence and recovery, and not during the induction of general anesthesia [52]. After surgery, the patient should still be kept in a closely monitored environment.

## CONCLUSIONS

To date, RSII with cricoid pressure is the standard procedure for cesarean section under general anesthesia. It has been used safely for a long time with thiopental-succinylcholine-inhalational anesthetics and has not undergone procedural changes. However, induction of general anesthesia for cesarean section is relatively uncommon worldwide; hence, trainees have lesser experience with general anesthesia now than they had in the past. Therefore, it is essential that all obstetric anesthesiologists maintain their skills by regularly practicing drills, including perioperative difficult/failed airway management, and updating their knowledge of drugs, instruments, monitors, and even legal permissions.

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## CONFLICTS OF INTEREST

No potential conflict of interest relevant to this article was reported.

## DATA AVAILABILITY STATEMENT

Data sharing not applicable to this article as no datasets were generated or analyzed during the current study.

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## REFERENCES

1. Kim AM, Park JH, Kang S, Yoon TH, Kim Y. An ecological study of geographic variation and factors associated with cesarean section rates in South Korea. *BMC Pregnancy Childbirth* 2019; 19: 162.
2. Chien P. Global rising rates of caesarean sections. *BJOG* 2021; 128: 781-2.
3. Practice guidelines for obstetric anesthesia: an updated report by the American Society of Anesthesiologists Task Force on Obstetric Anesthesia and the Society for Obstetric Anesthesia and Perinatology. *Anesthesiology* 2016; 124: 270-300.
4. Mhyre JM, Sultan P. General anesthesia for cesarean delivery: occasionally essential but best avoided. *Anesthesiology* 2019; 130: 864-6.
5. Hawkins JL, Koonin LM, Palmer SK, Gibbs CP. Anesthesia-related deaths during obstetric delivery in the United States, 1979-1990. *Anesthesiology* 1997; 86: 277-84.
6. Ikeda T, Kato A, Bougaki M, Araki Y, Ohata T, Kawashima S, et al. A retrospective review of 10-year trends in general anesthesia for cesarean delivery at a university hospital: the impact of a newly launched team on obstetric anesthesia practice. *BMC Health Serv Res* 2020; 20: 421.
7. Cobb BT, Lane-Fall MB, Month RC, Onuoha OC, Srinivas SK, Neuman MD. Anesthesiologist specialization and use of general anesthesia for cesarean delivery. *Anesthesiology* 2019; 130: 237-46.
8. Traynor AJ, Aragon M, Ghosh D, Choi RS, Dingmann C, Vu Tran Z, et al. Obstetric anesthesia workforce survey: a 30-year update. *Anesth Analg* 2016; 122: 1939-46.
9. Guglielminotti J, Landau R, Li G. Adverse events and factors associated with potentially avoidable use of general anesthesia in cesarean deliveries. *Anesthesiology* 2019; 130: 912-22.
10. Hawkins JL, Chang J, Palmer SK, Gibbs CP, Callaghan WM. Anesthesia-related maternal mortality in the United States: 1979-2002. *Obstet Gynecol* 2011; 117: 69-74.
11. Ring L, Landau R, Delgado C. The current role of general anesthesia for cesarean delivery. *Curr Anesthesiol Rep* 2021; 11: 18-27.
12. Chen Y, Liu W, Gong X, Cheng Q. Comparison of effects of general anesthesia and combined spinal/epidural anesthesia for cesarean delivery on umbilical cord blood gas values: a double-blind, randomized, controlled study. *Med Sci Monit* 2019; 25: 5272-9.
13. Robbins LS, Blanchard CT, Biasini FJ, Powell MF, Casey BM, Tita AT, et al. General anesthesia for cesarean delivery and childhood neurodevelopmental and perinatal outcomes: a secondary analysis of a randomized controlled trial. *Int J Obstet Anesth* 2021; 45: 34-40.
14. Park JI, Park SH, Kang MS, Kang GW, Kim ST. Evaluation of changes in anesthetic methods for cesarean delivery: an analysis for 5 years using the big data of the Korean Health Insurance Review and Assessment Service. *Anesth Pain Med (Seoul)* 2020; 15: 305-13.
15. Park SH, Kim DJ, Kim WY, Kim JH, Lee YS, Park YC. Clinical evaluation of anesthesia for cesarean section at tertiary medical center: retrospective study for 5 years (2009-2013). *Anesth Pain Med* 2016; 11: 49-54.
16. Johnson RV, Lyons GR, Wilson RC, Robinson AP. Training in obstetric general anaesthesia: a vanishing art? *Anaesthesia* 2000; 55: 179-83.
17. Searle RD, Lyons G. Vanishing experience in training for obstetric general anaesthesia: an observational study. *Int J Obstet Anesth* 2008; 17: 233-7.
18. Palanisamy A, Mitani AA, Tsen LC. General anesthesia for cesarean delivery at a tertiary care hospital from 2000 to 2005: a retrospective analysis and 10-year update. *Int J Obstet Anesth* 2011; 20: 10-6.
19. Odor PM, Bampoe S, Moonesinghe SR, Andrade J, Pandit JJ, Lucas DN; Pan-London Perioperative Audit and Research Network (PLAN), for the DREAMY Investigators Group. General anaesthetic and airway management practice for obstetric surgery in England: a prospective, multicentre observational study. *Anaesthesia* 2021; 76: 460-71.
20. Desai N, Wicker J, Sajayan A, Mendonca C. A survey of practice of rapid sequence induction for caesarean section in England. *Int J Obstet Anesth* 2018; 36: 3-10.
21. Sumikura H, Niwa H, Sato M, Nakamoto T, Asai T, Hagihira S. Rethinking general anesthesia for cesarean section. *J Anesth* 2016; 30: 268-73.
22. Kosaka Y, Takahashi T, Mark LC. Intravenous thiobarbiturate anesthesia for cesarean section. *Anesthesiology* 1969; 31: 489-506.
23. Abouleish E, Abboud T, Lechevalier T, Zhu J, Chalian A, Alford K. Rocuronium (Org 9426) for caesarean section. *Br J Anaesth* 1994; 73: 336-41.
24. Sabetian G, Zand F, Mirhadi F, Hadavi MR, Asadpour E, Dehghanpisheh L, et al. Adequacy of maternal anesthesia depth with two sodium thiopental doses in elective caesarean section: a randomized clinical trial. *BMC Anesthesiol* 2021; 21: 201.
25. Odor PM, Bampoe S, Lucas DN, Moonesinghe SR, Andrade J, Pandit JJ; Pan-London Peri-operative Audit and Research Network (PLAN), for the DREAMY Investigators Group. Incidence

- of accidental awareness during general anaesthesia in obstetrics: a multicentre, prospective cohort study. *Anaesthesia* 2021; 76: 759-76.
26. Park HS, Kim YS, Kim SH, Jeon AR, Kim SE, Choi WJ. Comparison of electroencephalogram between propofol- and thiopental-induced anesthesia for awareness risk in pregnant women. *Sci Rep* 2020; 10: 6192.
  27. Duggal K. Propofol should be the induction agent of choice for caesarean section under general anaesthesia. *Int J Obstet Anesth* 2003; 12: 275-6.
  28. Russell R. Propofol should be the agent of choice for caesarean section under general anaesthesia. *Int J Obstet Anesth* 2003; 12: 276-9.
  29. Tumukunde J, Lomangisi DD, Davidson O, Kintu A, Joseph E, Kwizera A. Effects of propofol versus thiopental on Apgar scores in newborns and peri-operative outcomes of women undergoing emergency cesarean section: a randomized clinical trial. *BMC Anesthesiol* 2015; 15: 63.
  30. Nayar R, Sahajanand H. Does anesthetic induction for cesarean section with a combination of ketamine and thiopentone confer any benefits over thiopentone or ketamine alone? A prospective randomized study. *Minerva Anesthesiol* 2009; 75: 185-90.
  31. Moradkhani M, Hejri P, Nadri S, Beiranvand S. Effects of ADJUVANT ketamine on induction of anesthesia for the cesarean section. *Curr Rev Clin Exp Pharmacol* 2021; 16: 197-200.
  32. McGuigan PJ, Shields MO, McCourt KC. Role of rocuronium and sugammadex in rapid sequence induction in pregnancy. *Br J Anaesth* 2011; 106: 418-9; author reply 419-20.
  33. Pühringer FK, Kristen P, Rex C. Sugammadex reversal of rocuronium-induced neuromuscular block in caesarean section patients: a series of seven cases. *Br J Anaesth* 2010; 105: 657-60.
  34. Moya F, Kvisselgaard N. The placental transmission of succinylcholine. *Anesthesiology* 1961; 22: 1-6.
  35. Vanner R. Cricoid pressure. *Int J Obstet Anesth* 2009; 18: 103-5.
  36. Tsen LC, Bateman BT. Anesthesia for cesarean delivery. In: Chestnut's obstetric anesthesia: principles and practice. 6th ed. Edited by Chestnut DH, Wong CA, Tsen LC, Ngan Kee WD, Beilin Y, Mhyre JM, et al.: Amsterdam, Elsevier. 2019, pp 568-626.
  37. Morgan M. The confidential enquiry into maternal deaths. *Anaesthesia* 1986; 41: 689-91.
  38. Priebe HJ. Cricoid pressure: an expert's opinion. *Minerva Anesthesiol* 2009; 75: 710-4.
  39. Hansel J, Rogers AM, Lewis SR, Cook TM, Smith AF. Videolaryngoscopy versus direct laryngoscopy for adults undergoing tracheal intubation. *Cochrane Database Syst Rev* 2022; 4: CD011136.
  40. Browning RM, Rucklidge MW. Tracheal intubation using the Pentax Airway Scope videolaryngoscope following failed direct laryngoscopy in a morbidly obese parturient. *Int J Obstet Anesth* 2011; 20: 200-1.
  41. Howle R, Onwochei D, Harrison SL, Desai N. Comparison of videolaryngoscopy and direct laryngoscopy for tracheal intubation in obstetrics: a mixed-methods systematic review and meta-analysis. *Can J Anaesth* 2021; 68: 546-65.
  42. Timmermann A. Supraglottic airways in difficult airway management: successes, failures, use and misuse. *Anaesthesia* 2011; 66 Suppl 2: 45-56.
  43. Frerk C, Mitchell VS, McNarry AF, Mendonca C, Bhagrath R, Patel A, et al. Difficult Airway Society Intubation Guidelines Working Group. Difficult Airway Society 2015 guidelines for management of unanticipated difficult intubation in adults. *Br J Anaesth* 2015; 115: 827-48.
  44. Apfelbaum JL, Hagberg CA, Connis RT, Abdelmalak BB, Agarkar M, Dutton RP, et al. 2022 American Society of Anesthesiologists practice guidelines for management of the difficult airway. *Anesthesiology* 2022; 136: 31-81.
  45. Mushambi MC, Kinsella SM, Popat M, Swales H, Ramaswamy KK, Winton AL, et al. Obstetric Anaesthetists' Association; Difficult Airway Society. Obstetric Anaesthetists' Association and Difficult Airway Society guidelines for the management of difficult and failed tracheal intubation in obstetrics. *Anaesthesia* 2015; 70: 1286-306.
  46. Delgado C, Ring L, Mushambi MC. General anaesthesia in obstetrics. *BJA Educ* 2020; 20: 201-7.
  47. Lyons G, Macdonald R. Awareness during caesarean section. *Anaesthesia* 1991; 46: 62-4.
  48. Bogod D, Plaat F. Be wary of awareness--lessons from NAP5 for obstetric anaesthetists. *Int J Obstet Anesth* 2015; 24: 1-4.
  49. Mankowitz E, Brock-Utne JG, Downing JW. Nitrous oxide elimination by the newborn. *Anaesthesia* 1981; 36: 1014-6.
  50. Yoo KY, Lee JC, Yoon MH, Shin MH, Kim SJ, Kim YH, et al. The effects of volatile anesthetics on spontaneous contractility of isolated human pregnant uterine muscle: a comparison among sevoflurane, desflurane, isoflurane, and halothane. *Anesth Analg* 2006; 103: 443-7.
  51. Kimizuka M, Tokinaga Y, Azumaguchi R, Hamada K, Kazuma S, Yamakage M. Effects of anesthetic agents on contractions of the pregnant rat myometrium in vivo and in vitro. *J Anesth* 2021; 35: 68-80.
  52. Mhyre JM, Riesner MN, Polley LS, Naughton NN. A series of anesthesia-related maternal deaths in Michigan, 1985-2003. *Anesthesiology* 2007; 106: 1096-104.