Comparison of recovery profiles of propofol and sevoflurane anesthesia with bispectral index monitoring in percutaneous nephrolithotomy

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**Background:** The aim of the study was to evaluate the comparative effects of propofol infusion versus sevoflurane for maintenance of anesthesia with respect to hemodynamics, recovery characteristics, nausea and vomiting in patients undergoing percutaneous nephrolithotomy.

**Methods:** Forty American Society of Anesthesiologists physical status I-II patients, aged between 22 and 65 years were randomly divided to receive either intravenous anesthesia with propofol (group P) or sevoflurane (group S). Cardiovascular variables, peripheral oxygen saturation (SpO₂), end-tidal carbon dioxide (ETCO₂), bispectral index (BIS) and train-of-four (TOF) values were recorded at intervals throughout the procedure. Time to spontaneous respiration, eye opening, extubation, obey commands, hand squeezing, Aldrete Score > 9 and the incidence of postoperative nausea and vomiting were recorded.

**Results:** Early recovery times [spontaneous respiration (P = 0.002), eye opening (P = 0.006), extubation (P = 0.013), obey commands (P < 0.05), hand squeezing (P = 0.005)] were significantly longer in group P. The incidence of vomiting was significantly higher in group S (P < 0.05). Hemodynamic parameters, levels of SpO₂, ETCO₂, and BIS and TOF values were not significantly different between the groups (P > 0.05).

**Conclusions:** The present study which adjusted sevoflurane concentration and propofol infusion rate according to BIS values revealed that maintenance of anesthesia with sevoflurane is associated with faster recovery than anesthesia with propofol. Propofol resulted in a significantly lower incidence of postoperative nausea and vomiting. Hemodynamic parameters and levels of SpO₂ and ETCO₂ were comparable between the groups during percutaneous nephrolithotomy. (Korean J Anesthesiol 2013; 64: 223-228)

**Key Words:** Percutaneous nephrolithotomy, Propofol, Sevoflurane.
Propofol and sevoflurane for PCNL

Introduction

Percutaneous nephrolithotomy (PCNL) is a less invasive alternative procedure to open surgery for removal of medium-sized or larger renal calculi from the urinary tract using a nephroscope into the kidney through a track created in the patient’s back. This procedure is advantageous in that it reduces the duration of hospitalization, results in less morbidity and postoperative pain, and minimal scarring [1].

However, hemodynamic changes may occur during the procedure because continuous irrigation of the kidneys may cause excessive fluid absorption and increase hydrostatic pressure [2,3]. Also, continuous irrigation makes monitoring the amount of bleeding not possible. Significant bleeding may occur during the procedure and maintenance of hemodynamic stability may be too difficult. For this reason, choosing an anesthetic agent is very critical, because it must have a minimal effect on hemodynamic parameters. There are a limited number of studies about hemodynamic responses of anesthesia on this procedure. Therefore, the present study investigated the hemodynamic changes and recovery of this procedure and the results were compared with other studies [4-7].

Hemodynamic changes may also be caused by inadequate anesthesia levels in patients undergoing operation. It is important to know which is responsible for this condition: significant bleeding or inadequate anesthesia. If it is due to bleeding crystalloid, colloid or blood replacement should be performed. However, if it is from inadequate anesthesia, the depth of anesthesia should be reevaluated.

During the last decade, an increasing number of monitoring systems have been designed to estimate the depth of anesthesia. One of these systems is the bispectral index (BIS) monitor; different studies have shown that BIS may help to assess the hypnotic component of anesthesia, reduce drug consumption and shorten recovery time compared with the standard practice protocol [8-11].

Propofol and sevoflurane are anesthetic agents known to have minimal effects on hemodynamic parameters [12-16]. The aim of the study was to determine a more suitable anesthetic for better maintenance in terms of hemodynamic characteristics and recovery profile for the duration of PCNL which bears the difficulty of uncontrollable bleeding for anesthesiologists. Also, unlike previous studies, this study was intended to compare propofol and sevoflurane by adjusting the depth of anesthesia according to BIS.

Materials and Methods

After obtaining Institutional Medical Ethics Committee approval and written informed consent, 40 American Society of Anesthesiologists physical status I-II patients aged between 22 and 65 years undergoing general anesthesia for elective PCNL were enrolled in the study and randomly assigned (by opening of a sealed envelope) into two groups receiving sevoflurane (group S) or propofol infusion (group P) for maintenance of anesthesia. Patients who had cardiovascular, renal, hepatic or endocrine disorders were excluded from the study. In the operating room after establishment of standard monitoring, arterial and venous cannulation, the skin of the forehead was degreased with 70% isopropanol; a BIS electrode (Aspect Medical Systems, Norwood, MA, USA) was positioned as recommended by the manufacturer. Impedances were measured to ensure optimal electrode contact defined as ≤7.5 kΩ for the BIS as required by the manufacturer. EEG was continuously recorded using a BIS monitor (Aspect 2000 XP BIS monitor, P/N 185-0070, Host Rev. 3.12, Aspect Medical Systems, Newton, MA, USA). All patients were then premedicated with midazolam 0.03 mg/kg and fentanyl 1 µg/kg intravenously ten minutes before induction. Propofol (2 mg/kg) was used for induction in all patients.

After induction, skin surface electrodes for neurostimulation were placed on the volar forearm along the course of the ulnar nerve, close to the proximal wrist crease to stimulate adductor pollicis. The other ends of electrodes were connected to a respective train-of-four (TOF) Guard acceleromyography monitor (TOF Watch SX, Organon-Teknika, Boxtel, Netherlands). Nerve was stimulated with TOF stimulation (a series of four twitches in 2 sec, 2 Hz frequency, each 0.2 ms long) every 12 seconds after loss of the eyelash reflex. A current intensity of 50 mA was used. Intubation was performed when all four TOF responses from the adductor pollicis muscle were disappeared. Vecuronium (0.1 mg/kg for tracheal intubation; thereafter 2 mg as needed) was used as a neuromuscular blocking agent for both groups. After this, all patients were manually ventilated by an oxygen mask for 2 minutes and then were intubated. Patients in each group received intermittent positive pressure ventilation with N₂O 50% in an oxygen mixture via a rebreathing system at a tidal volume of 6–8 ml/kg and a frequency of 12 per minute with an end tidal CO₂ (ETCO₂) target of 30–35 mmHg.

Continuous monitoring included heart rate, invasive measurements of systolic arterial blood pressure, oxygen saturation (SpO₂), ETCO₂, BIS and TOF values. The criteria for supplementary doses of vecuronium were assessed according to TOF monitoring.

For maintenance of anesthesia, propofol infusion at 3–12 mg/kg/hr or 0.5–2 vol% of sevoflurane was administered. The concentration of sevoflurane used and the infusion rate of propofol were adjusted according to an EEG target value of 40–60 for BIS. BIS values were continuously recorded at intervals of 10 minutes. If BIS values were over 60, the infusion rate of propofol or concentration of sevoflurane was gradually...
increased. If was under 40, the infusion rate of propofol or concentration of sevoflurane was decreased. In the case of more than 20% reduction in the mean arterial pressure from baseline levels, the infusion rate of crystalloid solution was increased. If this was not sufficient, the infusion rate of propofol or the concentration of sevoflurane was reduced. If hypotension was due to bleeding, colloids and blood were administered. Severe hypotension was finally treated with a vasopressor.

About fifteen minutes before the end of surgery, sevoflurane and propofol were reduced to facilitate rapid emergence from anesthesia. They were adjusted to a BIS value of 70.

Hemodynamic parameters, SpO₂, and ETCO₂ levels were noted before induction, before and after intubation, before prone position and at 5, 10, 30, 60, 90 and 120 minutes after prone position, and before extubation and at 5, 10, 30, 60, 90 and 120 minutes after extubation. BIS values were measured and recorded every 10 minutes. The time from discontinuation of drugs to eye opening in response to verbal stimulus, hand squeezing, spontaneous ventilation, tracheal extubation, stating name, stating date of birth, time to Aldrete score > 9, nausea and vomiting were recorded.

Anesthetic agents were discontinued when the skin incision was being sutured. After suturation, the patient was turned into the supine position. When neuromuscular recovery was completed and adequate depth of breathing was reached (after the return of adequate ventilator drive, tidal volume > 8 ml/kg, respiratory rate > 12/min, normal breathing patterns and good oxygenation [SpO₂ > 98%]) and the patient could obey the verbal commands ("open your eyes", "elevate your head"; commanded every 30 seconds), tracheal extubation was done. The adequacy of recovery following reversal was assessed according to TOF monitoring. A TOF ratio of greater than 90% was correlated with the criteria of adequate clinical recovery. Nausea and vomiting were evaluated every fifteen minutes after extubation according to the score of nausea and vomiting (0 = no emetic symptoms, 1 = nausea, 2 = vomiting). Ondansetron (0.1 mg/ kg i.v.) was only used for patients with a score of 2. Warmed blankets were used in the recovery room. For postoperative pain relief, meperidine 0.5 mg/kg was administered to all patients intravenously just before the anesthetic agents were discontinued as part of the standard analgesic protocol of our institute.

NCSS for windows NCSS 2007 software (NCSS LLC, Kaysville, UT, USA) was used for statistical analysis. Repeated measures of ANOVA were used to determine differences overtime, and between multiple time periods for each anesthetic agent. Post hoc Newman-Keuls multiple comparison analysis was used. An unpaired t-test was used in each anesthetic agent group. A chi-square test was performed for the evaluation of available data. The statistically significant level was established at 0.05.

**Results**

There were no significant differences between the two groups regarding age, body weight, gender and duration of surgery (P > 0.05) (Table 1). The measurements of hemodynamic parameters were statistically similar between group P and group S (P > 0.05). A significant rise and fall was observed in each group during the measurement of mean arterial pressure and heart rate in the course of operative period. Although there were statistically significant differences between baseline values and intraoperative hemodynamic parameters, all values were within acceptable limits. All significant changes in groups P and S in addition to the differences compared to baseline values of each group are graphically represented in Fig. 1 and 2. BIS values were extremely stable and similar between the two groups throughout the study (P ≥ 0.05).

There were significant differences in recovery time after anesthesia with sevoflurane versus propofol. Early recovery time after sevoflurane anesthesia was significantly shorter than propofol infusion except Aldrete recovery scores (Table 2).

![Fig. 1. Comparisons of the changes in mean arterial blood pressure (*indicates P < 0.05), and differences compared to baseline values of each group (‡ indicates P < 0.05 for group S, †indicates P < 0.05 for group P). S: Sevoflurane, P: Propofol, ind: induction, int: intubation, ext: extubation, min: minute.](www.ekja.org)

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**Table 1. Characteristics of Patients**

| Characteristic     | Group S (n = 20) | Group P (n = 20) | P value |
|--------------------|-----------------|-----------------|---------|
| Age (yr)           | 41.4 ± 11.7     | 42.3 ± 10.0     | 0.796   |
| Weight (kg)        | 80 ± 14.4       | 72.5 ± 13.7     | 0.104   |
| Sex (M/F)          | 12/8            | 9/11            | 0.342   |
| OP time (min)      | 157.5 ± 48.5    | 159 ± 45.7      | 0.92    |

There were no significant differences between the groups. S: Sevoflurane, P: Propofol. OP time: duration of operation.
The incidence of nausea and vomiting was significantly higher in group S compared to group P ($P < 0.05$). One patient in group P and 4 patients in group S had nausea. Vomiting was not observed in any patient in group P but in 3 patients in group S; these patients were treated with ondansetron 0.1 mg/kg intravenously. VAS score was ≤ 4 in all patients in both groups.

**Discussion**

Percutaneous nephrolithotomy is widely used for the treatment of renal stones. It is a relatively less invasive endoscopic intervention compared to open surgery [17]. Although it is accepted as a safer method, irrigation of the kidneys and drugs used for anesthesia may cause hemodynamic disturbances [4-7].

A recent study was designed to determine cognitive and clinical outcomes after sevoflurane compared with propofol-based anesthesia for on-pump cardiac surgery. Patients assigned to sevoflurane-based anesthesia were associated with better results in all cognitive tests than those in the propofol group. There was no difference in general clinical outcome [14]. Robinson et al. [18] reported a more rapid recovery following sevoflurane anesthesia than that of propofol. Gupta et al. [19] reported that no time difference was found in eye opening time between sevoflurane and propofol in their systematic review, but the time period to obeying commands was faster in the sevoflurane group. Postoperative nausea and vomiting were significantly greater with sevoflurane compared with propofol. In the present study, almost all recovery events following anesthesia including eye opening, hand squeezing, spontaneous ventilation, extubation, stating name and stating date of birth were achieved earlier in the sevoflurane group. Wandel et al. [20] reported that following general anesthesia, patients who received sevoflurane for maintenance could be extubated earlier and regained cognitive functions much earlier than those on propofol infusion. In some other studies, investigators observed a shorter recovery time in patients given sevoflurane anesthesia, which is similar to the results of our study [21,22]. Samantaray and Rao [23] observed that maintenance of anesthesia with sevoflurane was associated with faster recovery than propofol anesthesia in patients who underwent spine surgery, and sevoflurane was associated with postoperative nausea and vomiting as with any other inhalational anesthetic.

According to the results of a previous study, there was a significant decrease in heart rate during PCNL in the propofol group [4]. This decrease can be related to the use of alfentanil. In the present study, only $\text{N}_2\text{O}$ was used and no decrease in heart rate was observed in the propofol group. Opioid supplementation or increased delivery of anesthetic agents instead of $\text{N}_2\text{O}$ has resulted in delayed awareness and recovery [24]. Several investigators found that recovery was more rapid, the incidence of postoperative nausea and vomiting was lower, and peroperative hemodynamic stability was better in the propofol group compared to the sevoflurane group [25-27]. A recent study compared the incidence and degree of postoperative nausea and vomiting in patients who received general anesthesia with propofol and those with sevoflurane. The propofol group had a statistically lower incidence of post-operative nausea and vomiting [15]. This finding is in accordance with the current study. Small and clinically unimportant differences were found by some investigators in recovery time between the two groups of patients who received propofol or sevoflurane [28,29]. On the contrary, Husedžinović et al. [30] did not find any significant difference in hemodynamic parameters between propofol and sevoflurane groups in patients undergoing open cholecystectomy.

In endoscopic procedures, it is obligatory to control blood
pressure in order to reduce bleeding and thus improve the surgeon’s view of the operative field. As endoscopic procedures end rapidly after removal of the endoscope, the anesthesiologist has to ensure an adequate level of anesthesia until completion of endoscopic maneuvers but at the same time must prevent prolonged recovery time at the end of the surgery. In the present study, an adequate level of anesthesia and hemodynamic stability were achieved simply by the BIS monitor system. Also, BIS monitor system helped to shorten the duration of recovery. In previous studies which compared the effects of sevoflurane and propofol infusion anesthesia on hemodynamic changes and recovery, BIS monitors were not used and the depth of anesthesia was not monitored objectively [4,8,14,18-23,25-30]. However, some of the changes in cardiovascular parameters might have been associated with the depth of anesthesia. Also, in some patients, the level of anesthesia might be deeper at the end of the surgery and the recovery time might be longer. In the current study, the depth of anesthesia was maintained at the same level for the two anesthetic agents and the effects of the anesthetics on hemodynamic parameters and recovery were assessed under similar circumstances.

Introduction of more rapid and short-acting volatile anesthetics and intravenous anesthetics has allowed anesthesiologists to achieve a more consistent recovery profile that facilitates fast-tracking after general anesthesia. Anesthetic techniques that optimize intraoperative surgical conditions while providing rapid, early recovery have assumed increased importance.

Cicek et al. [5] compared the effects of propofol-alfentanil and propofol-remifentanil anesthesia on hemodynamic factors and recovery characteristics during PCNL. Both groups provided stable hemodynamic parameters during PCNL, whereas time to recovery of spontaneous ventilation, extubation and eye opening were significantly shorter in the propofol-remifentanil group than the propofol-alfentanil group. In a previous report, inhalation anesthesia with sevoflurane and N₂O, and total intravenous anesthesia with propofol and alfentanil was found to be effective in controlling mean arterial pressures during PCNL. The mean heart rate was lower during PCNL in the propofol-alfentanil-N₂O group compared with the sevoflurane group (P < 0.01). The mean systolic and diastolic arterial pressures were not different between both groups at any stage of measurement (P < 0.05). In the sevoflurane group, the concentrations of renin, aldosterone and adrenocorticotrophic hormone were significantly higher compared with the propofol-alfentanil-N₂O anesthesia group (P < 0.05) [7]. In our study, although there were changes in intraoperative hemodynamic parameters within the groups, all the values were in the acceptable limits and there was no significant difference between the groups.

In conclusion, recovery time of sevoflurane anesthesia was significantly shorter than propofol infusion anesthesia in the current study, and the incidence of nausea and vomiting was significantly higher in the sevoflurane group compared with the propofol infusion group. These results collectively show that both sevoflurane and propofol are convenient anesthetic techniques for percutaneous nephrolithotomy. Both anesthetics achieved circulatory stability, rapid titration in relation to clinical needs, acceptable surgical field and fast recovery. Faster recovery time of sevoflurane is especially useful for endoscope treatments in which the surgical procedure ends abruptly.

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