

The Thai Anesthesia Incidents (THAI Study) of Anesthetic Risk Factors related to Perioperative Death and Perioperative Cardiovascular Complications in Intracranial Surgery

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Objective: To evaluate the correlations between anesthetic risk factors and perioperative cardiovascular complications as well as perioperative death within 72 hours.

Material and Method: This case controlled took the data from the Thai Anesthesia Incidents Study (THAI Study), a prospective multi-centered registry of anesthesia in Thailand. The authors included all the patients who received intracranial surgery from 20 hospitals throughout Thailand. The present study was divided into two groups and focused on anesthetic factors that possibly related to perioperative cardiovascular complications or perioperative death. The statistical analysis were Chi Square test and logistic regression model with the statistical significance if p -value < 0.05 demonstrated in Odds ratio (OR) and 95% confidence interval.

Results: From the 7,430 patients, there were 63 patients (0.85%) with perioperative cardiovascular complication. The American Society of Anesthesiologists (ASA) physical status 3-5 (OR 5.77, 95%CI 2.33-14.27) and the absence of anesthesiologists (OR 2.19, 95%CI 1.06-4.54) had statistical correlation with the cardiovascular complication. Eighty-four patients (1.13%) who died within 72 hours post operatively were found. The ASA physical status 3-5 (OR 10.14, 95%CI 3.42-30.02), the emergency circumstance (OR 3.55, 95%CI 1.31-9.60), and the absence of endtidal carbondioxide monitor (OR 2.27, 95%CI 1.26-4.09) had statistical correlation with the perioperative death.

Conclusion: Predictors of perioperative cardiovascular complications in intracranial surgical patients were ASA physical status 3-5 and absence of certified board anesthesiologists. Risk factors of perioperative death were ASA physical status 3-5, emergency condition, and absence or no monitoring of capnometer.

Keywords: Anesthesia, Risk factor, Cardiovascular complication, Perioperative death, Intracranial surgery, Certified board anesthesiologist

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The disease prognosis, quality of life, and survival rate are all influenced by perioperative compli-

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cations. In 2005, a multi-centered study with a title of the Thai Anesthesia Incidents Study (THAI Study) of anesthetic adverse outcomes^(1,2) was accomplished by Thai anesthesiologists. The perioperative cardiovascular complications (e.g. myocardial ischemia and

myocardial infarction) were found in 2.7/10000. Neurovascular complications (e.g. coma, cerebrovascular accident, seizure) were found in 4.8/10000, respiratory complications (e.g. hypoxia) were found in 31.9/10000. Dead within 24 hours postoperatively was found in 28.3/10000. The incidences of each complication occurred frequently and were severe⁽¹⁾. The preventive strategy can be developed from anesthetic risk analysis, especially on those jeopardy patients, the neurosurgical patients.

The perioperative risk factors are generally concerned by surgical risk factors, for example, postoperatively rebleeding⁽³⁾, intracerebral hemorrhage, pre-operative vasospasm⁽³⁻⁵⁾, and the duration and strengthening of intra-operative brain retraction⁽⁶⁾. The correlation analysis between anesthetic techniques, monitoring, special technique in anesthesia, medications including anesthetic factors, and perioperative complications have been less published even though those patients were taken care of by the majority of anesthesiologists and nurse anesthetists. Therefore, the authors performed this case controlled study to evaluate the correlations between anesthetic risk factors and perioperative cardiovascular complications, and perioperative death for further improvement of anesthesia service.

Material and Method

This case controlled study took the data from the THAI Study under the process of quality assurance from February 1, 2003 to August 31, 2004. The authors included all the patients who received intracranial surgery from 20 hospitals throughout Thailand, seven university hospitals, five regional hospitals, four provincial hospitals, and four community hospitals. The authors focused on anesthetic factors that were possibly related to perioperative cardiovascular complications or perioperative death like age, sex, American Society of Anesthesiologist (ASA) physical status, emergency situation, anesthetic durations, preoperative respiratory conditions, preoperative cardiovascular conditions, pre-operative neurological conditions, preoperative hematological conditions, preoperative endocrine conditions, premedications, the presence of anesthesiologist in anesthetic team, the usage of steroid drugs preoperatively, invasive monitoring (arterial blood pressure, central venous pressure and pulmonary wedge pressure monitoring), endtidal carbon dioxide monitoring, special anesthetic techniques (hypotensive and hypothermic technique), induction agents, muscle relaxant, opioids, and inhalation agents.

The study design was achieved as follows.

1. To determine factors associated with the perioperative cardiovascular complication, the authors separated intracranial surgical patients into two groups by simple randomization with SPSS software version 11.5. The control group was composed of 189 patients who did not have perioperative cardiovascular complications. The other group (the study group) was composed of 63 patients with perioperative cardiovascular complications. The number of patients in the study group should not be less than 23. The sample size was calculated from the study of Akavipat P, et al⁽⁷⁾ who found that the ASA physical status 3-5 patients have a powerful correlation with post intracranial aneurysmal surgery complications and the incidence studied by Charuluxananan S et al⁽²⁾ ($p_1 = 0.89$, $p_2 = 0.50$, OR = 8.16, level of significance = 5%, power of the test = 80%). The definition of perioperative cardiovascular complications was the episode of myocardial injury, myocardial ischemia, and cardiac arrest diagnosed by clinical and confirmed by electrocardiogram and/or cardiac enzyme. A statistical analysis was used to identify anesthetic risk factors related to the complications.

2. To determine factors associated with the perioperative death, the authors separated intracranial surgical patients into two groups by simple randomization with SPSS software version 11.5. The control group was composed of 252 patients who survived at least 72 hours postoperatively. The study group was composed of 84 patients who died within 72 hours postoperatively. The number of patients in the study group should not be less than 23. The sample size was calculated from the study of Wolters U, et al⁽⁸⁾ who found that the ASA physical status 3-5 patients have a powerful correlation with perioperative death and the incidence studied by Charuluxananan S, et al⁽²⁾ ($p_1 = 0.94$, $p_2 = 0.49$, OR = 16.22, level of significance = 5%, power of the test = 80%). A statistical analysis was used to identify anesthetic risk factors related to perioperative death.

The quantitative data are shown as number, percent, and standard deviations. The qualitative data were analyzed by Chi-Square test with the statistical significance at p-value of less than 0.05. Then, the multiple logistic regression analysis was used to identify risk factors and the strength of the association was shown as Odds ratio with 95% confidence intervals.

Results

The seven thousand four hundred and thirty patients were categorized into perioperative cardiovascular complications and perioperative death within

Table 1. Demographic and administrative characteristics of intracranial surgical patients

Data	Total intracranial surgical patients	Perioperative cardiovascular complication	Perioperative death
Sex			
Male	4473 (60.2)	46 (73.0)	61 (72.6)
Female	2957 (39.8)	17 (27.0)	23 (27.4)
Age (year)	41.6 ± 21.4	42.2 ± 20.6	39.3 ± 20.0
Weight (Kg)	54.3 ± 16.7	54.5 ± 16.5	55.4 ± 15.4
Anesthetic duration (minute)	165.6 ± 132.5	121.6 ± 91.1	109.7 ± 80.8
ASA physical status			
1	998 (13.4)	2 (3.2)	1 (1.2)
2	2722 (36.6)	5 (7.9)	4 (4.8)
3	2616 (35.2)	16 (25.4)	24 (28.6)
4	1022 (13.8)	29 (46.0)	46 (54.8)
5	72 (1.0)	11 (17.5)	9 (10.7)

Values shown as frequency (%), mean ± SD

Table 2. Anesthetic characteristics of intracranial surgical patients

Anesthetic factors	Total intracranial surgical patients	Perioperative cardiovascular complication	Perioperative death
Absence of anesthesiologists	1695 (22.8)	24 (38.1)	30 (35.7)
Invasive monitoring	2423 (32.6)	21 (33.3)	21 (25.0)
Endtidal carbon dioxide monitoring	4456 (60.0)	27 (42.9)	29 (34.5)
Anesthetic technique			
- Hypotensive	88 (1.2)	3 (4.8)	3 (3.6)
- Hypothermia	12 (0.2)	0 (0)	1 (1.2)
Anesthetic agents			
- Pentothal	1157 (15.6)	7 (11.1)	7 (8.3)
- Propofol	4972 (66.9)	26 (41.3)	39 (46.4)
- Ketamine	19 (0.3)	0 (0)	0 (0)
- Midazolam	1238 (16.7)	16 (25.4)	21 (25.0)
- Succinyl Choline	848 (11.4)	1 (1.6)	2 (2.4)
- Pancuronium	2537 (34.1)	25 (39.7)	31 (36.9)
- Atracurium	933 (12.6)	6 (9.5)	6 (7.1)
- Cis-atracurium	482 (6.5)	1 (1.6)	0 (0)
- Vecuronium	2806 (37.8)	27 (42.9)	41 (48.8)
- Rocuronium	560 (7.5)	2 (3.2)	2 (2.4)
- Nitrous oxide	5517 (74.3)	39 (61.9)	54 (64.3)
- Halothane	247 (3.3)	2 (3.2)	3 (3.6)
- Isoflurane	6195 (83.4)	49 (77.8)	67 (79.8)
- Sevoflurane	653 (8.8)	1 (1.6)	2 (2.4)
- Morphine	340 (4.6)	0 (0)	0 (0)
- Fentanyl	6749 (90.8)	54 (85.7)	75 (89.3)
- Prostigmine	3551 (47.8)	11 (17.5)	17 (20.2)

Values shown as frequency (%)

72 hours as a demographic data in Table 1.

The emergency cases were found in 4128 (55.6%) with 78 (92.9%) deaths and 56 (88.9%) had perioperative cardiovascular complications. There were 96.4% in the perioperative death group and 93.7% in the perioperative cardiovascular complication group had abnormal pre-anesthetic conditions which were assessed by anesthetic personnel. The patients who received benzodiazepine for premedication were found in 6.3%, steroid 6.3% in the perioperative cardiovas-

cular complication group and 9.5% and 3.6% in the perioperative death orderly. The presence of anesthesiologists, monitoring, special anesthetic technique and anesthetic agents are shown in Table 2.

From univariate analysis, factors were related to perioperative cardiovascular complications and perioperative death. The results that had statistical significance are illustrated in Table 3-4.

Then the authors used the logistic regression to find the correlation factors without any confound-

Table 3. Anesthetic factors related to perioperative cardiovascular complications in intracranial surgery (univariate analysis)

Anesthetic factors	Odds ratio	95% Confidence interval	p-value
Female	2.23	1.19-4.18	0.011
ASA physical status 3-5	8.44	3.67-19.76	<0.001
Emergency case	6.82	2.96-15.75	<0.001
Absence of anesthesiologist	3.14	1.66-5.94	<0.001
Absence of endtidal carbondioxide monitor	1.98	1.11-3.53	0.019
Non nitrous oxide usage	1.91	1.04-3.51	0.041

Table 4. Anesthetic factors related to perioperative death in intracranial surgery analyzed (univariate analysis)

Anesthetic factors	Odds ratio	95% Confidence interval	p-value
Male	1.89	1.10-3.25	0.022
ASA physical status 3-5	15.80	6.19-40.33	<0.001
Anesthetic duration less than 6 hours	4.32	1.00-18.67	0.032
Emergency case	10.74	4.51-25.55	<0.001
No midazolam premedication	6.39	1.50-27.22	0.004
Absence of anesthesiologist	2.36	1.37-4.08	0.002
No steroid usage	6.03	1.82-19.94	0.001
Absence of endtidal carbondioxide monitor	2.74	1.64-4.59	<0.001
Non nitrous oxide usage	2.09	1.22-3.58	0.009

Table 5. Anesthetic factors which related to perioperative cardiovascular complications and perioperative death (multivariate analysis)

Anesthetic factors	Odds ratio	95% Confidence interval	p-value
Cardiovascular complication			
ASA physical status 3-5	5.77	2.33-14.27	<0.001
Absence of anesthesiologist	2.19	1.06-4.54	0.034
Perioperative death			
ASA physical status 3-5	10.14	3.42-30.02	<0.001
Emergency case	3.55	1.31-9.60	0.013
Absence of endtidal carbondioxide monitor	2.27	1.26-4.09	0.006

ing. This is illustrated in Table 5. The stepwise multiple logistic regression was evaluated and initiated the equations as follows. $Y = 1.75 \text{ ASA physical status} + 0.79 \text{ anesthesiologist} - 4.023$ for the correlation of anesthetic risk factors and perioperative cardiovascular complications and $Y = 2.32 \text{ ASA physical status} + 1.27 \text{ emergency case} + 0.82 \text{ endtidal carbondioxide monitor} - 4.223$ for the correlation of anesthetic risk factors and perioperative death.

Discussion

Among the intracranial surgery in Thailand between February 2003 and August 2004, the incidence of perioperative cardiovascular complication and perioperative death were found in 0.85% and 1.13% by sequence. These incidences are possibly not comparable to the other sources because virtually all of the researches are studied in general surgery not particularly on every procedure that takes place in the intracranium. Aarabi B et al⁽⁹⁾ described the incidence of perioperative death in severe traumatic brain injury was 28%. Sperling MR et al⁽¹⁰⁾ found 0.85 in 1000 patient-year in epilepsy surgery. Marina G et al⁽¹¹⁾ found 3% in intracerebral aneurysm surgery and Fiorani P et al⁽¹²⁾ reported 0.7% after carotid endarterectomy and revealed 0.8% of perioperative cardiovascular complication.

In the present study, a large number of patients were categorized into the ASA physical status over 2, which defined the severity of disease. That was the reason why the incidence of perioperative death was above average similar to the previous report⁽¹³⁾. Furthermore, the emergency circumstance can increase mortality and morbidity⁽¹⁴⁾ but the process of pre-anesthetic evaluation can minimize and improve good surgical outcome⁽¹⁵⁻¹⁷⁾.

From univariate analysis, steroids generated a worse outcome but after being adjusted any confounding factors from multivariate analysis there was no statistical significance. Some advised steroids in craniopharyngioma⁽¹⁸⁾ and idiopathic increase intracranial pressure⁽¹⁹⁾, some did not^(20,21) and some claimed as the justification of avascular necrosis of the hip⁽²²⁾.

There was a strong relation between perioperative cardiovascular complications and perioperative death⁽²³⁾. The influence factor is the absence of anesthesiologists from the anesthetic team. In a preceding study they exemplified the same conclusion⁽²⁴⁻²⁶⁾ for example, Silber JH et al reported the upsurge of perioperative death from 2.5/1000 to 3.8/1000⁽²⁷⁾. Besides, from anesthetic personnel the medical equipment like

capnometer also plays an important role in anesthesia. Several countries consider it as a basic monitor^(28,29). Endtidal carbondioxide monitoring does not prevent the post operative complication but reduces the incidence by warning⁽³⁰⁾. The others asserted the intensity of morbidity and mortality for the nonappearance of effective monitors^(31,32).

Nowadays the authors confronted the limitation of resources, manpower, and financial statement. The alternative of anesthetic trained from other medical recruits, referral system and competency for each professional would have an advantage. However, the beneficial evaluation to the patients and national health management should be deliberated. In summary, the present study, as part of the Thai Anesthesia Incidents Study, revealed that high ASA physical status, emergency condition, absence of certified board anesthesiologists, and absence or no monitoring of capnometry were predictors of cardiovascular complications and or perioperative death in intracranial surgical patients.

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ปัจจัยทางวิสัญญีวิทยาที่เกี่ยวข้องกับการเสียชีวิตและภาวะแทรกซ้อนทางระบบหัวใจและหลอดเลือด ระหว่างและหลังการผ่าตัดกะโหลกศีรษะในโครงการศึกษาภาวะแทรกซ้อนทางวิสัญญีวิทยาในประเทศไทย

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วัตถุประสงค์: เพื่อประเมินความสัมพันธ์ของปัจจัยเสี่ยงด้านวิสัญญีวิทยาที่เกี่ยวข้องกับภาวะแทรกซ้อนทางระบบหัวใจและหลอดเลือด และภาวะการเสียชีวิตระหว่างและหลังการผ่าตัดภายใน 72 ชั่วโมง

วัสดุและวิธีการ: ทำการศึกษาย้อนหลังในผู้ป่วยทั้งหมดที่ได้รับการผ่าตัดภายในกะโหลกศีรษะที่ได้ จากฐานข้อมูลของ Thai Anesthesia Incidents Study (THAI Study) ที่ทำการศึกษาในโรงพยาบาลในระดับต่าง ๆ 20 แห่งทั่วประเทศ โดยการวิเคราะห์ แบบ case controlled 1:3 เพื่อหาปัจจัยเสี่ยงต่าง ๆ ทางด้านวิสัญญีวิทยา ที่อาจเกี่ยวข้องกับภาวะแทรกซ้อนทางระบบหัวใจและหลอดเลือด และหาปัจจัยเสี่ยงที่อาจเกี่ยวข้องกับภาวะการเสียชีวิตระหว่างและหลังการผ่าตัด โดยใช้หลักการวิเคราะห์ทางสถิติแบบ Chi Square test และ multiple logistic regression โดยมีนัยสำคัญทางสถิติเมื่อ $p\text{-value} < 0.05$ และแสดงผลในรูปของ Odds ratio และ 95% confidence intervals

ผลการศึกษา: ผู้ป่วยทั้งหมดจำนวน 7,430 คน เป็นผู้ป่วยที่มีภาวะแทรกซ้อนทางระบบหัวใจและหลอดเลือด จำนวน 63 คน (0.85%) พบว่าการจำแนกผู้ป่วยตาม American Society of Anesthesiologists ระดับ 3-5 (OR 5.77, 95%CI 2.33-14.27) และการไม่มีวิสัญญีแพทย์ในทีมผู้ให้ยาระงับความรู้สึก (OR 2.19, 95%CI 1.06-4.54) มีความสัมพันธ์กับการเกิดภาวะแทรกซ้อนอย่างมีนัยสำคัญ ส่วนผู้ป่วยที่เสียชีวิตระหว่างและหลังการผ่าตัดภายใน 72 ชั่วโมงนั้นมีจำนวน 84 คน (1.13%) พบว่าการจำแนกผู้ป่วยตาม American Society of Anesthesiologists ระดับ 3-5 (OR 10.14 95%CI 3.42-30.02) ภาวะชุกเฉิน (OR 3.55, 95%CI 1.31-9.60) และการไม่มีเครื่องเป่าระงับปริมาณคาร์บอนไดออกไซด์ในลมหายใจออก (OR 2.27, 95%CI 1.26-4.09) มีความสัมพันธ์กับการเสียชีวิตอย่างมีนัยสำคัญ

สรุป: ปัจจัยเสี่ยงของการเกิดภาวะแทรกซ้อนทางระบบหัวใจและหลอดเลือด ได้แก่ ASA physical status 3-5 และการไม่มีวิสัญญีแพทย์ในทีมให้การระงับความรู้สึก และปัจจัยเสี่ยงของการเสียชีวิตระหว่างและหลังการผ่าตัด ได้แก่ ASA physical status 3-5, ภาวะชุกเฉินและการไม่มีหรือไม่ได้ใช้เครื่องวัดระดับคาร์บอนไดออกไซด์ในลมหายใจออก
