

RESEARCH

Open Access



Association of intraocular pressure and postoperative nausea and vomiting after microvascular decompression - a prospective cohort study

Yuantao Hou¹, Hansheng Liang¹, Cungang Fan², Ruen Liu^{2*} and Yi Feng^{1*}

Abstract

Background: Postoperative nausea and vomiting is common in patients receiving microvascular decompression. In the current study, we examined whether postoperative nausea and vomiting is associated with reduced intraocular pressure (IOP) after microvascular decompression, a measure that reflects intracranial pressure.

Methods: This is a prospective cohort study. Adult patients scheduled for microvascular decompression surgery for hemifacial spasm between January 2020 and August 2020 were eligible. IOP was measured immediately before anesthesia induction and 30 min after patients regained complete consciousness using non-contact tonometry. IOP reduction was defined by at least 1 mmHg decrease vs. preoperative baseline. The primary outcome was vomiting on postoperative day 1.

Results: A total of 103 subjects were enrolled. IOP was reduced in 56 (54.4%) subjects. A significantly greater proportion of patients with IOP reduction had vomiting on postoperative day 1 (51.8% (29/56) vs. 23.4% (11/47) in those without IOP reduction; $p = 0.003$). In the multivariate regression analysis, vomiting on postoperative day 1 was associated with female sex [odds ratio = 7.87, 95% CI: 2.35–26.32, $p = 0.001$] and IOP reduction [odds ratio = 2.93, 95% CI: 1.13–7.58, $p = 0.027$].

Conclusions: In patients undergoing microvascular decompression surgery, postoperative IOP reduction is associated with postoperative vomiting.

Trial registration:: Chinese Clinical Trial Registry: [ChiCTR2000029083](https://www.clinicaltrials.gov/ct2/show/study?term=ChiCTR2000029083). Registered 13 January 2020.

Keywords: Hemifacial spasm, Intraocular pressure, Microvascular decompression, Nausea, Vomiting

Background

Microvascular decompression (MVD) is the standard treatment for hyperactive dysfunctional cranial nerve syndromes (such as trigeminal neuralgia, hemifacial spasm, and glossopharyngeal neuralgia) in patients who

do not respond to or tolerate pharmacological treatments [1, 2]. A significant proportion of patients experience severe postoperative nausea and vomiting (PONV) in the first 24 h after surgery [3–5]. PONV increases complications (e.g., surgical site bleeding, electrolyte disturbance, dehydration, and aspiration) and delays postoperative recovery [2, 6].

PONV has been partly attributed to loss of cerebrospinal fluid (CSF) and sudden reduction of intracranial pressure (ICP) [2]. This circumstance is similar to the

*Correspondence: liure@126.com; doctor_yifeng@sina.com

¹ Department of Anesthesiology, Peking University People's Hospital, No. 11 Xizhimen South Street, Xicheng District, Beijing, China

² Department of Neurosurgery, Peking University People's Hospital, No. 11 Xizhimen South Street, Xicheng District, Beijing, China



post-dural puncture headache, in which CSF loses rapidly, and is often accompanied by nausea and vomiting [7]. Treatments for PONV, including prone position, rehydration, and autologous epidural blood patch, are based on restoring ICP [8].

ICP is often determined via lumbar puncture, an invasive procedure with a risk of nerve injury and infection. Intraocular pressure (IOP) is widely used as a surrogate monitoring [9]. Compared with other methods, IOP monitoring is easy, fast, and inexpensive. We conducted a prospective study to examine PONV in patients undergoing MVD surgery using air puff tonometry. The rate of PONV was compared between patients with significant IOP reduction (at least 1 mmHg decrease from the preoperative baseline) vs. those without IOP reduction after MVD surgery.

Methods

Ethics, consent and permission

This study was approved by the Ethics Committee of Peking University People's Hospital (#2019PHB271-01; December 31st 2019). Written informed consents were obtained from all participants. The trial was registered at the Chinese Clinical Trial Registry (ChiCTR2000029083; January 13th 2020) (<http://www.chictr.org.cn/edit.aspx?pid=48279&htm=4>).

Participants

This prospective study was conducted in Peking University People's Hospital between January 2020 and August 2020. Adult patients (18–75 years of age) scheduled for MVD surgery for hemifacial spasm were eligible. The diagnosis of hemifacial spasm was established according to medical history, clinical manifestation of involuntary facial movements, and neurological imaging [10]. The main exclusion criteria were (1) body mass index (BMI) at <18 or >30 kg/m², (2) preoperative diagnosis of motion sickness or vertigo, (3) ophthalmic diseases (e.g., glaucoma, cataract, eye trauma) or previous eye surgery, (4) uncontrolled hypertension (systolic blood pressure >180 mmHg and/or diastolic blood pressure >110 mmHg despite of treatment), uncontrolled diabetes (fasting blood glucose >10 mmol/l despite of treatment), severe cardio-cerebrovascular disease, and mental diseases, and (5) pre-planned immediate return to the intensive care unit after surgery.

IOP measurement

IOP was measured using a Pulsair Intellipuff portable noncontact tonometer (Keeler Ltd., Windsor, UK), immediately prior to anesthesia induction and 30 min after the patients gained complete consciousness in a supine position, by an experienced doctor not involved in the study

otherwise. Two to three measurements were taken for each eye in each patient, and averaged. IOP reduction was defined as IOP decrease by at least 1 mmHg from the preoperative baseline.

Anesthesia

Anesthesia was induced with intravenous midazolam (0.03 mg/kg), propofol (1.5–2.5 mg/kg), sufentanil (0.3–0.4 µg/kg), and rocuronium (0.8 mg/kg), and maintained with propofol and remifentanil at the bispectral index score between 45 and 55. PetCO₂ was maintained at 35–45 mmHg. For PONV prevention, patients received 1 mg droperidol and 40 mg methylprednisolone intravenously after intubation, and 5 mg tropisetron intravenously before skin suture. After the resumption of spontaneous respiration, patients received 1 mg neostigmine, 0.5 mg atropine and 0.5 mg flumazenil intravenously, and were extubated. Oxygen supplementation was conducted at a rate of 2 L/min for 6 h.

PONV assessment

PONV events were evaluated over consecutive 24 h periods, at 9 am on postoperative day (POD) 1, 2 and 3. Severity of nausea was evaluated using a 0–10 numerical rating scale (NRS), with 10 for unbearable nausea [11]. Vomiting included actual vomiting and retching. Rescue tropisetron (5 mg) was given intravenously when nausea score was ≥ 7 , or upon repeated episodes of vomiting. Dizziness was also assessed using an NRS.

Endpoints

The primary endpoint was the rate of vomiting on POD 1. Secondary endpoints were the rate and severity of nausea and dizziness, and postoperative tropisetron rescue on POD 1–3.

Sample size calculation and statistical analysis

We conducted a preliminary study in 51 patients. The results showed 55.9% (19/34) rate of vomiting on POD 1 in subjects with IOP reduction vs. 11.8% (2/17) in subjects with no IOP reduction. Assuming 90% power, and α at 0.05, a total of 82 subjects were required. Assuming a dropout rate of 20%, 103 subjects would be needed.

Continuous variables are presented as mean \pm standard deviation (SD) and analyzed using Student's *t*-test if distributed normally, and presented as median (interquartile range) and analyzed using Mann-Whitney U test otherwise. Categorical variables are presented as number and percentage, and analyzed using Chi-square test. Variables with $p < 0.20$ in univariate analysis in comparison between subjects with or without PONV were entered into a multivariable logistic regression analysis to identify the variables associated with PONV. All statistical

analyses were conducted using SPSS version 25.0 software (IBM, New York, NY, USA). $p < 0.05$ (2-sided) was considered statistically significant.

Results

A total of 189 patients were screened, and 103 were enrolled (Fig. 1). The final analysis included 103 patients. The mean age was 52.12 ± 9.09 years, with a

1:2 male-female ratio (Table 1). Fifty-six patients (54.4%) had IOP reduction (at least 1 mmHg decrease from the baseline).

Changes in IOP

Operative characteristics are shown in Table 2. The preoperative IOP was significantly higher in patients with IOP reduction vs. patients without IOP reduction ($p < 0.001$). Postoperative IOP was significantly lower

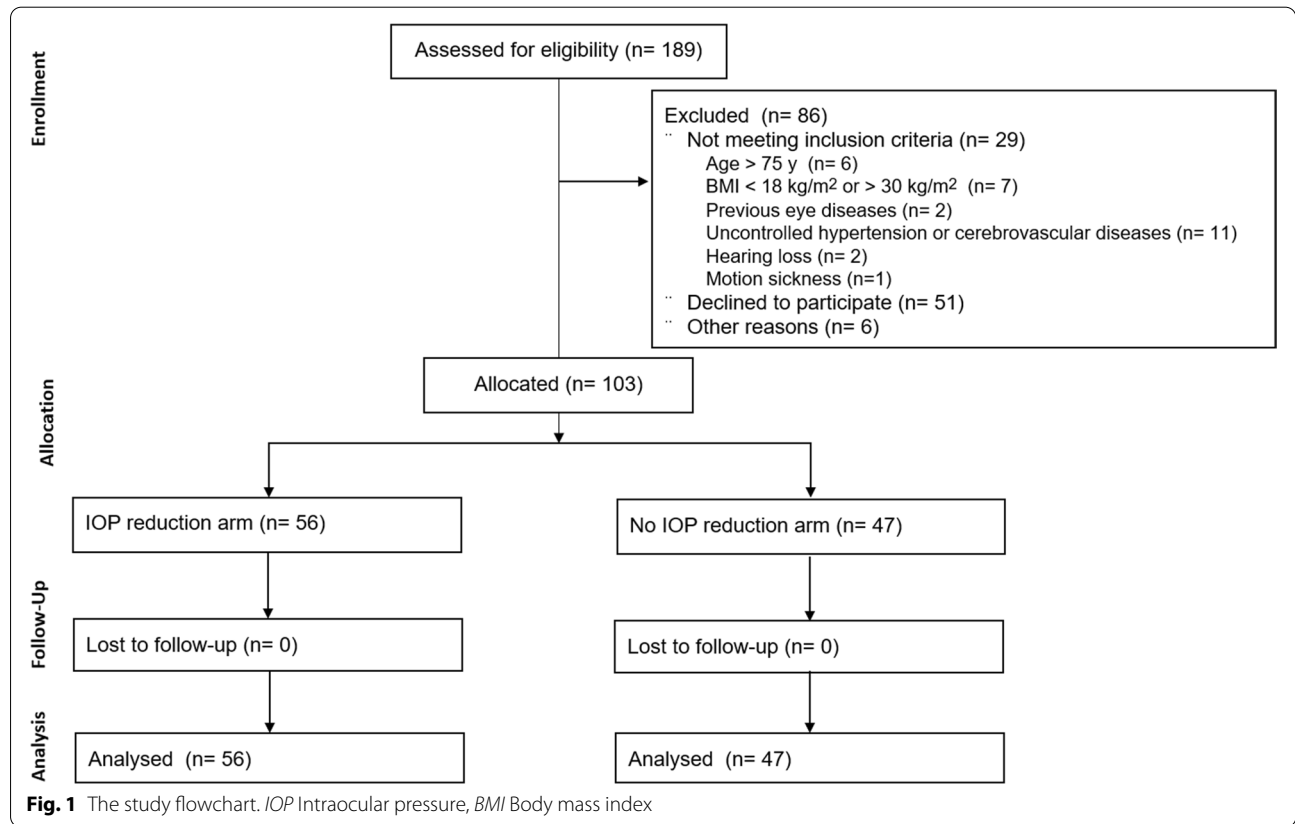


Table 1 Demographic and baseline characteristics of the study population

	All (n = 103)	IOP reduction (n = 56)	No IOP reduction (n = 47)	P value
Age, years, M ± SD	52.12 ± 9.09	51.71 ± 8.85	52.60 ± 9.43	0.626
Female, N (%)	70 (68.0)	40 (71.4)	30 (63.8)	0.410
BMI, kg/m ² , M ± SD	24.55 ± 3.00	24.65 ± 2.67	24.43 ± 3.37	0.179
Smoking, N (%)	18 (17.5)	9 (16.1)	9 (19.2)	0.682
Hypertension, N (%)	33 (32.0)	17 (30.4)	16 (34.0)	0.690
Hemoglobin, g/L, M ± SD	138.76 ± 15.76 [#]	138.82 ± 16.99	138.70 ± 14.30 [*]	0.968
Preoperative IOP				
mmHg, M ± SD	16.93 ± 3.11	18.31 ± 3.06	15.30 ± 2.28	< 0.001
>21 mmHg, N (%)	15 (14.6)	13 (23.2)	2 (4.3)	0.007

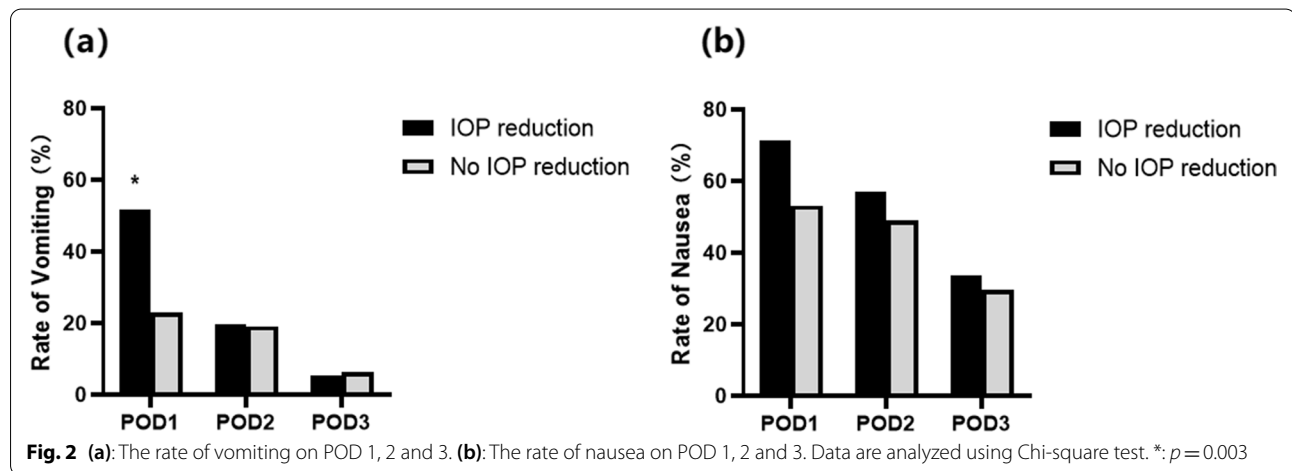
M ± SD Mean ± standard deviation, IOP Intraocular pressure, BMI Body mass index

^{*} n = 46, 1 missing. [#] n = 102, 1 missing

Table 2 Operative characteristics of the study population

	Total (n = 103)	IOP reduction (n = 56)	No IOP reduction (n = 47)	P value
Operative time, min, Median (IQR)	59.00 (51.00, 66.00)	58.00 (48.00, 64.00)	60.00 (53.00, 73.00)	0.098
Intraoperative sufentanil dose, mg, Median (IQR)	20.00 (16.00, 25.00)	20.00 (15.25, 25.00)	20.00 (16.00, 25.00)	0.777
Intraoperative fluid, mL, Median (IQR)	800.00 (600.00, 900.00)	750.00 (600.00, 900.00)	800.00 (700.00, 900.00)	0.459
Intraoperative output, mL, Median (IQR)	155.00 (110.00, 310.00)	120.00 (110.00, 220.00)	160.00 (110.00, 320.00)	0.190
Postoperative IOP, mmHg, M ± SD	15.68 ± 2.38	14.99 ± 2.26	16.49 ± 2.28	0.003

M ± SD Mean ± standard deviation, IQR Interquartile range, IOP Intraocular pressure



in patients with IOP reduction vs. patients without IOP reduction ($p = 0.003$).

Vomiting and tropisetron rescue

Forty (38.8%) patients experienced vomiting on POD 1. The rate of vomiting was significantly higher in patients with IOP reduction than those without IOP reduction on POD 1 [51.8% (29/56) vs. 23.4% (11/47), $p = 0.003$], but did not differ on POD 2 [19.6% (11/56) vs. 19.2% (9/47), $p = 0.950$] and POD 3 [5.4% (3/56) vs. 6.4% (3/47), $p = 0.825$] (Fig. 2). Tropisetron rescue during the first 3 postoperative days did not differ between patients with vs. without IOP reduction (21.4% (12/56) vs. 17.0% (8/47), $p = 0.573$).

In the univariate analysis, patients with vomiting on POD 1 had higher female ratio ($p < 0.001$) and higher rate of preoperative IOP > 21 mmHg ($p = 0.006$), IOP reduction ($p = 0.004$), smoking ($p = 0.017$) and higher preoperative IOP ($p = 0.011$). (Additional file 1). In the multivariate regression analysis, IOP reduction and female sex were independent risks of vomiting on POD 1 (Table 3). The area under the curve was 0.781 (Fig. 3).

Table 3 Multivariate logistic regression analysis of risks of vomiting on postoperative day 1

	Odds ratio (95% CI)	P value
IOP reduction	2.93 (1.13–7.58)	0.027
Preoperative IOP > 21 mmHg	4.05 (0.98–16.69)	0.053
Female sex	7.87 (2.35–26.32)	0.001
Smoking	-	0.555
Preoperative IOP	-	0.707

95% CI 95% confidence interval, IOP Intraocular pressure

Nausea

A total of 65 (63.1%), 55 (53.4%) and 33 (32.0%) patients experienced nausea on POD 1, 2 and 3, respectively. No significant difference was observed between patients with IOP reduction and those without IOP reduction [71.4% (40/56) vs. 53.2% (25/47) on POD 1, $p = 0.056$; 57.1% (32/56) vs. 48.9% (23/47) on POD 2, $p = 0.406$; and 33.9% (19/56) vs. 29.8% (14/47) on POD 3, $p = 0.654$] (Fig. 2). The NRS nausea score did not differ between patients with vs. without IOP reduction [2.50 (0, 4.00) vs. 1.00 (0, 3.00) on POD 1, $p = 0.087$; 1.00 (0, 2.00) vs. 0 (0, 2.00) on POD 2, $p = 0.409$; and 0 (0, 1.00) vs. 0 (0, 1.00) on POD 3, $p = 0.453$].

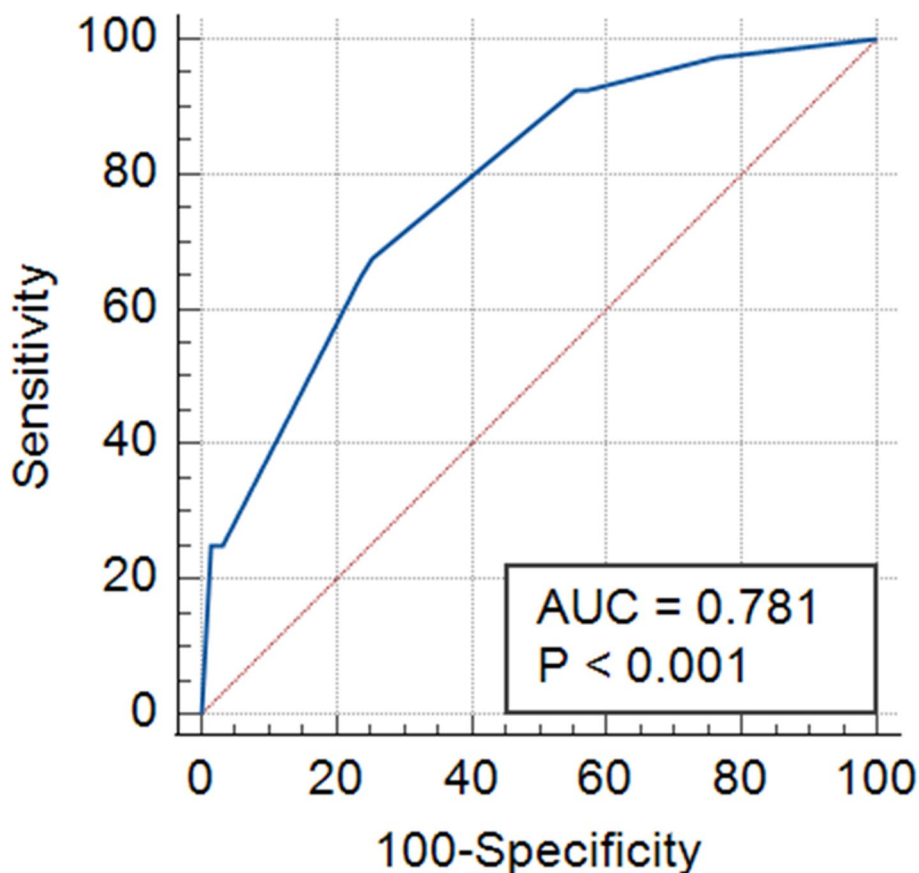


Fig. 3 The ROC curve for vomiting on POD 1. The area under the curve on POD 1 is 0.781. ROC, receiving operating characteristics; POD, postoperative day

Discussion

In the current study, approximately 40.0% patients experienced vomiting and 60.0% patients experienced nausea on POD 1 after MVD surgery, despite pre-emptive treatments to prevent PONV. This PONV rate observed in our study is similar to previous studies [3–5]. We also found that IOP reduction after MVD surgery is an independent risk factor for vomiting on POD 1. To our knowledge, this is the first study observing the relationship between IOP reduction and PONV.

From an anatomic point of view, the optic nerve is surrounded by the optic nerve sheath, which is continuous with the dura mater, arachnoid membrane, and pia mater [12], the CSF surrounds the optic nerve sheath up to the point where the optic nerve enters the orbit [13]. Though the intraocular space does not exchange fluid with the retrobulbar subarachnoid space significantly, the IOP can be influenced through the deformation of the lamina cribrosa (a barrier between the intraocular space and the extraocular cerebrospinal fluid space) provided by the pressure difference between these spaces [14]. For

IOP measurement, the Goldmann applanation tonometry (GAT) is the gold standard [15], but requires direct contact with cornea. Previous studies showed that, IOP measured with an air puff tonometer agrees well with the results obtained with GAT in both normotensive and hypertensive patients [16–19]. So, we chose a portable noncontact tonometer in our study.

Previous studies have examined the relationship between ICP and IOP. An animal study conducted in dogs showed that, when ICP remained above 70 mmH₂O, ICP decrease was significantly correlated with a decrease in IOP [20]. Further study conducted in male Sprague-Dawley rats showed that, stimulation of the dorsomedial hypothalamus/perifomical region led to increases in both ICP and IOP, indicating the presence of common regulatory regions of ICP and IOP in the brain [21]. A study of 50 patients showed significant correlation between ICP (as measure with lumbar puncture) and IOP independent of BMI, age and disease type [12]. A meta-analysis that included 546 subjects examined the correlation between ICP and IOP. They found

moderate correlation between IOP and ICP, and suggested IOP could be used for intracranial hypertension diagnosis [22]. The included studies in this meta-analysis showed significant heterogeneity, and further studies are needed before using IOP as routine evaluation of intracranial hypertension. The results of our study indicated that, Theoretically, measures that target intracranial hypotension (e.g., prolonged bed rest and fluid infusion) should be considered in patients with robust IOP reduction after MVD surgery. However, multi-centered trials with bigger sample size and more solid study design are needed before translation into clinical practice, and potential risks such as deep venous thrombosis and heart failure should be carefully weighed.

The approximately 40.0% rate of vomiting and 60.0% rate of nausea in the current study was very high, considering the fact that all study subjects received methylprednisolone, droperidol and tropisetron. In contrast, the rate of PONV in western countries is similar despite of less-potent anti-emetic regimen [4]. The reason for such a discrepancy is unknown, but has been previously attributed to ethnicity [23]. With regards to the use of tropisetron as rescue treatment, patients undergoing microvascular depression are highly susceptible to severe PONV, and require prophylaxis using multiple antiemetic agents. Option of selecting an agent from a different class is thus limited. A previous study observed significant IOP decrease when using droperidol 5 mg intravenously [24]. In our study, droperidol dosage was considerably lower at 1 mg, with expected less effect on IOP.

Our study has several limitations. First, exclusion of patients with uncontrolled hypertension or diabetes, or severe cardio-cerebrovascular diseases, factors which possibly influence IOP value, could decrease generality of our results. Atropine and neostigmine were given for muscle relaxant antagonism after general anesthesia, which may affect IOP. However, all patients were treated with the same drugs and at identical doses, so the impact was minimal. Not measuring ICP directly represents another inherent weakness.

Conclusions

In conclusion, PONV occurs in a significant proportion of patients undergoing MVD surgery, and postoperative IOP reduction is an independent and significant predictor of vomiting on POD 1.

Abbreviations

MVD: Microvascular decompression; PONV: Postoperative nausea and vomiting; CSF: Cerebrospinal fluid; ICP: Intracranial pressure; IOP: Intraocular pressure; BMI: Body mass index; NRS: Numerical rating scale; POD: Postoperative day; SD: Standard deviation; IQR: Interquartile range; OR: Odds ratio; CI: Confidence interval.

Supplementary information

The online version contains supplementary material available at <https://doi.org/10.1186/s12871-022-01665-x>.

Additional file 1: Table 1. Univariate analysis of risks of vomiting on postoperative day 1.

Acknowledgements

We would like to thank Huixin Liu, PhD, from Peking University People's Hospital, for providing assistance in statistical analysis.

Authors' contributions

Planning the study: YF, RL. Conduction of the study: YH, HL, CF. Drafting the article: YH. Revising the article for important intellectual content: YF, RL. All authors approved the final version of the manuscript.

Funding

The authors declare that they have no funding.

Availability of data and materials

The datasets used and/or analysed during the current study are available from the corresponding author on reasonable request.

Declarations

Ethics approval and consent to participate

This study was approved by the Ethics Committee of Peking University People's Hospital (#2019PHB271-01; December 31th 2019). Written informed consents were obtained from all participants. We confirm that all methods were performed in accordance with the relevant guidelines and regulations.

Consent for publication

Not applicable.

Competing of interests

The authors declare that they have no competing interests.

Received: 4 August 2021 Accepted: 18 April 2022

Published online: 30 April 2022

References

- Miller LE, Miller VM. Safety and effectiveness of microvascular decompression for treatment of hemifacial spasm: a systematic review. *Br J Neurosurg.* 2012;26:438–44. <https://doi.org/10.3109/02688697.2011.641613>.
- Sato K, Sai S, Adachi T. Is microvascular decompression surgery a high risk for postoperative nausea and vomiting in patients undergoing craniotomy? *J Anesth.* 2013;27:725–30. <https://doi.org/10.1007/s00540-013-1621-9>.
- Thongrong C, Chullabodhi P, Kasemsiri P, Kitkhuandee A, Plailaharn N, Sabangban L, et al. Effects of intraoperative dexamethasone and ondansetron on postoperative nausea and vomiting in microvascular decompression surgery: a randomized controlled study. *Anesthesiol Res Pract.* 2018;2018:6297362. <https://doi.org/10.1155/2018/6297362>.
- Meng L, Quinlan JJ. Assessing risk factors for postoperative nausea and vomiting: a retrospective study in patients undergoing retromastoid craniectomy with microvascular decompression of cranial nerves. *J Neurosurg Anesthesiol.* 2006;18:235–9. <https://doi.org/10.1097/00008506-200610000-00003>.
- Venkatraghavan L, Li L, Bailey T, Manninen PH, Tymianski M. Sumatriptan improves postoperative quality of recovery and reduces postcraniotomy headache after cranial nerve decompression. *Br J Anaesth.* 2016;117:73–9. <https://doi.org/10.1093/bja/aew152>.
- Cheong KB, Zhang JP, Huang Y, Zhang ZJ. The effectiveness of acupuncture in prevention and treatment of postoperative nausea and vomiting—a systematic review and meta-analysis. *PLoS One.* 2013;8:e82474. <https://doi.org/10.1371/journal.pone.0082474>.

7. Turnbull DK, Shepherd DB. Post-dural puncture headache: pathogenesis, prevention and treatment. *Br J Anaesth*. 2003;91:718–29. <https://doi.org/10.1093/bja/aeg231>.
8. Iga K, Murakoshi T, Kato A, Kato K, Terada S, Konno H, et al. Repeat epidural blood patch at the level of unintentional dural puncture and its neurologic complications: a case report. *JA Clin Rep*. 2019;5:14. <https://doi.org/10.1186/s40981-019-0232-3>.
9. Price DA, Grzybowski A, Eikenberry J, Januleviciene I, Verticchio Vercellin AC, Mathew S, et al. Review of non-invasive intracranial pressure measurement techniques for ophthalmology applications. *Br J Ophthalmol*. 2020;104:887–92. <https://doi.org/10.1136/bjophthalmol-2019-314704>.
10. Yaltho TC, Jankovic J. The many faces of hemifacial spasm: differential diagnosis of unilateral facial spasms. *Mov Disord*. 2011;26:1582–92. <https://doi.org/10.1002/mds.23692>.
11. Hou Y, Yan Q, An H, Wang J, Tian M, Zhao W, et al. The use and protective effects of transcutaneous electrical acupoint stimulation during abdominal surgery: study protocol for a multicenter randomized parallel controlled trial. *Trials*. 2019;20:462. <https://doi.org/10.1186/s13063-019-3558-2>.
12. Sajjadi SA, Harirchian MH, Sheikhabahaei N, Mohebbi MR, Malekmadani MH, Saberi H. The relation between intracranial and intraocular pressures: study of 50 patients. *Ann Neurol*. 2006;59:867–70. <https://doi.org/10.1002/ana.20856>.
13. Salman MS. Can intracranial pressure be measured non-invasively? *Lancet*. 1997;350:1367. [https://doi.org/10.1016/S0140-6736\(05\)65138-0](https://doi.org/10.1016/S0140-6736(05)65138-0).
14. Kaskar OG, Fleischman D, Lee YZ, Thorp BD, Kuznetsov AV, Grace L. Identifying the critical factors governing translaminar pressure differential through a compartmental model. *Invest Ophthalmol Vis Sci*. 2019;60:3204–14. <https://doi.org/10.1167/iovs.18-26200>.
15. Prabhakar SK, Mahesh BS, Shanthamallappa M. A comparative study of intraocular pressure measurement by three tonometers in normal subjects. *Nepal J Ophthalmol*. 2013;5:201–6. <https://doi.org/10.3126/nepjoph.v5i2.8729>.
16. Ogbuehi KC, Almubrad TM. Accuracy and reliability of the Keeler Pulsair EasyEye non-contact tonometer. *Optom Vis Sci*. 2008;85:61–6. <https://doi.org/10.1097/OPX.0b013e31815ed742>.
17. Hubanova R, Aptel F, Zhou T, Arnol N, Romanet JP, Chiquet C. Comparison of intraocular pressure measurements with the Reichert Pt100, the Keeler Pulsair Intellipuff portable noncontact tonometers, and Goldmann applanation tonometry. *J Glaucoma*. 2015;24:356–63. <https://doi.org/10.1097/01.jg.0000435776.99193.41>.
18. Babalola OE, Kehinde AV, Iloegbunam AC, Akinbinu T, Moghalu C, Onuoha I. A comparison of the Goldmann applanation and non-contact (Keeler Pulsair EasyEye) tonometers and the effect of central corneal thickness in indigenous African eyes. *Ophthalmic Physiol Opt*. 2009;29:182–8. <https://doi.org/10.1111/j.1475-1313.2008.00621.x>.
19. Parker VA, Herrtage J, Sarkies NJ. Clinical comparison of the Keeler Pulsair 3000 with Goldmann applanation tonometry. *Br J Ophthalmol*. 2001;85:1303–4. <https://doi.org/10.1136/bjo.85.11.1303>.
20. Hou R, Zhang Z, Yang D, Wang H, Chen W, Li Z, et al. Pressure balance and imbalance in the optic nerve chamber: The Beijing Intracranial and Intraocular Pressure (iCOP) Study. *Sci China Life Sci*. 2016;59:495–503. <https://doi.org/10.1007/s11427-016-5022-9>.
21. Samuels BC, Hammes NM, Johnson PL, Shekhar A, McKinnon SJ, Allingham RR. Dorsomedial/Perifornical hypothalamic stimulation increases intraocular pressure, intracranial pressure, and the translaminar pressure gradient. *Invest Ophthalmol Vis Sci*. 2012;53:7328–35. <https://doi.org/10.1167/iovs.12-10632>.
22. Yavin D, Luu J, James MT, Roberts DJ, Sutherland GR, Jette N, et al. Diagnostic accuracy of intraocular pressure measurement for the detection of raised intracranial pressure: meta-analysis: a systematic review. *J Neurosurg*. 2014;121:680–7. <https://doi.org/10.3171/2014.4.JNS13932>.
23. Klenke S, Frey UH. Genetic variability in postoperative nausea and vomiting: A systematic review. *Eur J Anaesthesiol*. 2020;37:959–68. <https://doi.org/10.1097/EJA.0000000000001224>.
24. Harris A, Zalish M, Kagemann L, Siesky B, Migliardi R, Garzozzi HJ. Effect of intravenous droperidol on intraocular pressure and retrobulbar hemodynamics. *Eur J Ophthalmol*. 2002;12:193–9. <https://doi.org/10.1177/112067210201200304>.

Publisher's note

Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.

Ready to submit your research? Choose BMC and benefit from:

- fast, convenient online submission
- thorough peer review by experienced researchers in your field
- rapid publication on acceptance
- support for research data, including large and complex data types
- gold Open Access which fosters wider collaboration and increased citations
- maximum visibility for your research: over 100M website views per year

At BMC, research is always in progress.

Learn more biomedcentral.com/submissions

