

# Oculocardiac reflex and oculo-respiratory reflex during strabismus surgery under general anesthesia using the laryngeal mask airway with maintenance of spontaneous respiration: A retrospective study

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## Abstract

**Objective:** To determine (1) how the specific muscle undergoing surgical treatment affects the occurrence of the oculocardiac reflex (OCR) and oculo-respiratory reflex (ORR) and (2) whether the depth of anesthesia influences the occurrence of the OCR and ORR in patients undergoing strabismus surgery with a laryngeal mask airway with maintenance of spontaneous respiration.

**Methods:** The medical records of patients who underwent strabismus surgery on the lateral rectus (LR) and medial rectus (MR) muscles from January 2017 to December 2017 were retrospectively reviewed.

**Results:** The incidence of the OCR was not significantly different between LR and MR operations in either pediatric or adult patients. The incidence of the ORR as indexed by the tidal volume (TV) was significantly higher during MR than LR surgery in pediatric patients (29.3% vs. 10.1%, respectively). The change in TV during muscle traction and the bispectral index were significantly correlated in both pediatric and adult patients ( $r^2 = 0.034$  and  $0.058$ , respectively).

**Conclusions:** The OCR was not influenced by the specific muscle undergoing surgery or the depth of anesthesia. The incidence of the ORR as indexed by the TV was higher during MR surgery in pediatric patients and was positively correlated with the depth of anesthesia.

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## Keywords

Strabismus surgery, laryngeal mask airway, oculocardiac reflex, oculorespiratory reflex, depth of anesthesia, spontaneous respiration

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## Introduction

The oculocardiac reflex (OCR) is evoked by parasympathetic stimulation induced by manipulation of the eye or extraocular muscles; thus, it typically occurs during strabismus surgery.<sup>1</sup> The afferent pathway of the OCR runs from the ophthalmic branch of the trigeminal nerve to the vagus nuclei, while the efferent pathway runs from the vagus nerve to the heart.<sup>2,3</sup> The OCR is defined as a 10% to 20% reduction in the heart rate (HR) from the baseline value or the development of dysrhythmia.<sup>1,2,4</sup> The incidence of the OCR ranges from 14% to 90% in previously published studies depending on the definition used.<sup>1,5–7</sup> In most cases, the OCR resolves without treatment, but some patients develop cardiac arrest and, in very rare cases, sudden death.<sup>8,9</sup> The OCR occurs more often in pediatric patients and in those undergoing surgery on the medial rectus (MR) muscle; however, it is less prevalent in pediatric patients when deeper anesthesia is induced.<sup>5,7,10–12</sup>

The oculorespiratory reflex (ORR) manifests as shallow respiratory movement, bradypnea, and respiratory arrest caused by pressure in or manipulation of the eye.<sup>13,14</sup> The afferent pathway of the ORR is identical to that of the OCR, but the efferent pathway has not yet been fully elucidated.<sup>13</sup> Additionally, the exact definition of, incidence of, and risk factors for the ORR have not yet been clearly described. Because most strabismus surgeries are performed under general anesthesia with mechanical ventilation that requires neuromuscular blocking agents (e.g.,

rocuronium), very little clinical research has focused on the ORR.

Most strabismus surgeries are performed on an outpatient basis. Therefore, anesthesia using a laryngeal mask airway (LMA) to maintain spontaneous respiration has been suggested in patients undergoing strabismus surgery.<sup>10</sup> However, few studies have focused on the OCR and ORR during strabismus surgery in relation to the depth of anesthesia, especially in terms of the specific extraocular muscle undergoing surgery.<sup>3,7,11,13</sup> Therefore, we designed this retrospective study primarily to elucidate the occurrence of the OCR and ORR according to the specific extraocular muscle being operated on in both pediatric and adult patients undergoing strabismus surgery with an LMA and maintenance of spontaneous respiration. The secondary aim was to determine whether the depth of anesthesia influences the occurrence of the OCR and ORR.

## Patients and methods

### *Study population and ethical approval*

The Institutional Review Board of Seoul St. Mary's Hospital, The Catholic University of Korea approved this study (approval no. KC17RESI0365) and waived the requirement for obtaining informed consent because of the retrospective nature of the study. We reviewed the medical records of patients who underwent strabismus surgery at Seoul St. Mary's Hospital from January to December 2017. Data abstraction was performed using the hospital's electronic

charting system. Patients were excluded if they had an American Society of Anesthesiologists physical status of >III; underwent strabismus surgery on the superior/inferior rectus and oblique muscles; had bronchial asthma, chronic obstructive pulmonary disease, sleep apnea, or heart disease; or had no OCR and ORR data. Data on sex, age, body weight, the specific muscles operated on, HR, tidal volume (TV), respiratory rate (RR), minute ventilation (MV), and muscle traction time were collected. Patients aged <15 years were defined as children.

### **Anesthetic management**

All patients scheduled to undergo strabismus surgery under general anesthesia at our hospital were routinely managed as follows. The patients were allowed to eat and drink until 8 hours before surgery. Intravenous access was established with a 20- to 24-gauge angiocatheter at the day surgery center (DSC). For pediatric patients who refused to enter the operating room alone, 0.05 to 0.10 mg of midazolam was intravenously administered at the DSC; otherwise, no premedication was administered. Basic monitoring, including electrocardiography, noninvasive sphygmomanometry, pulse oximetry, and measurement of the bispectral index (BIS), was performed in the operating room. For anesthetic induction, 1.5 to 2.0 mg/kg of propofol was administered, and an LMA (LMA<sup>®</sup> Flexible<sup>™</sup> Airway; Teleflex, Westmeath, Ireland) fitted for age and weight was inserted when the patients were fully sedated. Anesthesia was maintained with sevoflurane at 2.0 to 2.5 vol%, 50% nitrous oxide, and 50% oxygen to maintain the BIS at 40 to 60. Spontaneous respiration was maintained with application of pressure support at 5 to 10 cmH<sub>2</sub>O, preventing the peak pressure from exceeding 20 cmH<sub>2</sub>O. At the end of surgery, the

sevoflurane was discontinued and the LMA was removed. The patients were transferred to the post-anesthesia care unit and then to the DSC for discharge.

### **Data collection and statistical analyses**

The ophthalmologist informed the anesthesiologist about the progress of the strabismus surgery, such as the extraocular muscle hooking and release, and the anesthesiologist recorded the cardiovascular and respiratory parameters at each point of the strabismus surgery. The lowest HR, TV, RR, and MV were recorded during hooking of the rectus muscle; all parameters were recorded at the moment of traction release. Changes in the HR, TV, RR, and MV were calculated as the lowest value during lateral rectus (LR) or MR muscle traction divided by the baseline value. The OCR was defined as a >20% reduction in the HR from the baseline value. The baseline HR was defined as the HR before surgical incision. The ORR was defined as a >20% reduction in the TV, RR, and MV from the baseline values. The baseline values of the respiratory parameters were defined as the values before surgical incision.

All data were analyzed using SPSS software ver. 20.0 (IBM Corp., Armonk, NY, USA). For demographic data, the  $\chi^2$  test or Fisher's exact test and the t-test were used. The incidence rates of the OCR and ORR were compared between the LR and MR muscles using the  $\chi^2$  test. Linear logistic regression was performed to analyze changes in the HR, TV, RR, and BIS during muscle traction. Data are presented as mean  $\pm$  standard deviation or number and percentage. A p-value of <0.05 was considered statistically significant.

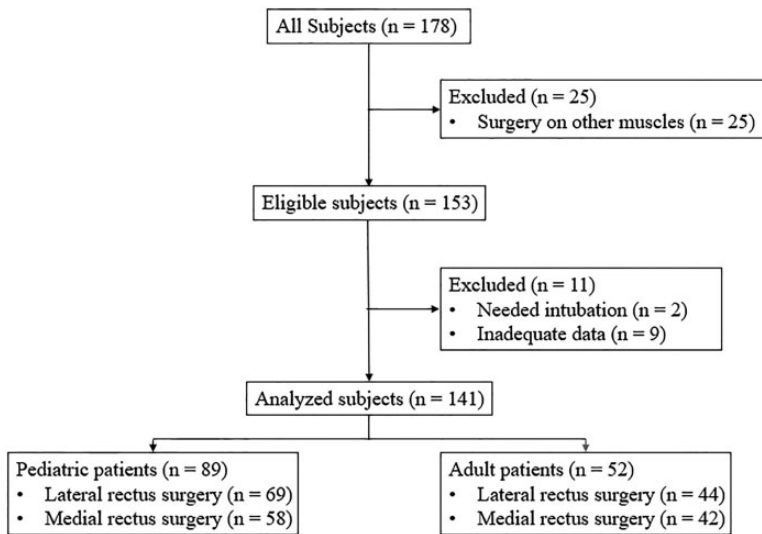
### **Results**

A total of 175 patients underwent strabismus surgery during the study period.

The data of 141 patients (89 children and 52 adults) who met the inclusion criteria were analyzed. The pediatric patients underwent 69 LR and 58 MR operations, while the adult patients underwent 44 LR and 42 MR operations (Figure 1). The mean muscle traction time was  $161 \pm 50.0$  s in children and  $156.1 \pm 42.5$  s in adults (Table 1).

The incidence of the OCR during LR versus MR operations was not significantly

different in either pediatric or adult patients (42.0% vs. 46.6% and 34.1% vs. 38.1%, respectively). The overall incidence of the ORR in terms of the TV was 12.2%. The incidence of the ORR in terms of the TV was significantly higher during MR surgery (29.3%) than during LR surgery (10.1%) in children ( $p < 0.05$ ); however, there was no significant difference in adult patients. The incidence of the ORR in terms of the RR



**Figure 1.** Patient flow diagram.

**Table 1.** Patient characteristics.

	Pediatric	Adult
Total	89	52
Sex, male/female	41/48	26/26
Age, years	$6.6 \pm 2.8$	$39.2 \pm 16.6$
Body weight, kg	$26.6 \pm 10.9$	$63.7 \pm 7.7$
Number of operated muscles, LR/MR	69/58	44/42
Baseline heart rate, beats/minute	$97.7 \pm 15.1$	$72.6 \pm 11.6$
Baseline tidal volume, mL/kg	$4.0 \pm 1.5$	$4.4 \pm 1.2$
Baseline respiratory rate, breaths/minute	$24.3 \pm 6.7$	$17.7 \pm 7.1$
Baseline minute ventilation, L/minute	$2.6 \pm 1.0$	$4.0 \pm 1.5$
Baseline BIS	$82.3 \pm 17.5$	$97.5 \pm 1.0$
Muscle traction time, s	$161.0 \pm 50.0$	$156.1 \pm 42.5$

Categorical variables are shown as number, and other variables are shown as mean  $\pm$  standard deviation. LR, lateral rectus; MR, medial rectus; BIS, bispectral index.

and MV was not significantly different between LR and MR surgeries in either pediatric or adult patients (Table 2). The overall incidence of the OCR was not significantly different between pediatric and adult patients. The incidence of the ORR in terms of the TV was significantly higher in pediatric patients than that in adult patients (18.9% vs. 2.3%,  $p < 0.001$ ), while there was no significant difference in the incidence of the ORR in terms of the RR and MV between pediatric and adult patients (Table 3).

The change in the HR during muscle traction and the BIS showed a negative correlation in pediatric patients, but this was not statistically significant (Figure 2a). The change in the TV during muscle traction and the BIS were significantly correlated in both pediatric and adult patients ( $r^2 = 0.034$  and  $0.058$ , respectively;  $p < 0.05$ ) (Figures 2b and 3b), while the change in the RR during muscle traction

and the BIS showed no significant correlation in either group (Figures 2c and 3c). The change in the MV during muscle traction was correlated with the BIS in pediatric patients ( $r^2 = 0.043$ ,  $p < 0.05$ ), but not in adult patients (Figures 2d and 3d).

## Discussion

Strabismus surgery is one of the most common ophthalmologic surgeries performed under general anesthesia, especially in pediatric patients. To the best of our knowledge, this is the first study to compare the incidence rates of the OCR and ORR during LR and MR strabismus surgeries in patients undergoing general anesthesia using an LMA with spontaneous respiration; moreover, it is the first study to investigate the incidence rates of the OCR and ORR in relation to the depth of anesthesia. This study showed that the incidence of the OCR was not significantly different between LR and MR surgeries in either pediatric or adult patients; in addition, it was not correlated with the depth of anesthesia. In pediatric patients, the incidence of the ORR in terms of the TV was higher during MR surgery than during LR surgery. Changes in the TV during rectus muscle traction were correlated with the depth of anesthesia, while changes in the RR during rectus muscle traction were not

**Table 2.** Incidence of OCR and ORR.

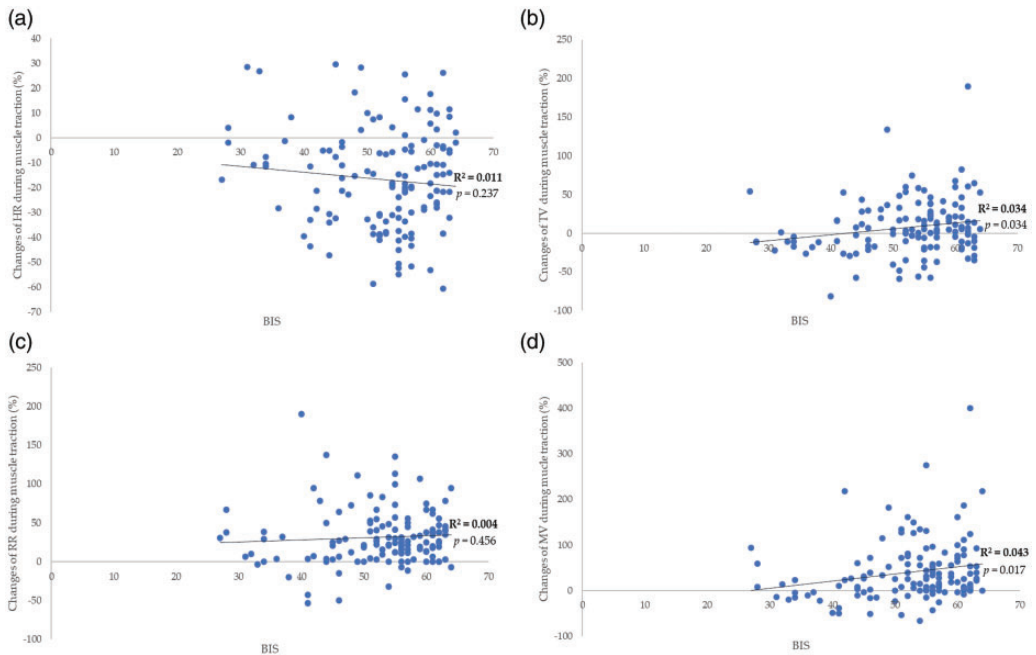
	LR muscle	MR muscle	p value
OCR			
Total	44 (38.9)	43 (43.0)	0.737
Pediatrics	29 (42.0)	27 (46.6)	0.546
Adults	15 (34.1)	16 (38.1)	0.709
ORR			
TV			
Total	8 (7.1)	18 (18.0)	0.312
Pediatrics	7 (10.1)	17 (29.3)	0.008*
Adults	1 (2.3)	1 (2.4)	0.722
RR			
Total	1 (0.9)	2 (2.0)	0.192
Pediatrics	1 (1.4)	1 (1.7)	0.532
Adults	0 (0.0)	1 (2.4)	0.307
MV			
Total	3 (2.7)	7 (7.0)	0.217
Pediatrics	2 (2.9)	6 (10.3)	0.106
Adults	1 (2.3)	1 (2.4)	0.722

Variables are shown as number (percentage). LR, lateral rectus; MR, medial rectus; OCR, oculocardiac reflex; ORR, oculorespiratory reflex; TV, tidal volume; RR, respiratory rate; MV, minute ventilation. \* $p < 0.05$ .

**Table 3.** Comparison between pediatric and adult patients.

	Pediatric	Adult	p value
OCR	56 (44.1)	31 (36.0)	0.259
ORR			
TV	24 (18.9)	2 (2.3)	0.000*
RR	2 (1.6)	1 (1.2)	1.000
MV	8 (6.3)	2 (2.3)	0.322

Variables are shown as number (percentage). OCR, oculocardiac reflex; ORR, oculorespiratory reflex; TV, tidal volume; RR, respiratory rate; MV, minute ventilation. \* $p < 0.05$ .



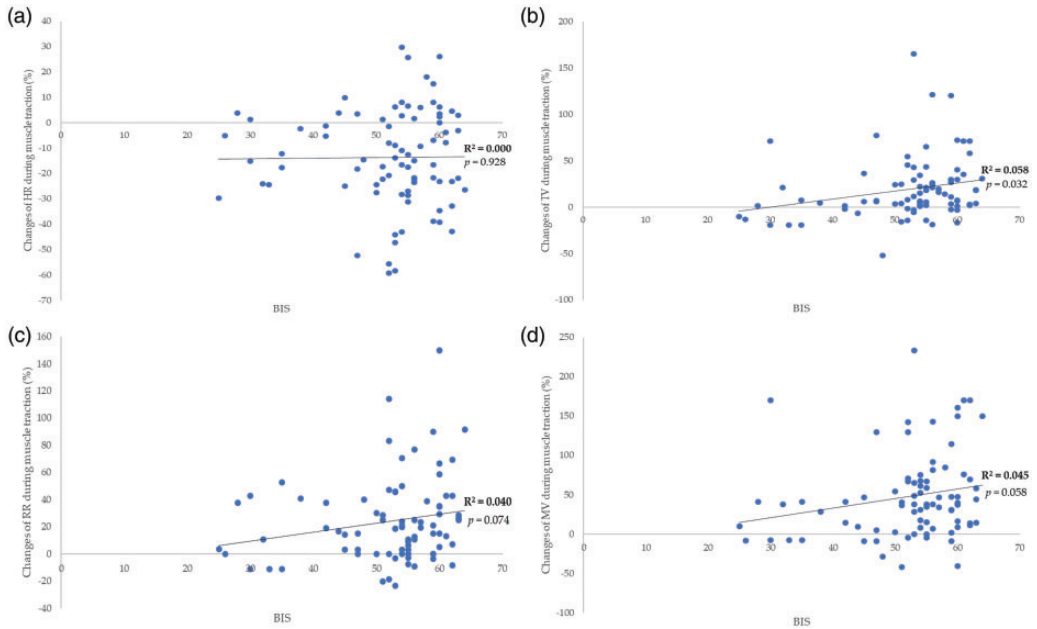
**Figure 2.** Relationship between BIS and changes in (a) HR, (b) TV, (c) RR, and (d) MV during muscle traction in pediatric patients. BIS, bispectral index; HR, heart rate; TV, tidal volume; RR, respiratory rate; MV, minute volume.

correlated with the depth of anesthesia in either pediatric or adult patients.

The OCR is an important phenomenon during strabismus surgery; thus, both anesthesiologists and ophthalmic surgeons closely monitor potential signs of the OCR to prevent its occurrence. Retrobulbar block, premedication with anticholinergics, and gentle tension during muscle traction have been suggested to reduce the incidence of the OCR.<sup>12,16,17</sup> Previous studies of the association between the OCR and operations on specific extraocular muscles have produced conflicting results. In some studies, the OCR occurred more frequently during MR surgery than during LR surgery because of the differences in the afferent pathways involved.<sup>3,4,7,18</sup> Other studies showed no relationship between the OCR and the specific extraocular muscle on which the operation was

performed.<sup>1,6,19,20</sup> In our study, LR and MR surgeries showed no differences in the incidence of the OCR in either pediatric or adult patients; moreover, changes in the HR during muscle traction were not associated with the depth of anesthesia. Changes in the HR during muscle traction were negatively correlated with the depth of anesthesia in pediatric patients, as in previous studies; however, this correlation was not statistically significant. We presume that this discrepancy with previous results was due to the maintenance of spontaneous respiration during surgery in our study population. Notably, respiration modulates sympathetic nerve activity, and a more rapid RR is associated with a higher level of sympathetic activity.<sup>21</sup> Additionally, negative intrathoracic pressure during spontaneous inspiration causes a slight increase in the HR because of vagal withdrawal and





**Figure 3.** Relationship between BIS and changes in (a) HR, (b) TV, (c) RR, and (d) MV during muscle traction in adult patients. BIS, bispectral index; HR, heart rate; TV, tidal volume; RR, respiratory rate; MV, minute volume.

lung hyperinflation, which may occur during mechanical ventilation and leads to bradycardia due to vagal overstimulation.<sup>22</sup> In contrast to the findings of previous studies involving general anesthesia with mechanical ventilation, the patients in our study maintained their spontaneous respiration; the resulting sympathetic stimulation offset the parasympathetic reflex that occurred during extraocular muscle manipulation.

The ORR has not been well characterized for anesthesiologists or ophthalmologic surgeons because most studies to date have been experimental in nature. Although the efferent pathway of the ORR has not been clearly elucidated, it appears to be independent of the vagus nerve because intravenous atropine enhances the ORR; moreover, as in the OCR, the afferent pathway stimulates the pneumotaxic respiratory center to send a signal to

the medullary respiratory area through the phrenic nerve and other respiratory nerves.<sup>13</sup> The ORR is prevented by retrobulbar block.<sup>15</sup>

Allison et al.<sup>11</sup> reported that the ORR occurred in a manner that led to a reduced TV without significant declines in the RR in pediatric patients who underwent strabismus surgery with the use of an LMA to maintain spontaneous respiration; this result is identical to the result of our study. We used sevoflurane, the respiratory effect of which has not yet been fully elucidated,<sup>11,23–26</sup> and we found that the overall incidence of the ORR in terms of the TV was 12.2% in pediatric patients. The incidence rate was higher during MR surgery than during LR surgery, which might have been a result of the different afferent pathways involved.<sup>19</sup> However, the overall incidence rate of the ORR in terms of the RR and MV was <5% and did not differ

between the LR and MR surgeries. Based on these findings, the ORR may be defined as a change of in the TV during muscle traction. Meanwhile, maintenance of the MV during traction of the MR and LR muscles indicates that muscle traction does not interfere with spontaneous respiration in a comprehensive sense.

Among the respiratory parameters, only the TV decreased with a greater depth of anesthesia in both pediatric and adult patients, although the correlations were marginal. In other words, the TV increased as the depth of anesthesia decreased (i.e., became lighter), which might be considered a result of stimulation of muscle hooking, such as pain. However, all patients were anesthetized deep enough to maintain a BIS of <60 even without using rocuronium, which may lower the BIS regardless of the actual depth of anesthesia.<sup>27</sup> We consider that the higher incidence of the ORR in terms of the TV in pediatric than adult patients is the reason that the BIS and MV showed a significant correlation only in pediatric patients.

One limitation of this study is that the OCR and ORR occurring during superior rectus, inferior rectus, and oblique muscle surgery were not analyzed because of the lower number of procedures performed on these muscles. Most strabismus surgeries are performed on the horizontal rectus extraocular muscles, such as the LR and MR muscles. Therefore, we only included patients who had undergone surgeries on two horizontal rectus muscles. Ha et al.<sup>6</sup> reported that the incidence of the OCR was significantly higher in the first operated muscle than in the second operated muscle in patients who underwent two muscle surgeries because of the adaptation to the subsequent stimulation. In the present study, we focused only on the effect of the MR and LR muscles on the OCR and ORR, including patients undergoing surgery on one or two muscles. We did not analyze

whether the OCR or ORR differed according to the order of the operated muscles in our study. This might have influenced the results of this study by introducing bias. Nevertheless, this study is considered sufficiently meaningful as the first study to analyze the OCR and ORR depending on the specific muscle being operated on and the depth of anesthesia in patients undergoing strabismus surgery with an LMA to maintain spontaneous respiration.

In summary, the OCR was not influenced by either the specific muscle being operated on or the depth of anesthesia. The ORR in terms of the TV occurred more frequently during MR surgery in pediatric patients and was positively correlated with the depth of anesthesia in both pediatric and adult patients. Thus, we propose that young age and MR surgery are risk factors for the ORR during strabismus surgery under general anesthesia with spontaneous respiration. Additionally, we do not advise deepening anesthesia to prevent the ORR when using an LMA with spontaneous respiration because the OCR is not correlated with the depth of anesthesia. However, further prospective observational studies should be performed to clarify the relationship between the ORR and the depth of anesthesia as well as to validate the risk factors for the ORR during strabismus surgery with the use of an LMA to maintain spontaneous respiration.

#### **Declaration of conflicting interest**


The authors declare that there is no conflict of interest.

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