ORi™: a new indicator of oxygenation

Yusuke Ishida1 · Toshio Okada1 · Takayuki Kobayashi1 · Hiroyuki Uchino1

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Abstract
In the perioperative period, hypoxemia and hyperoxia are crucial factors that require attention, because they greatly affect patient prognoses. The pulse oximeter has been the only noninvasive monitor that can be used as a reference of oxygenation in current anesthetic management; however, in recent years, a new monitoring method that uses the oxygen reserve index (ORi™) has been developed by Masimo Corp. ORi is an index that reflects the state of moderate hyperoxia (partial pressure of arterial oxygen [PaO2] between 100 and 200 mmHg) using a non-unit scale between 0.00 and 1.00. ORi monitoring performed together with percutaneous oxygen saturation (SpO2) measurements may become an important technique in the field of anesthetic management, for measuring oxygenation reserve capacity. By measuring ORi, it is possible to predict hypoxemia and to detect hyperoxia at an early stage. In this review, we summarize the method of ORi, cautions for its use, and suitable cases for its use. In the near future, the monitoring of oxygen concentrations using ORi may become increasingly common for the management of respiratory function before, after, and during surgery.

Keywords Oxygenation reserve · Hypoxemia · Hyperoxia · Oxygen reserve index (ORi)

Introduction
Over the years, intraoperative anesthetic monitoring techniques have been improved similarly to the improvement of surgical instruments. Blood pressure, heart rate, electrocardiography, and blood oxygen saturation level are mainly monitored during intraoperative anesthesia. In the present article, we review the oxygen reserve index (ORi™), which is a measurement that enables the assessment of oxygen reserve capacity, and discuss its various applications and future outlooks.

What is ORi
During anesthetic management, predicting the partial pressure of arterial oxygen (PaO2) is difficult when percutaneous oxygen saturation (SpO2) is 100%. In usual cases, in which arterial lines of the patients are secured, arterial blood gas analysis for the appropriate management of PaO2 is possible. However, as this is an invasive method, it cannot be performed on all patients. In most patients undergoing general anesthesia, tracheal intubation and artificial ventilation are carried out to manage positive-pressure ventilation. However, during surgery, atelectasis and pulmonary injury may occur owing to multiple factors [1–3], which may cause sudden hypoxemia during or after surgery, making it difficult to manage the patient [1, 4, 5]. Therefore, PaO2 during surgery is a useful indicator for anesthesiologists, particularly in patients with low respiratory function (e.g., obstructive/restrictive ventilatory disorder) and low oxygen reserve (e.g., pregnant women and obese patients) [6, 7]. In addition, controlling oxygen concentrations during surgery in patients with diseases in which hyperoxia should be avoided, such as interstitial pneumonia and retinopathy of prematurity [8], is a difficult problem. Although hypoxia can lead to adverse effects, hyperoxia during surgery can also induce pulmonary injury and absorption atelectasis, resulting in postsurgical...
respiratory complications [1–3]. Furthermore, hyperoxia can also increase the risk of other adverse events [9] (Table 1). Unfavorable clinical outcomes owing to a high fraction of inspired oxygen (FiO2) have been recently reported in critically ill adults, including patients with chronic obstructive pulmonary disease, myocardial infarction, cardiac arrest, stroke, and traumatic brain injury [10–15].

Until recently, the pulse oximeter was the only noninvasive instrument to measure arterial blood oxygen saturation (SaO2), which is an index of blood oxygenation. During the 1970s, the world’s first pulse oximeter that was able to measure SaO2 from the fingertip was commercialized. Since then, this device has been used in a wide range of situations for the noninvasive monitoring of blood oxygenation [16]. However, this device has its flaws. Although SpO2 and PaO2 are correlated at most points, when SpO2 reaches 100%, estimating PaO2 becomes difficult [17] (Fig. 1). Thus, other methods, such as arterial blood gas analysis, are often performed to measure PaO2 for the evaluation of oxygenation.

The oxygen reserve index (ORi) is a noninvasive and continuous parameter that can be measured by connecting a multi-wavelength sensor to Root® with Radical-7®, Pulse CO-Oximeter® (Masimo Corp., Irvine, CA, USA). ORi is an index measured using a non-unit scale between 0.00 and 1.00, which reflects the range of moderate hyperoxia (PaO2 between 100 and 200 mmHg), and is defined as oxygen reserve reflecting the level of venous blood oxygen saturation (SvO2). ORi remains at 0.00 in a room air environment. With supplemental oxygen, after SpO2 reaches 100%, SvO2 also increases as PaO2 does, and usually reaches a plateau at about a PaO2 of 200 mmHg [18, 19] (Fig. 2). In general, a pulse oximeter displays the SpO2 value measured from light absorption of the pulsatile blood at the site where the sensor is placed. Pulsatile changes are observed in the arteries, capillaries, and venules. Masimo developed rainbow SET® which can analyze changes in the absorption of incident light of both arterial and venous blood using a multi-wavelength sensor, and display them as ORi, which is a unitless scale between 0.00 and 1.00. Thus, ORi begins to increase from 0.00 at about a PaO2 of 100 mmHg, and reaches a plateau of 1.00 at approximately 200 mmHg [18–21]. However, there are reports stating that in cases where FiO2 is 1.0, even when

**Table 1** Adverse events caused by exposure to high oxygen concentrations

| Adverse events                                    |
|---------------------------------------------------|
| Absorption atelectasis                            |
| Vasoconstriction                                  |
| Central nervous system toxicity                   |
| Alveolar edema                                    |
| Pulmonary fibrosis                                |
| Hyperoxic acute lung injury                       |
| Increase in reactive oxygen species               |
| Increased mortality after resuscitation           |
| Higher incidence of reinfarction after acute myocardial infarction |
| Increased cancer recurrence                       |
| Increased risk of fire during certain procedures   |

(Cited from Anesth Analg. 2017;125:682–687 and modified)
Oxygenation is complete, the peak value is different for each individual and reaches a plateau at about 0.60 [22]. Applegate et al. analyzed the association between ORi and PaO₂ during surgery, and found that if PaO₂ is below 240 mmHg, ORi and PaO₂ show a positive correlation ($r^2 = 0.536$) [23] (Fig. 3). However, if PaO₂ is higher than 240 mmHg, there was no correlation ($r^2 = 0.0016$). Furthermore, when ORi was more than 0.24, PaO₂ was more than 100 mmHg. As the report showed that 96.6% of the samples had a PaO₂ of ≥ 150 mmHg when ORi was above 0.55, the authors concluded that ORi can be used as an indicator of PaO₂. Vos et al. investigated the correlation between PaO₂ and ORi during the administration of oxygen in 20 healthy volunteers, and they also reported that the two values have a strong positive correlation [24]. In addition, regarding the correlation between PaO₂ and ORi during anesthesia under oxygenation, when PaO₂ was less than 240 mmHg ($n = 69$), linear regression analysis showed a relatively strong positive correlation ($r^2 = 0.706$) [25].

### Suitable cases for the use of ORi with regard to anesthetic management

The following is a list of cases for which the use of ORi is suitable.

- **Rapid sequence induction**

  Normally in a room air environment, ORi is 0.00. After the administration of oxygen, even after SpO₂ reaches 100%, ORi will continue to increase. Rapid sequence induction (RSI) is appropriate for patients in whom the aspiration of gastric contents may occur, including those with a full stomach, ileus, and expectant mothers [26–28].

  ![Fig. 3 Correlation between PaO₂ and ORi. A positive correlation is observed between ORi and PaO₂ below 240 mmHg ($r^2 = 0.536$). The dotted lines indicate the 95% confidence interval of the regression line. (Modified and reproduced with permission) (Wolters Kluwer Health: Anesth Analg) [23]. PaO₂ partial pressure of arterial oxygen, ORi oxygen reserve index](image)

  In RSI, tracheal intubation is performed without ventilation after the occurrence of apnea. In such cases, ORi may be an indicator of the time it will take for SpO₂ to decrease during tracheal intubation. Yoshida et al. investigated the usefulness of ORi during RSI [29]. They reported that in 10 out of 17 patients (77%) in whom preoxygenation was performed, ORi started to decrease 32.5 s (IQR: 18.1–51.3) before a decrease in SpO₂ post-apnea. By monitoring the ORi, it is possible to predict a decrease in oxygenation about 30 s before SpO₂ starts to decrease. From this finding, Yoshida concluded that it may be possible to lower the incidence rate of hypoxemia during RSI. Thus, ORi is expected to be useful for RSI.

  In addition, owing to the recent outbreak of coronavirus disease 2019 (COVID-19), RSI has been recommended to prevent the infection of anesthesiologists during intubation [30, 31]. Furthermore, the use of aerosol boxes for the prevention of COVID-19 infection makes intubation difficult [32–34]. Thus, aerosol boxes may increase intubation times and therefore increase the risk of hypoxia in patients [32]. Using ORi may hence reduce the risk of hypoxemia and enable safe tracheal intubation.

- **Pediatric applications**

  In pediatric patients, oxygen consumption is higher, and the closing volume is also relatively larger than the functional residual capacity compared with adult patients. Owing to this fact, the development of hypoxemia in pediatric patients is more rapid than in adult patients during apnea. Previous studies investigating ORi in pediatric patients during anesthetic induction showed that ORi monitoring enabled detection of a reduction in oxygenation approximately 30 s prior to a rapid change in SpO₂ [21]. These studies suggest that prediction of the onset of a rapid decrease in oxygen saturation enables the prompt implementation of appropriate treatments, which is an advantage. In fact, the use of ORi for tracheoplasty in children was found to enable the earlier prediction of hypoxemia [35]. In addition, as the trachea is shorter in children than in adults, when performing tracheal intubation and head bending during surgery, there is a higher possibility of unilateral intubation in children [36, 37]. Therefore, in the former case, comparison of ORi before and after intubation may be useful, and in the latter case, a sudden drop in ORi during the operation may be a useful indicator of subsequent hypoxemia. Therefore, the use of ORi may enable the earlier implementation of appropriate treatments. All of the above demonstrate that ORi is a useful indicator in pediatric anesthesia during the perioperative period.

- **Other suitable cases**
ORi is also useful for patients in whom respiration management is required during one-lung ventilation (OLV). Hypoxemia may still occur in about 10% of such patients [38]. When switching from two-lung to OLV, the shunt fraction increases, oxygenation is impaired, and hypoxemia may occur [39]. Studies monitoring ORi during the anesthetic management of OLV showed that decreases in PaO₂ could be detected much earlier than when only SpO₂ was monitored. These studies suggest that ORi can reduce the risk of complications during OLV [40]. Another study reported that the monitoring of ORi during OLV could predict hypoxemia [41]. These findings may be helpful to adjust FiO₂ individually in patients undergoing OLV, and to avoid unnecessarily high concentrations of oxygen.

In addition, the use of ORi is suitable for surgeries that have a high probability of leading to atelectasis, such as surgeries on obese patients [42, 43], surgeries in the Trendelenburg position, and laparoscopic surgeries [43–45]. Other studies also suggested that ORi is useful in situations in which awake intubation is performed [22]. ORi monitoring may be able to avoid hypoxemia in cases in which bronchial stenting using a rigid endoscope is performed while the patient continues to breathe spontaneously [46]. For surgeries in which patients’ respiratory statuses can vary, ORi monitoring can be an indicator to decide whether or not early intervention is required to avoid hypoxemia [47]. ORi monitoring during the induction of anesthesia can be an indicator of whether preoxygenation is being carried out appropriately.

In addition, hyperoxia can be detected by monitoring ORi in patients under general anesthesia, and thus the administration of unnecessarily high concentrations of oxygen can possibly be avoided. In fact, there are several reports demonstrating that the use of ORi in managing oxygen levels during and after surgery, and for patients in the intensive care unit (ICU) can prevent exposure to excess oxygen [25, 48, 49]. In general, high oxygen concentrations are more harmful to infants, children, and older patients than to adults [50]. Oxidative stress that occurs as a consequence of treatment with high oxygen concentrations during the perinatal period notably affects the lung and brain development of premature and newborn infants. For instance, bronchopulmonary dysplasia is caused by hyperoxic lung injury [51–53]. In older patients, the production and accumulation of reactive oxygen species (ROS) are largely increased, and antioxidative functions are decreased, resulting in various aging-associated diseases [54, 55]. Hyperoxia promotes ROS production, suggesting that perioperative hyperoxia might enhance the adverse effects of aging-associated diseases, particularly in lung and brain diseases. Therefore, adequate oxygenation using ORi may lead to a reduction in complications.

**Methods and cautions of the use of ORi**

ORi is measured by placing an RD rainbow Lite SET® sensor, which is a multiwavelength sensor, on a patient’s fingertip and connecting it to Root® with Radical-7®. In addition to ORi, this sensor can measure a patient’s SpO₂, pulse rate (PR), Perfusion Index (PI), and Pleth Variability Index (PVI®) (Fig. 4). In addition, ORi can be measured by a RD rainbow SET®-2 sensor, which can measure Noninvasive and Continuous Hemoglobin (SpHb®). However, it is necessary to block any outside light when using the SpHb sensor, as this sensor is easily affected by light owing to its high sensitivity to long wavelengths (Fig. 4). Furthermore, special attention is needed to make sure that the SpO₂ is greater than 98% because if this criterion is not established, ORi does not show a positive number. One caution during monitoring is that measurement is difficult when the patient’s body is moving or when the blood perfusion rate is very low. Furthermore, ORi values may fluctuate if a pigmented drug is administered intravenously. There is a report showing that ORi rapidly decreased after indigo carmine was administered.
Outlooks of ORi

As mentioned above, in recent years, ORi has been used as an index of oxygenation during surgery. However, in the near future, ORi may also be used in situations in which changes in oxygenation occur frequently, such as in patients in the ICU and general wards. Yoshida et al. suggested the usefulness of ORi for patients in the ICU, when suctioning through an endotracheal tube is performed during ventilation [58]. Furthermore, the fact that the ORi device is noninvasive and easy-to-wear will enable its use in various clinical situations. Recently, COVID-19 has become a worldwide epidemic. ORi might be useful to prevent infections, such as COVID-19, by reducing the frequency of contact of clinical workers with patient blood when performing blood gas analysis or securing an arterial line. The use of ORi will not only enable the early prediction of hypoxemia, but may also be useful for avoiding hyperoxia. In recent years, there have been reports that the mortality rate of patients managed for hyperoxia is high [59]. Thus, in the near future, ORi monitoring may become an effective method to lower the adverse effects caused by high oxygen concentrations. ORi not only determines the optimal inhaled oxygen concentration during the perioperative period, but also acts as an index of proper oxygenation, and has the potential to help avoid complications caused by hypoxemia and hyperoxia. Studies on how the use of ORi can change patient prognoses should be performed in the near future.

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