EFFECT OF "LIDOCAINE" INFUSION ON SEVOFLURANE REQUIREMENT DURING BIS GUIDED CARDIAC SURGERY

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ABSTRACT

The need for sevoflurane is reduced in the patients by intravenous lidocaine, in an amazing way when compared to the patients who did not obtained lidocaine. The double-spectrum indicator is a non-interventional device. When anesthesia is given for the patients during surgery or severe diseases or injuries, the brain-mapping device is used to indicate the patients' severity. This is done by reversing the signals, received by the device.

The reason of the study

The effect of lidocaine, when added to the amount of sevoflurane administered to the patients is demonstrated in this study. The tests are done by comparing the lidocaine received patients, with the patients who were given sevoflurane without lidocaine, with the use of the double-spectrum indicator.

Pathology and Methods of Work

The research was performed from June 2014 to December 2014 at the Erbil Center for Cardiothoracic Surgery with 50 patients, who were separated into two different groups. Lidocaine was administered to the first group of 25 patients and the second group of 25 patients was not treated with lidocaine.

Results

In group A patients, the rate of sevoflurane concentrations was 1.68. This reduced to 1.44% at sternotomy and 1.2% during bypass. The rate of sevoflurane administered in group B was 1.36%, as soon as anesthesia was given. This has reduced to 0.84% at sternotomy and 0.4 during bypass.

Conclusions

The amount of sevoflurane that needs to be administered to the patients through assessment using the double-spectrum indicator is drastically reduced by the intravenous use of lidocaine, while giving anesthesia.

KEYWORDS: Sevoflurane, Lidocaine Received & Anesthesia

INTRODUCTION

Bispectrol Index

Previously, the effects of anesthetics on the brain were not monitored by the anesthesiologists, in terms of
"adequacy" or "depth" of anesthesia. Indirect assessment has been done on sedation by using subjective sedation or vital signs scales. However, under-sedation and over-sedation remains a major challenge, due to the restrictions in the subjective assessment tools. It is assumed that the consciousness is blocked by general anesthetics by suppressing the central nervous system. It is anticipated that the anesthesia’s adequacy is related to some components of the electroencephalogram (EEG), since it measures the electrical motion of the brain. The hypnotic element of a continuous EEG parameter is the bi spectral index (BIS) that ranges from an awake, no drug effect value of 95 to 100 to no measurable activity of EEG with a zero value.

The BIS monitor system includes a digital signal converter, a sensor, and a monitor. The electrical signals are picked up from the cerebral cortex by the sensor and passed on to the digital signal converter. Then, the digitized signals move to the pre-processor of the device, which screens the “artifacts” (stray high-frequency signals), which are formed as a result of electro cautery equipment or patient movement. To determine the bi spectral index, the filtered EEG data is then subjected to a sophisticated algorithm, which is in the numerical level between 0-100. The BIS readings are as follows: 90-100 = fully awake; 80-90 = light sedation; 60-80 = moderate sedation; 40-60 = deep sedation; 0 = no EEG activity;

It has been recommended that the BIS can be used for titrating volatile anesthetics accurately according to specific requirements. This helps to ensure that the exposure to unwanted increased concentrations of anesthetics while decreasing the chances of mindfulness during anesthesia.

![Diagram of BiSpectral Index Values](image)

**Figure 1: Diagrammatic Representation of BiSpectral Index Values**

**LIDOCALNE**

**Pharmakeokinetics**

As a local anesthetic, the efficiency of lidocaine is categorized by a speedy action in-between the period of efficacy. Thus, lidocaine is appropriate for block, infiltration, and surface anesthesia. Numerous available formulations can be utilized before intubations, endoscopies, etc. pH buffering of lidocaine results in less painful local freezing. For short ophthalmic procedures, lidocaine drops are used on the eyes.
Lidocaine is one of the most significant class-Ib antiarrhythmic drug; for treating ventricular arrhythmias (for digoxin poisoning, cardio version, acute myocardial infarction, and cardiac catheterization) it is used intravenously if amiodarone is contra-indicated or not available. For this indication, it should be given after defibrillation. Vasopressors and CPR have been started. Lidocaine can be used as a cough suppressor (antitussive). When inhaled, it acts peripherally and reduces the reflex of cough.

This application can be used as a security and relief for intubated patients, since it decreases the coughing frequency and any other damage to trachea, it could cause when developing from anesthesia.

Pharmacodynamics

Signal conduction in neurons is altered by anaesthesia lidocaine by hindering the speed voltage-gated Na+ channels in the cell membrane of neurons that are accountable for the propagation of signal. With adequate blockage, the postsynaptic neuron’s membrane will not depolarize and therefore will fail to conduct an voltage-gated Na+ channels. The anesthetic effect is created by not only avoiding pain signals from spreading to the brain, but also by preventing them before they could start. A high degree of selectivity is allowed by the cautious titration in the sensory neurons’ blockage, whereas advanced concentrations also affect neuron signaling’s other modalities.

Antiarrhythmic

For the action of drug in the heart, the same principle is applied. The depolarization threshold is raised by blocking the sodium channels in the conduction system as well as the heart’s muscle cells, which will make the heart to decrease to start or perform early action of 241 potentials that causes an arrhythmia.

Sevoflurane

Pharmacology

Sevoflurane acts mainly as GABA_A receptor’s positive allosteric modulator. It also potentiates glycine receptor currents, acts as an antagonist for NMDA receptor, inhibits nACh and 5-HT_3 receptor. Sevoflurane is breath in as an aesthetic, which is used to make children asleep during surgery. However, while waking up from the medication, it is observed that it causes delirium and agitation. Still, it is not known whether this can be stopped.

Pharmacokinetics

Among the recently released anesthetics for clinical use, Sevoflurane is comparatively added recently to the range of inhaling aesthetics. When compared to the older inhalational agents such as halothane or isoflurane, the most significant property of sevoflurane is that it is less soluble in the blood. The effects are rapid uptake and stimulation than the other inhalational agents, faster removal and rescue, and improved depth control of an anesthesia. The faster pharmacokinetics is an outcome of the gas partition/low blood coefficient of 0X.9. The minimum alveolar concentration (MAC) of sevoflurane is 2.05%, with an oil/gas partition coefficient of 47.2. Liver metabolizes 2-5% of the drug utilized. The pharmacokinetics of sevoflurane do not change in obese patients, children and in patients with renal insufficiency.

Mask induction is made feasible by pharmacokinetics and pleasant odour of sevoflurane, which is a benefit in pediatrician aesthesia. The formation of inorganic fluoride is resulted from the hepatic metabolism of sevoflurane. A minor amount of sevoflurane is decreased upon contact with alkaline CO2 absorbent, and the
formation of a metabolite (compound A) is seen, which is inhaled in minor amounts. The question whether the compound A and inorganic fluoride are nephrotoxic is currently a subject of controversy.

**Pharmacodynamics**

Sevoflurane is an inhalation anesthetic agent that is used for induction and for maintaining general anesthesia. The administration of Sevoflurane is connected with a smooth, fast loss of consciousness during inhalation and a rapid retrieval when anesthesia is discontinued.

For a 40-year-old adult, minimum alveolar concentration (MAC) of sevoflurane in oxygen is 2.1%. The MAC of sevoflurane reduces with the addition of nitrous oxide and with age.

Induction is accompanied with a minimum of upper respiratory irritation signs or excitement, no central nervous system stimulus and no indication of too much secretion within the trachea bronchial tree. Inspired concentration is rapidly followed by the changes in the depth of sevoflurane anesthesia. Induction and recovery times were decreased in child patients who received sevoflurane.

**Adverse Effects**

Intracranial pressure is raised by sevoflurane and it can lead to respiratory despair. Studies inspecting a current important health concern, neurotoxicity induced by anesthetia (includes sevoflurane, especially with infants and children) are troubled with confounders, and most are statistically underpowered, and so are debated to request more data either to back up or contradict the potential link.

The safety of anesthesia when used in infants and children is concerning. The preclinical evidence from appropriate animal models have shown that the commonly used clinical essential agents, including sevoflurane, could turn to be neurotoxic to the developing brains, thus, causing neurobehavioral anomalies in the elongated term. As of 2010, two studies, PANDA and GAS (large-scale clinical studies) are going on; to supply further important data on neuro developmental effects of general anaesthesia on young children and infants, even when sevoflurane is used.

**PATIENTS AND METHODS**

**Study Population**

This study was conducted from June 2014 to December 2014 in Erbil cardiac center after getting permission from its ethical committee. For this study, 50 patients were selected as the candidates. Selection was done with exclusion criteria consisting of patients with history of seizures, patients with history of lidocaine reaction, or the ones using drugs that affect BIS. Standard monitors were used including non-invasive and invasive arterial pressure, ECG, central venous pressure and pulse oximetry. BIS was also used to evaluate anesthesia’s depth. Anesthesia was induced after 8 hours of fasting using fentanyl 2 mcg/kg i.v. followed after 3 min by propofol 1.5 -- 2.5_1 mg kg_1 iv. When BIS touched less than 50, atracurium 0.5 mg kg i.v. was given to permit tracheal incubation. With a tidal volume of 7-8 ml/kg and an oxygen fraction of 0.4 in air, the lungs were ventilated. To maintain an end tidal CO2 between 30 and 35 mm Hg, the respiratory rate was adjusted.

Patients were randomly distributed into 2 groups. Group B patients received 1.5 mg/ kg bolus of 2% lidocaine i.v. followed by 2 mg/kg/h infusion, whereas the patients in group A received saline in equal volumes.
The concentration of Sevoflurane was adjusted to keep BIS between 40 and 60 during anesthesia maintenance. Sevoflurane was decreased or increased by 0.5% if BIS goes out of the range. End-tidal sevoflurane concentration and BIS values were also noted as per the stages of the surgery. Lidocaine and sevoflurane were discontinued, before the Cardiopulmonary Bypass. Using independent t-test, normally distributed continuous data were analyzed.

A P value of < 0.05 was measured as statistically significant.

**Demographics and Statistical Analysis**

To collect the data, a data collection sheet was designed (Fig. 6). The collected information was managed via SPSS v.19 computer program. From each parameter, data was expressed as mean± standard deviation. The measure for statistical significance is independent t-test at p value < 0.05.

**Table 1: Date Collection Sheet**

<table>
<thead>
<tr>
<th>Name:</th>
<th>age:</th>
<th>sex:</th>
<th>weight:</th>
<th>Group:</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Date of operation:</td>
<td>Type of operation:</td>
<td></td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

**RESULTS**

Among the 50 patients involved in our study, 9 (18%) were female and 41 (82%) were male. Mean weight was 7+2 and the mean age was 65.26 ± 34.22 years.

50 patients were separated into 2 groups: Control (A) group (n= 25) and Lidocaine (B) group (n=25). In this study, no patients were excluded. As per the data available in table 2, the mean dose of total bolus and infusion lidocaine was 398 ± 11(mg). Immediately after induction of anesthesia, the mean concentration of Sevoflurane in the control group (A) was 1.68, 1.2 at pre-cardiopulmonary bypass stages and 1.44 at sternotomy.

In the lidocaine group (B), the mean concentration of Sevoflurane was 1.36 directly after anesthesia induction, 0.4 at pre-cardiopulmonary Bypass stages and 0.84 at sternotomy (see table 3).

A significant difference was observed between the 2 groups based on the independent t-test concerning the doses of sevoflurane immediately after induction of anesthesia, stjerniSicony and pro-cardiopulmonary Bypass stages.

**Table 2: Patients and Lidocaine Parameters**

<table>
<thead>
<tr>
<th>Parameter</th>
<th>Control (group A) (N=25)</th>
<th>LIDOCAINE (GROUPE B) (N=25)</th>
<th>Total (n=50)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age (Years)</td>
<td>67.32 ± 28.3</td>
<td>65.92±30.2</td>
<td>65.26±34.2</td>
</tr>
<tr>
<td>Weight (Kg)</td>
<td>78+3</td>
<td>73+3</td>
<td>76+2</td>
</tr>
<tr>
<td>Gender Male</td>
<td>21(84%)</td>
<td>20 (80%)</td>
<td>41 (82%)</td>
</tr>
<tr>
<td>Female</td>
<td>4(16 %)</td>
<td>5 (20%)</td>
<td>9(18%)</td>
</tr>
<tr>
<td>Lidocaine dose</td>
<td>398+11</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

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Table 3: Mean Dose of Sevoflurane

<table>
<thead>
<tr>
<th>Stage of Operation</th>
<th>Control (Group A) (n=25)</th>
<th>Lidocaine (group B) (n=25)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Immediately after induction of anesthesia</td>
<td>1.68±0.25</td>
<td>1.36±0.1</td>
</tr>
<tr>
<td>Sigmoidomy</td>
<td>1.44±0.06</td>
<td>0.84±0.07</td>
</tr>
<tr>
<td>Pre-cardiopulmonary Bypass</td>
<td>1.2±0.1</td>
<td>0.4±0.01</td>
</tr>
</tbody>
</table>

DISCUSSIONS

This research shows that when lidocaine is given at a dose of 1.5 mg/kg bolus followed by 2 mg/kg/h, it reduces the requirement of sevoflurane while keeping the BIS score between 40 and 60, during anesthesia maintenance in patients going through cardiac surgeries.

In this research, a significant statistical difference between the 2 groups have been found, which approve Wilson ET at who noticed in a research that lidocaine, without or with ketamine considerably decreased the MAC of sevoflurane in dogs.

Matsubara et al. also observed that intravenous lidocaine reduced MAC of sevoflurane in the dogs that are anesthetized. Another study showed that, lidocaine dose reduced the MAC of Sevoflurane in cats (Pypendop and 1lkiw).

The research conducted by us doesn’t agree with Ahmed M. Omar et al. as it shows no statistical difference of MAC of sevofluranc between the 2 groups in human.

When cardiac surgery is performed for the patients included in this research, there was a reduction in ETSesaduraneduring the operation or along the stages of operation, also due to the use of fentanyl 10 meg/kg/dose with midazolam 10 ingi dose at stematomy. Due to lidocaine’s action at the spinal level, by reducing the motor response, the effect of lidocaine on MAC of Sevofluranc is noted.

However, it was detected that the intravenous infusion of lidocaine decreased the requirement of bispectral index-guided sevoflurane during anesthesia.

The rate of infusion of lidocaine and the bolus dose used in this research was (398 + 11 mg) that was on par with the earlier studies which showed that this dosage did not end in plasma concentration more than 4 Meg/ml, which is less than the toxic levels.

CONCLUSIONS

During the maintenance of general anesthesia, intravenous administration of lidocaine can significantly reduce the requirements of BIS-guided sevoflurane

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