Intraoperative neuromonitoring alerts in a pediatric deformity center

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OBJECTIVE Intraoperative neuromonitoring (IONM) involves the use of somatosensory evoked potentials (SSEPs) and transcranial electric motor evoked potentials (TceMEPs). In this retrospective study the authors examined the sensitivity and specificity of both SSEPs and TceMEPs during pediatric spinal deformity surgeries.

METHODS The authors performed a retrospective quantitative analysis of data obtained in 806 patients (197 males and 609 females) treated from December 2011 until October 2015. All patients were diagnosed with scoliosis that was classified as one of the following: adolescent idiopathic scoliosis (AIS) (38%), congenital scoliosis (22%), or syndromic scoliosis (40%). Also, 53 patients underwent vertebral column resection (VCR). All surgeries were monitored by high-level neuromonitoring specialists and were performed with total intravenous anesthesia. Alerts were described as a decrease in amplitude by 50% or greater (bilateral or unilateral) in SSEPs, TceMEPs, or both.

RESULTS True-positive alerts for TceMEPs were observed in 60 of the 806 patients (7.4%). True-positive alerts for SSEPs were observed in 7 of the 806 patients (0.9%). In contrast, there were no false-positive or false-negative outcomes. Only 1 case (0.1%) was reported with a permanent postoperative deficit. No reported false negatives or false positives were observed, and thus sensitivity was 100% and specificity was 93%–100% for TceMEPs. The rate of sensitivity was 13.2% and the rate of specificity was 100% for SSEPs. The breakdown of total alert was as follows: 6.6% in AIS cases, 24.5% in congenital scoliosis cases, and 10.2% in syndromic scoliosis cases. Neurological injury rates were significantly lower than in previous studies, as there were 0% for AIS cases (p = 0.12), 0.6% for congenital scoliosis cases (p = 0.17), and 0% for syndromic scoliosis cases (p = 0.07). One injury in a patient with congenital scoliosis occurred during a VCR procedure, which brought the injury rate to 1.9% (p < 0.005). IONM alerts occurred during 34% of rod/correction cases, 25% of thoracic screw placements, 20% of the osteotomies, 17% of the resections, 3% of the cage insertions, and 2% of the sublaminar wiring procedures.

CONCLUSIONS The authors hypothesize that the results of this study will support the necessity, as a standard of care, of multimodality neuromonitoring during high-risk pediatric spinal deformity surgery because of the decrease in postoperative deficits. Their data suggest that the TceMEPs are more sensitive than SSEPs, but when used in combination, they offer the patient a level of safety that would otherwise not exist. Last, these findings support the notion that better outcomes are achieved with high-level IONM professionals.

KEY WORDS intraoperative neuromonitoring; transcranial electric motor evoked potential; somatosensory evoked potential

In the United States, the vast majority of pediatric deformity surgeries are performed while using intraoperative neuromonitoring (IONM). If reliable transcranial electric motor evoked potentials (TceMEPs) are present, they can be used intraoperatively as a substitute for performing the wake-up test.1 However, multiple factors may influence the ability to generate baseline evoked potentials in certain pediatric populations.1,2,12 Because of their associated risks of complete or partial paralysis, surgical procedures that may benefit from the use of evoked potential monitoring include the initial placement of both expandable spinal rods and vertical expandable prosthetic titanium rods (VEPTRs), the initial lengthening of both rod types, and vertebral column resections.1,2
Certain variables may influence the ability to generate baseline evoked potentials, including the patient's chronological age, cognitive ability, and a history of neurological disorders. With regard to a patient's chronological age, the current literature suggests neuronal pathways and the myelination of these pathways do not fully develop until after the age of 2 years. Therefore, obtaining evoked potentials in this young population poses a great challenge for the neurophysiology professional. However, both somatosensory evoked potentials (SSEPs) and TceMEPs are deemed safe for this population, and by all means monitoring should be attempted prior to incision. Also, cognitive ability has been associated with obtaining baseline potentials that can be monitored. Specifically, the quality of the cortex being stimulated may determine the robustness of baseline data. Last, neurological disorders, such as seizures, were once thought to be a contraindication to TceMEPs. According to MacDonald and colleagues, the incidence of seizures was 0.03% and the incidence of bite injuries was 0.2%. Therefore, a risk/benefit analysis should be performed prior to utilizing TceMEPs.

For over 20 years, surgeons have had access to modalities such as TceMEPs and SSEPs during their surgical procedures. In high-risk surgeries, such as those used in spinal deformity correction (i.e., scoliosis reduction) a 0.3%–0.8% chance of a postoperative deficit has been observed. Multimodality neuromonitoring of SSEPs and TceMEPs allows for a greater level of specificity regarding patient outcome after spinal deformity surgery. Nuwer et al. observed a sensitivity level of 92% and a specificity level of 98% for SSEP monitoring, in terms of new postoperative motor deficits. SSEP monitoring evaluates the integrity of the dorsal column's medial lemniscal pathway. The dorsal columns are located in the posterior two-thirds of the spinal cord, and blood is supplied by 2 posterior spinal arteries. Three limitations of SSEP monitoring exist. First, the small computerized averaged response lacks immediate feedback through data collection. Second, SSEPs do not reveal the integrity of motor tracts. Third, there may be a delay in response time. The current literature suggests that such a lag time may be 5 to 16 minutes.

In contrast, TceMEP monitoring allows for evaluation of motor function through electrical stimulation of the cerebral cortex, and also for evaluation of the integrity of the anterolateral cortical spinal tracts. Additionally, this tract is located in the anterior third of the spinal cord, and the blood is supplied through the anterior spinal artery. Moreover, because this tract is rich in cell bodies and synapses, it is more metabolically active. These factors suggest that monitoring TceMEPs can provide greater sensitivity for detecting ischemic changes to the spinal cord. Optimal recording sites for TceMEPs may include the abductor pollicis, abductor longus, and other long extensor/flexor muscles. These muscles are the preferred recording sites because of their rich corticospinal tract innervation within the cortex. However, the sensitivity of TceMEPs may be influenced by both hypoperfusion and hypothermia, and an effort to control these factors should be attempted. Moreover, the use of total intravenous agents (TIVAs) and a multipulse stimulation train allow for a greater advantage in overcoming anesthetic agents. Motor function may be monitored using TceMEPs, but TceMEP monitoring is sensitive to inhalation agents. Therefore, a TIVA anesthetic protocol is an optimal choice for maintaining sedation throughout the procedure. According to MacDonald and colleagues, the anterior portion of the cord contains a large number of cell bodies and synapses that are highly sensitive to ischemic changes. Last, the anterior portion of the cord has greater metabolic demands and fewer radicular arteries than the posterior portion of the cord. All of these factors suggest an increased sensitivity to ischemic changes that can be detected through TceMEP monitoring.

Although the benefits of multimodality monitoring are positive, some institutions still do not use this approach in pediatric spinal surgeries. This paper demonstrates the benefits of a multimodality approach and reports the levels of specificity that TceMEP and SSEP monitoring have achieved at our institution, which is a regional spinal deformity center.

### Methods

After obtaining approval from Temple University’s Institutional Review Board, we performed a retrospective review of the neuromonitoring and surgical records; the search yielded 809 patients who had undergone scoliosis surgery in which neuromonitoring was used during the period from December 2011 until October 2015. Both male and female patients were examined, and ages ranged from 1 to 20 years. Patients excluded from the study were those with a preexisting spinal cord injury or preexisting lower-extremity paralysis. Patients included in the study population had one of the following diagnoses: idiopathic, congenital, or syndromic scoliosis, which included neuromuscular and various syndromes. A data entry system was created for these patients’ cases and included the following variables: diagnosis, TceMEP alerts, SSEP alerts, surgical or anesthetic interventions, and time required for the alert to resolve. Alerts were defined as a drop of 50% in amplitude of either TceMEPs or SSEPs. In addition, an alert needed to be accompanied by a surgical event that posed a risk of spinal cord trauma (e.g., screw placement, osteotomy, rod correction, or derotation).

Continuous neurophysiological monitoring was performed from the time the patient was positioned until 30 minutes after the last major corrective force was applied to the rods that were placed. Both SSEPs and TceMEPs were monitored using a Cascade Elite neuromonitoring system (Cadwell). Potentials were recorded and interpreted by a group of 2 nationally certified trained individuals who had successfully completed multiple graduate-level neuroanatomy/physiology educational courses and who were present throughout the entire procedure. All IONM changes were
evaluated by these individuals utilizing the alert criteria. Anesthesia consisted of TIVAs administered in all of the patients after positioning. Inhalational agents and neuromuscular blockade were administered for induction only. The level of the neuromuscular blockade was measured by the neurophysiologist using the train-of-four technique. Train-of-four data were recorded on the Cascade system from both the ulnar nerve and the posterior tibial nerve. Other IONM modalities recorded included triggered and spontaneous electromyography for lumbar and thoracic pedicle screw testing and nerve root irritation detection, respectively. Electroencephalography was used in an effort to correlate the frequency of brain activity with the amplitude of TceMEPs and cortical SSEPs.

SSEPs of both upper and lower extremities were monitored by stimulation of the ulnar nerve and posterior tibial nerve, respectively. Also, subdermal needle electrodes were positioned on the posterior area of the tibial femoral joint in an effort to record from the popliteal nerve. Data on this nerve were collected to ensure that proper technical setup was achieved. Cortical recording electrodes were placed using standard international 10/20 system sites and consisted of Cp3, Cp4, and Cpz, with reference to Fpz. Subcortical and cervical volleys were recorded from the additional placement of a cervical electrode over the area encompassing C-5, and again referenced to the Fpz electrode. SSEPs were considered suitable for monitoring if bilateral lower-extremity responses were present and at least 1 μV in amplitude.

TceMEPs were recorded on both the upper and lower extremities. Relative contraindications were investigated, but all patients underwent TceMEP monitoring. Stimulation was achieved through subdermal bent needles placed anterior to the spinal sulcus. The upper-extremity recording site consisted of the abductor pollicis brevis and served 2 purposes: to control for technical or anesthetic issues that may arise during the surgery and to detect a brachial plexopathy. Additionally, lower-extremity subdermal electrodes were positioned over the vastus lateralis, gastrocnemius, tibialis anterior, and abductor hallucis brevis muscles. Additional subdermal electrodes were placed in the intercostal and rectus abdominis regions, solely for the purpose of thoracic pedicle screw stimulation. TceMEPs were considered suitable for monitoring if they were bilaterally present in all monitored lower-extremity muscle groups with a minimum amplitude of 200 μV.

**Definition of an Alert**

A significant change was defined as a 50% decrease of TceMEP or SSEP amplitude during critical surgical times. The drop in amplitude was also significant in unilateral and/or bilateral data responses. Moreover, this drop in amplitude needed to be repeated twice and be unresponsive to an increase in voltage from stimulation. Latency was not used as a determinant of a significant change resulting in neurological impairment. Critical surgical times consisted of the following: drilling for and placing of pedicle screws, osteotomies, placement of rods, manipulation of rods, and placement of interbody spacers. Also, global decreases in amplitude were not considered significant changes and were correlated with changes in anesthetic infusion rates and/or blood loss, thus reducing false-positive alerts.

**Intervention**

Each significant change was subjected to an operating room-generated checklist (Fig. 1) pertaining to loss of IONM signals; the checklist was developed by the neurophysiology team based on the work of Vitale et al. The goal of the checklist was to encourage the surgical team to focus on increasing spinal cord perfusion for the patient. The first step consisted of a surgical pause, followed by the anesthesia team increasing the mean arterial pressure above 80–90 mm Hg or 10%–15% above the resting pressure. Next, hemoglobin was examined, and if the current recording was below 10 g/dl, blood was administered. Finally, if the surgical procedure allowed for reversal of the manipulation (i.e., removal or screw or rod), the surgeon would remove the instrumentation. However, if the procedure was irreversible (i.e., in the case of a vertebral column resection), the spine was stabilized and the procedure was aborted. Immediate postoperative evaluations were performed by the IONM professionals to ensure accurate measures.

**Statistical Analysis**

A database was generated from cases treated from
December 2011 to October 2015. Patient data included surgical procedure, TceMEP alerts, SSEP alerts, surgical intervention, anesthetic intervention, and time until recovery of evoked potentials. A significant IONM change was defined as a loss of 50% of amplitude of either TceMEPs or SSEPs coinciding with surgical interventions. Neuro-monitoring alerts were determined by the aforementioned criteria and then categorized utilizing Bradford Hill causality guidelines; Skinner and Sala determined that using causality guidelines can help determine a true-positive change based on the causal relationship between the intervention and recovery of the evoked response.\textsuperscript{10} Therefore, a true-positive change was defined as a decrease in amplitude of response based on the alert criteria, correlated with a surgical event, which either recovered with intervention or which remained at closing and correlated with a postoperative neurological impairment seen upon the patient’s emergence from anesthesia. A false-negative change was defined as a loss of amplitude that did not return at closing, yet the patient emerged from anesthesia with no new neurological impairment. A true-negative change was defined as no loss of amplitude and no new neurological impairments upon emergence from anesthesia. RSCs were all considered false positives; and in the third, causality guidelines were applied to the RSCs to classify them. For each group, specificity and PPV were calculated.

**Results**

A multimodality approach to neuromonitoring was used in all patients; patient and surgical procedure data are presented in Table 1. The total number of patients examined was 806 (197 male and 609 female). Ages ranged from 1 to 20 years (median 12.5 years). Cobb angles ranged from 21° to 180° (median 66.2°). Of the 806 patients, 511 underwent a posterior spinal fusion. The posterior spinal fusions were categorized into adolescent idiopathic (n = 136), congenital (n = 53), and syndromic (n = 322) scoliosis; also, 52 patients underwent vertebral column resection. Additional spinal procedures consisted of VEPTR (n = 68), growing rod (n = 59), vertebral body stapling (n = 41), and growth modulation (n = 127). TceMEP and SSEP monitoring were attempted pre-incision in all 806 patients.

For the 806 patients in the study population, the number of true-positive TceMEP alerts was 60, and the number of true-positive SSEP alerts was 7 (Table 2). In contrast, no reported false-positive or false-negative outcomes were observed. Additionally, there was 1 reported postoperative deficit. Sensitivity and specificity rates for TceMEP alerts were both 100%. In the above calculations, it was assumed that all RSCs were in fact true positives. To determine a range, the RSCs were broken into 3 groups. In the first group, assuming all RSCs were indeed true-positive TcemeP alerts, specificity was calculated to be 100%, with a PPV of 1. In the second group, assuming all RSCs were false-positive TceMEP alerts, specificity was 93%, with a PPV of 0.08. In the third group, applying causality guide-
lines to RSCs, specificity was 100%, with a PPV of 0.98. Thus, the specificity range for TceMEP alerts was 93% to 100%. In comparison, the sensitivity and specificity for SSEP alerts were 13.21 and 100%, respectively. IONM alert data according to surgery type are presented in Fig. 2 and Table 3.

The surgical causes for each neuromonitoring alert were thoracic screw placement (25%), osteotomy (20%), cage placement (3%), wiring (2%), resection (17%), and rod/correction (34%) (Fig. 3). Neurological deficit rates were determined to be as follows: idiopathic fusion, 0%; congenital fusion, 0%; syndromic fusion, 0%; and vertebral column resection, 1.9% (Table 4, Fig. 4).

**Discussion**

Multimodality monitoring that uses both SSEPs and TceMEPs has been shown to increase the sensitivity of this monitoring. This paper demonstrates the benefits of using the multimodality approach in severe spinal deformity surgery in a pediatric population. Through the use of this type of monitoring by well-trained and educated neurophysiology professionals, only 1 postoperative deficit was reported out of 806 cases studied; this isolated case occurred during a high-risk procedure involving a vertebral column resection, and interventions were performed by the surgical team at the time of TceMEP loss. The re-
sults of this study indicate that TceMEP monitoring had a sensitivity of 100% and a specificity ranging from 93% to 100% in detecting spinal cord injury. We also demonstrated that TceMEPs were more sensitive than SSEPs (13.21% sensitivity) in detecting spinal cord injury. Therefore, our findings suggest that TceMEP monitoring during pediatric spinal deformity surgery can significantly reduce the risk of postoperative deficits. Furthermore, TceMEPs are often used for high-risk spinal deformity surgery, and this is consistent with previous research.1,4

Unfortunately, a non-uniform system exists within the field of IONM. Variables that make IONM non-uniform may include the following: use of inhalational agents, slow-charge TceMEP stimulators, data not being interpreted in the room at the time of data loss, and a lack of standard criteria by which to identify and classify an alert. In our study, anesthesia was performed by a consistent group and was strictly managed throughout the procedure. The population base at our institution is often composed of patients who have multiple syndromes or cognitive impairments. Therefore, the use of inhalational agents may not allow for maximum data to be collected due to a lack of quality cortical matter, physiological abnormalities, and immature neurological pathways.1 A TIVA protocol was used in all 806 cases, and neuromuscular blockade was only administered for intubation and exposure. The level of neuromuscular blockade was measured using the train-of-four technique from both the ulnar and posterior tibial nerves. Additionally, all 4 twitches returned after exposure was complete. The level of blockade was monitored carefully, communicated clearly to the surgeons and anesthesia team, and documented frequently by the neurophysiology professional in the room. This allowed for in-the-room interpretation by the monitoring individual, ensuring that interventions were performed in a timely manner. No remote monitoring was involved in any of the 806 cases. To our knowledge, this model of IONM professionals is one of the few of its kind in the United States. More impressive is the ability of this team to generate TceMEPs in 98% of patients and to have a TceMEP-related specificity of 93%–100% and a sensitivity of 100%. Historically, this was the predominant IONM practice model prior to the mid-to late 1990s, when vendor remote models started to appear due to financial incentives. According to Skinner and Sala, surgeons should no longer accept “remote” monitoring as it is practiced in the United States.10 Our model is consistent with that used in Canada and Europe, and it is a more cost-effective model for any hospital that requires neuro-monitoring for its surgical procedures.

Limitations of this study are that it was performed at a single pediatric institution, which means that its findings may not be applicable to an adult center. This study is also limited by its retrospective design. Although all IONM data, surgical procedures, and immediate postoperative evaluations were performed by only 2 IONM professionals, long-term outcomes were subject to postoperative chart review. Also, IONM was conducted by professionals who had significant expertise in pediatrics, and the results of this study may not be able to be extrapolated to centers.

<table>
<thead>
<tr>
<th>Surgical Procedure</th>
<th>Total No.</th>
<th>TceMEP Alerts (%)</th>
<th>SSEP Alerts (%)</th>
<th>Injury Rate (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Posterior spinal fusion</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>For AIS</td>
<td>136</td>
<td>6.6</td>
<td>0.7</td>
<td>0.0</td>
</tr>
<tr>
<td>For congenital scoliosis</td>
<td>53</td>
<td>24.5</td>
<td>5.7</td>
<td>0.0</td>
</tr>
<tr>
<td>For syndromic scoliosis</td>
<td>322</td>
<td>10.2</td>
<td>0.9</td>
<td>0.0</td>
</tr>
<tr>
<td>VCR</td>
<td>53</td>
<td>34.0</td>
<td>5.7</td>
<td>1.9</td>
</tr>
<tr>
<td>Other procedures</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>VEPTR</td>
<td>68</td>
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<td>0.0</td>
<td>0.0</td>
</tr>
<tr>
<td>Growing rod</td>
<td>59</td>
<td>5.1</td>
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</tr>
<tr>
<td>VBS</td>
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<td>0.0</td>
<td>0.0</td>
<td>0.0</td>
</tr>
<tr>
<td>Growth modulation</td>
<td>127</td>
<td>1.6</td>
<td>0.0</td>
<td>0.0</td>
</tr>
</tbody>
</table>
whose monitoring staff does not have this level of expertise.

**Conclusions**

The authors hypothesize that the results of this study will support the necessity, as a standard of care, of multimodality neuromonitoring during high-risk pediatric spinal deformity surgery due to the decrease in postoperative deficits. Their findings suggest that TceMEP monitoring is more sensitive than SSEP monitoring, but when used in combination they offer the patient a level of safety that would otherwise not exist. Last, these findings support the notion that better outcomes are achieved with high-level IONM professionals.

**References**

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**Disclosures**

Dr. Hwang reports being on the speaker’s bureau of Zimmer Biomet. Dr. Pahys reports being a consultant for DePuy Synthes, Globus Medical, and Zimmer Biomet. Dr. Samdani reports being a consultant for DePuy Synthes Spine, Ethicon, Globus Medical, Misonix, Stryker, and Zimmer Biomet.

**Author Contributions**

Conception and design: J Zuccaro, M Zuccaro. Acquisition of data: J Zuccaro, M Zuccaro. Analysis and interpretation of data: J Zuccaro, M Zuccaro. Drafting the article: J Zuccaro, M Zuccaro. Critically revising the article: all authors. Reviewed submitted version of manuscript: all authors. Approved the final version of the manuscript on behalf of all authors: J Zuccaro. Statistical analysis: J Zuccaro, M Zuccaro. Study supervision: J Zuccaro.

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