A Comparison of Interside Asymmetries of Lower Extremity Somatosensory Evoked Potentials in Anesthetized Patients with Unilateral Lumbosacral Radiculopathy

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Study Design: Prospective cohort study.

Purpose: This study was to investigate interside asymmetries of three lower extremity somatosensory evoked potentials (SSEPs) in anesthetized patients with unilateral lumbosacral radiculopathy.

Overview of Literature: Although interside asymmetry is an established criterion of abnormal SSEP, little is known which of the lower SSEPs is more sensitive in detecting interside asymmetry in anesthetized patients.

Methods: Superficial peroneal nerve SSEP (SPN-SSEP), posterior tibial nerve SSEP (PTN-SSEP), and sural nerve SSEP were obtained in 31 lumbosacral surgery patients with unilateral lumbosacral radiculopathy, and compared with a group of 22 control subjects.

Results: The lumbosacral group showed significant larger interside asymmetry ratios of P37 latencies in SPN-SSEP and PTN-SSEP, and significant larger interside asymmetry ratio of P37-N45 amplitude in SPN-SSEP, when comparing with the control group. Within the lumbosacral group but not the control group, SPN-SSEP displayed significant larger interside asymmetry ratio in P37 latency. When referencing to the control group, more patients in the lumbosacral group displayed abnormal interside SPN-SSEP latency asymmetries which corroborated the symptom laterality.

Conclusions: The data suggested that SPN-SSEP was more sensitive in detecting interside latency asymmetry in anesthetized patients.

Keywords: Superficial peroneal nerve; Posterior tibial nerve; Sural nerve; Somatosensory evoked potential; Radiculopathy

Introduction

Interside asymmetry significantly beyond the control level is an important criterion of abnormal somatosensory evoked potential (SSEP) [1,2]. In theory, this parameter can be easily applied to anesthetized patients as the within-subject ratio is less likely subject to anesthetic influence.

For decades, post-tibial nerve SSEP (PTN-SSEP) has proven to be well suited for central nervous system evaluation [3]. However, it is still unclear of its adequacy in detecting lumbar nerve root injuries owing to the multiroot (L4–S3) inputs [4,5]. Superficial peroneal nerve and sural nerve mainly represent L4–5 and S1–2 dorsal roots,

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respectively [6]; these cutaneous nerve derived SSEPs have been successfully used in anesthetized patients with signal-to-noise ratios comparable to PTN-SSEP [7,8].

This study is to determine which of the lower SSEPs is more sensitive in detecting interside asymmetry in anesthetized patients by a comparison between PTN-SSEP, superficial peroneal nerve SSEP (SPN-SSEP), and sural nerve SSEP (SN-SSEP).

Materials and Methods

Lower extremity SSEPs were obtained from both lumbosacral and control patient groups. In the lumbosacral patient group, surgeries were performed in 31 patients who were unilateral symptomatic (mean age±standard deviation, 54.9±15.1; 19 females and 12 males). Of the 31 lumbosacral patients, sixteen were left symptomatic; 15 right symptomatic. Seventeen surgeries were performed at or above the L5 vertebral level; 14 surgeries involved the S1 and adjacent vertebral levels. Determination of the surgical level(s) was based on the patient's symptoms, neurological examination and medical imaging findings. The control group consisted of 22 patients (mean age±standard deviation, 54.4±8.0; 13 women and 9 men) who underwent anterior cervical discectomy and fusion. Clinical pictures of the control group were confined to cervical region without lumbosacral symptoms. All the surgical procedures were performed in a single facility under identical total intravenous anesthesia (TIVA) protocol. This study was approved by the Institutional Review Board and the informed consent was obtained from each patient.

Cortical SSEP waveforms derived from Cz' and Ci' montages were obtained by electrical stimulation of superficial peroneal, posterior tibial, and sural nerves at the ankle using a Cadwell Cascade (Cadwell Laboratories, Inc., Kennewick, WA). Specifically, the cathodes (stimulating electrodes) were placed (1) behind the medial malleolus for PTN-SSEP; (2) medial to the peroneus tertius at the level of lateral malleolus for SPN-SSEP; and (3) behind the lateral malleolus for SN-SSEP, respectively. Bilateral interleaving stimulation was sequentially presented to the three stimulus sites at intensity about 40 mA. The subdermal EEG electrodes were positioned at Cz'-Fpz, Ci'-Cc' for the cortical P37/N45 and at Fpz-C5Sp for subcortical P31/N34 recordings according to the international 10–20 standard. Electrode impedance was kept below five $k\Omega$

with inter-electrode balance within two k Ω . A total of 200 averages for each trial, filter bandpass of 30–1,000 Hz, and 100 milliseconds analysis epoch were used. Cortical SSEP peak (P37) latencies were measured from the offset of evoked stimulus; amplitudes were measured as the maximal magnitude of separation between peak and trough of P37-N45 complex. SSEPs were recorded after incision was completed but before the microscope was brought to the surgical field. To calculate the interside asymmetry ratios of P37 latency and P37-N45 amplitude, following equation was used: |(Lift-Right)|/(Left+Right).

The differences of interside asymmetry of SSEP P37 latencies and P37-N45 amplitudes between the lumbosacral and control groups were analyzed by independent *t* tests; whereas the interside asymmetries between SPN-, PTN-, and SN-SSEPs P37 latencies and P37-N45 amplitudes within either group were analyzed by repeated measures analysis of variance and *post-hoc* LSD pairwise comparison. Prolongation of interside latency or reduction of interside amplitude ratios beyond 2.5 standard deviations greater than the mean of control group was considered abnormal [1]. All statistical analyses were performed with SPSS ver. 19.0 (SPSS Inc., Chicago, IL, USA). Differences were considered significant when *p*<0.05.

Results

The interside asymmetry ratios of P37 latencies and P37-N45 amplitudes of SPN-, PTN-, and SN-SSEP were compared between the lumbosacral and control groups (Fig. 1). Significant difference in the interside asymmetry ratios of P37 latency between the groups were found in SPN-SSEP (t=2.105, p=0.04) and PTN-SSEP (t=2.079, p=0.043) (Fig. 1A). In addition, significant difference in the interside asymmetry ratio of P37-N45 amplitude between the groups were found in SPN-SSEP (t=2.142, p=0.037) (Fig. 1B). In contrast, SN-SSEP showed no significant difference in the interside asymmetry ratio between the groups.

The interside asymmetry ratios of P37 latency and P37-N45 amplitude were analyzed between SPN-, PTN- and SN-SSEP within either group (Fig. 2). In the lumbosacral group, significant differences in the interside asymmetry ratio of P37 latency were observed between SPN|PTN and between SPN|SN (SPN vs. PTN, p=0.026; SPN vs. SN, p=0.006) (Fig. 2A). No such differences were found in the control group (Fig. 2B). In addition, there was no significant difference in the interside asymmetry ratios of

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Fig. 1. Independent *t* test analysis of the interside asymmetry ratio of P37 latencies **(A)** and P37-N45 amplitudes **(B)** of superficial peroneal nerve (SPN)–, posterior tibial nerve (PTN)–, and sural nerve (SN)–somatosensory evoked potential (SSEP) between lumbosacral (L) and control (C) groups. Means and standard deviations were shown. **p*<0.05.



Fig. 2. Repeated measures analysis of variance analysis of the interside asymmetry ratios of superficial peroneal nerve (SPN)–, posterior tibial nerve (PTN)–, and sural nerve (SN)–somatosensory evoked potential (SSEP) P37 latencies in lumbosacral (A) and control groups (B), and P37-N45 amplitudes in lumbosacral (C) and control groups (D). Means and standard deviations were shown. *p<0.05.

Patient	Age/Sex	Surgical levels	SPN-SSEP			PTN-SSEP			SN-SSEP		
			Lt	Rt	Ratio	Lt	Rt	Ratio	Lt	Rt	Ratio
2	41/Male	Lt L4–5	48.4	43.7	0.051 ^{a)}	46.0	40.3	0.066 ^{a)}	42.1	41.4	0.009
6	73/Female	Lt L2-4	49.2	45.0	0.045 ^{a)}	49.0	45.2	0.041 ^{a)}	44.9	43.3	0.018
17	44/Female	Rt L4–5	42.7	48.4	0.063 ^{a)}	38.5	38.9	0.005	42.6	42.5	0.001
20	59/Female	Rt L3–5	42.8	46.6	0.042 ^{a)}	41.0	40.6	0.005	42.3	41.8	0.006
24	88/Female	Rt L4–5	51.1	44.2	0.072 ^{a)}	47.7	45.7	0.020	46.2	46.0	0.002

Table 1. Patients with abnormal interside latency asymmetry ratio

SPN, superficial peroneal nerve; SSEP, somatosensory evoked potential; PTN, posterior tibial nerve; SN, sural nerve; Lt, left; Rt, right. ^{a)}Exceed normative value. Latency in milliseconds.

 Table 2. Patients with abnormal interside amplitude asymmetry ratio

Patient	Age/Sex	Surgical levels	SPN-SSEP			PTN-SSEP			SN-SSEP		
			Lt	Rt	Ratio	Lt	Rt	Ratio	Lt	Rt	Ratio
2	41/Male	Lt L4–5	1.51	0.77	0.325	1.25	1.24	0.004	0.63	0.25	0.432 ^{b)}
19	49/Female	Rt L4–5	1.33	0.57	0.402 ^{a)}	1.36	1.51	0.052	1.15	0.81	0.173
31	65/Male	Rt L4–5	0.98	1.05	0.034	1.15	1.44	0.112	0.59	1.88	0.522 ^{b)}

SPN, superficial peroneal nerve; SSEP, somatosensory evoked potential; PTN, posterior tibial nerve; SN, sural nerve; Lt, left; Rt, right. ^{a)}Exceed normative value; ^{b)}Exceed normative value but opposite to the side of symptoms. Amplitude in millivolts.

P37-N45 amplitude between all the SSEPs in either the lumbosacral or control group (Fig. 2C, D).

When referencing to the normative values of interside latency (mean+2.5SD) and amplitude (mean-2.5SD) ratios derived from the control group, the patients with abnormal interside asymmetry ratios of P37 latency (Table 1) and P37-N45 amplitude (Table 2) were shown. A higher incidence of the abnormal latency asymmetry ratio was noted in SPN-SSEP (5/31, ~16%), followed by PTN-SSEP (2/31, ~6%). All the abnormal latency asymmetry ratios matched the symptom sides.

Discussion

As more advanced diagnostic tools have become available to aid the surgical decisions nowadays, SSEP is less frequently administered as part of the preoperative workups [4]. Nevertheless, studies had indicated that SSEP abnormalities strongly suggested a true compromise of the corresponding nerve root(s) in patients with radiculopathy, whereas the structural lesion on an imaging study did not always translate into abnormal axonal conductivity [9,10]. As such, our center uses the asymmetric SSEP as a surrogate marker of hidden culprits causing the root compression when the initial operative findings do not match the degree of SSEP abnormalities.

In lumbosacral patients, spondylotic radicular symptoms are often caused by L4 and L5 foraminal narrowing [9,10]. The higher degree and incidence of SPN-SSEP latency asymmetry was likely associated with its L4 and L5 origins. PTN-SSEP did not match the sensitive level of SPN-SSEP (PTN-SSEP ~6% vs. SPN-SSEP ~16%); it merely confirmed the PTN-SSEP findings. SN-SSEP showed the least amount of latency asymmetry, presumably due to the low incidence of S1 and S2 foraminal stenosis. Although S1 radiculopathy is common, its involvement is often bilateral due to the sacral fibers are more medially situated in the cauda equine and subject to midline compression [11]. The midline compression to bilateral S1 roots is unlikely to cause a notable interside difference in SN-SSEP.

In the patients with isolated compressive lumbosacral root lesions, the mixed nerve SSEP was almost always normal because of the multi-segment property of the mixed nerves [4]. To evaluate individual root function, techniques such as dermatomal somatosensory evoked potential (DSSEP) were developed to restrict SSEP input in a single nerve root. Although some reported that

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DSSEP was a sensitive measure for lumbosacral radiculopathies [12], other found DSSEP was rarely helpful in diagnosis [13]. The low signal-to-noise ratio rendered the on-line interpretation of DSSEP waveform difficult in the operating theater [14].

SSEPs elicited by cutaneous nerve stimulation are more segmentally specific than those from the mixed nerve stimulation, yet the waveforms are usually robust and consistent [4]. This study found SPN-SSEP was more sensitive in detecting unilateral compressive lesions than PTN-SSEP in anesthetized patients, which was largely in agreement with the earlier findings from non-anesthetized subjects [10,15-18]. In this study all SSEP recordings were performed under identical TIVA anesthesia protocol. Caution should be exercised when using inhalation gases, as the level of cortical suppression was a known factor altering interside interpeak latency difference [19].

The present study found SSEP latency asymmetries corroborated the clinical symptoms, suggesting latency asymmetry might be a good indicator in quantifying conduction delay in the anesthetic state. On the other hand, the incidence of abnormal interside amplitude asymmetries was low; when it did occur, the asymmetry preponderance did not always match the symptom side. Isolated abnormalities of SSEP amplitude should be interpreted with cautions, especially without concomitant latency delay [1].

The lower incidence of interside amplitude asymmetry in lumbosacral patients was presumably a result of central synaptic amplification which had, at least partially, compensated the nerve root axonal loss in the disease process [20]. This is in contrast to the acute laceration of nerve fibers during surgery, which may show an isolated attenuation of SSEP amplitude without concomitant latency delay [21].

Conclusions

This study shows that lower SSEPs are capable of detecting interside asymmetry in anesthetized patients with unilateral radiculopathy. Of the three lower extremity SSEPs, SPN-SSEP appears to be more sensitive than PTN- and SN-SSEP.

Conflict of Interest

No potential conflict of interest relevant to this article was reported.

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