INTRODUCTION

Stereotactic radiosurgery is being rapidly adopted for the treatment of various neurosurgical diseases including brain tumors, vascular malformations, trigeminal neuralgia, and even Parkinson’s disease due to its safety and effectiveness [1-5]. The accurate delivery of ionizing radiation to a specific focus as previously determined by MRI or CT is important for the effective and accurate treatment of intracranial lesions. Therefore, accurate headframe placement is a critically important aspect of gamma knife radiosurgery (GKRS). Headframe is rigidly attached to patients’ skull to establish a stereotactic coordination system and provide a means for precisely positioning the patient in stereotactic space. For example, the Leksell headframe (Elekta AB, Stockholm, Sweden) is fixed to the outer table of skull with two anterior and two posterior screws [6].

Unfortunately, most patients experience considerable dis-
comfort during frame fixation, and this discomfort has been overlooked, because surgeons naturally have focused on successfully conducting the procedure, sustaining stable vital signs, and avoiding problems. Nevertheless, reducing pain during frame fixation would reduce patient discomfort and tend to stabilize vital signs.

Traditionally, local anesthetic solution is injected subcutaneously at anticipated pinning sites using a 23-gauge spinal needle, while the stereotactic frame is held in position by an assistant. However, most patients complain of considerable pain despite of local anesthesia.

This study was designed to quantify and compare levels of pain experienced by patients when stereotactic headframes are placed using the conventional method or modified method designed to reduce pain during headframe fixation.

MATERIALS AND METHODS

We retrospectively studied 66 consecutive patients that underwent GKRS to treat intracranial brain lesions at either of two institutes between April 2015 and December 2015. This study was reviewed and approved by the Institutional Review Board of the Pusan National University Hospital (IRB No: H-1903-009-076). Patients that underwent GKRS due to trigeminal neuralgia and those receiving opioid medication to control pain were excluded. Demographic and baseline variables were collected including age at time of GKRS, gender, diagnosis, degree of pain [as assessed using visual analogue scale (VAS)] during intramuscular (IM) injections based on patients' past experiences. We also excluded patients with a VAS score of 0 during IM injection. For analytic purposes, diagnoses were categorized as brain tumors or vascular malformations. Brain tumors consisted of metastatic brain tumors, meningiomas, schwannomas, lymphomas, and gangliogliomas, and vascular malformations consisted of arteriovenous and cavernous malformations.

The 66 study subjects were divided into two groups according to the method of stereotactic frame fixation used. Group A consisted of 31 patients that received the conventional subcutaneous infiltration technique, and group B consisted of 35 patients that received the modified procedure.

The patients in group A received usual subcutaneous infiltration. Briefly, with the headframe held in position by an assistant, local anesthetic solution (2% lidocaine hydrochloride, 2–3 mL per pinning site) was injected subcutaneously at 4 pinning sites using a 23-gauge spinal needle. The assistant was hard to hold for a long time with the headframe, the 4 pins were applied immediately after local anesthesia according to measurement based on spinal needle depth. No patient received intravenous fentanyl or midazolam prior to headframe placement.

In group B, we performed local anesthesia using modified procedure during stereotactic frame fixation. Briefly, with the headframe held in a standard position by an assistant, each of anticipated pin sites was marked with surgical pen using a needle cap located in bar attached a headframe (Fig. 1). After marking pin sites, the stereotactic frame was removed, and local anesthetic solution (2% lidocaine hydrochloride, 2–3 mL per pinning site) was injected periosteally and subcutaneously at each marked pin site using a 26-gauge sharp-beveled needle (Fig. 2). The stereotactic frame was repositioned by an assistant under the same guidance of needle cap using pin site marks 5 minutes after local infiltration and gentle massage at

Fig. 1. Pin sites were marked with surgical pen under the guidance of needle cap positioned through the pin hole in the frame bar.

Fig. 2. Local anesthetic solution was injected periosteally and subcutaneously at each marked pin site using a 26 gauge sharp-beveled needle.
areas of intended local infiltration. Pins were applied simultaneously to marked sites according to measurement based on spinal needle depth (Fig. 3). No patients received intravenous fentanyl or midazolam prior to frame placement.

To evaluate the severity of pain during the procedures, patients' VAS (scale: 0–10) were assessed twice, that is, during local infiltration and pin insertion. In addition, to minimize group bias, we obtained VAS during IM injection based on patients' recollections. VAS of two groups was compared using the independent t-test. Statistical analysis was performed using SPSS version 18.0 for Windows (SPSS Inc., Chicago, IL, USA). p values of less than 0.05 were considered statistically significant.

RESULTS

Sixty-six patients were enrolled in this study. There were 16 male and 15 female patients in Group A (n=31) and 19 male and 16 female patients in Group B (n=35). For the 66 study subjects, mean average age was 56.4 years old and there were slightly more men (n=35) than women (n=31). The most common pathology treated was metastatic brain tumors (n=39). Age, gender, and diagnosis were non-significantly different in the two groups. Demographic characteristics are shown in Table 1.

The results obtained that mean VAS during headframe placement were significantly lower in group B than in group A (3.61 vs. 7.26; p<0.001). Additionally, mean VAS score was also significantly lower during local injection in Group B (3.74 vs. 4.74; p=0.008). However, mean group VAS scores at IM injection were non-significantly different (p=0.24). These findings are summarized in Table 2.

Table 1. Demographic data of the study subjects

|                         | Total (n=66) | Group A (n=31) | Group B (n=35) | p-value |
|-------------------------|-------------|----------------|----------------|---------|
| Age (yr), mean±SD       | 56.4±13.9   | 56±13.3        | 56.8±14.6      | 0.590   |
| Gender                  |             |                |                |         |
| Male                    | 35          | 16             | 19             | 0.704   |
| Female                  | 31          | 15             | 16             |         |
| Diagnosis               |             |                |                | 0.897   |
| Brain group             |             |                |                |         |
| Metastasis              | 39          | 14             | 25             |         |
| Meningioma              | 9           | 8              | 1              |         |
| Schwannoma              | 5           | 3              | 2              |         |
| Lymphoma                | 1           | 0              | 1              |         |
| Ganglioglioma           | 1           | 0              | 1              |         |
| Vascular group          |             |                |                |         |
| AVM                     | 5           | 3              | 2              |         |
| CM                      | 6           | 3              | 3              |         |

AVM, arteriovenous malformation; CM, cavernous malformation

Table 2. Comparisons of group mean visual analogue scale scores

|                         | Group A       | Group B       | p-value |
|-------------------------|---------------|---------------|---------|
| IM injection            | 2.67±1.24     | 2.31±1.23     | 0.240   |
| During local injection  | 4.74±1.46     | 3.74±1.49     | 0.008   |
| During frame placement  | 7.26±1.95     | 3.61±1.95     | <0.001  |

Results are presented as means±SDs.

DISCUSSION

Nowadays, GKRS is a major therapeutic modality in the neurosurgical field, and an accepted safe, effective procedure. However, many patients experience considerable pain and anxiety during stereotactic frame application, and some complain bitterly. Nevertheless, little attention has been paid to the pain that occurs when a frame is fixed and few studies have addressed the topic.

Liang et al. [7] reported on the effectiveness of topical anesthetic EMLA for postoperative wound pain after GKRS, but Duenas et al. [8] reported that topical EMLA did not significantly reduce pain when used preoperatively for frame fixation. Watson et al. [9] compared the effects of supraorbital and great occipital nerve block and subcutaneous infiltration on pain during stereotactic frame placement, and concluded supraorbital and greater occipital nerve block offered alternatives to subcutaneous infiltration for stereotactic frame placement, although no significant difference was detected between subcutaneous infiltration and nerve block. Wang et al. [6] investigated patient pain, when angled anterior or posterior posts were used to avoid temporalis muscle penetration.
during stereotactic headframe placement, but found no significant difference between conventional anterior posts and angled anterior posts.

The present study was used to objectively assess pain associated with stereotactic frame fixation and compare the pain associated with conventional frame fixation with that experienced using modified frame fixation method. In Group A, while maintaining the stereotactic frame, subcutaneous infiltration was performed at pinning sites using a 23-gauge spinal needle, and then frame fixation was performed promptly. This is because the assistant is hard to keep holding the headframe without changing its location. In Group B, after marking pinning sites with a surgical marking pen under frame guidance, the frame was removed and a local anesthetic solution was injected into the periosteal and subcutaneous layers at marked sites using a 26-gauge sharp-beveled needle. The headframe was repositioned 5 minutes later using the pen marks of patient's head and the 4 pins were applied. Mean VAS at local injection and frame fixation was found to be significantly lower in Group B. VAS was also obtained for IM injections based on patient recollections and no significant intergroup difference was found ($p=0.24$).

We suggest several hypotheses for the lower mean VAS in the group B. First, the onset time of local anesthesia will be important. Lidocaine is known to have an onset about 2 minutes after injection. So the timing of frame fixation and the technique of local anesthesia in group A might have been detrimental to its success for pain reduction during pin placement, relative to group B. Local anesthesia was performed without frame placement into subcutaneous and periosteal spaces in group B. In addition, when our technique is used, it would also be possible to inject several times around marked pinning sites using the pivot technique, and headframe fixation could also be performed after the local anesthetic has taken effect. On the other hand, subcutaneous infiltration was performed while the assistant held the headframe in group A, and pin placement was performed before the anesthetic had taken effect. Furthermore, local anesthetic was injected at a single point during the subcutaneous infiltration technique, which raised a tight bleb, and this is probably painful because scalp tissues are dense and the local increase in tissue pressure is likely to be considerable [9].

Second, the course of the supraorbital nerve and depth of local anesthesia might have affected pain severity. The supraorbital nerve usually exits via the supraorbital foramen in the medio orbital rim and then branches into medioal and lateral branches. The medial branch crosses and runs superficially over frontal muscle to supply the forehead skin and anterior scalp as far as the vertex, whereas the lateral branch lateralizes, traveling on peristeum 1 cm medial and parallel to the tempo-
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helps reduce pain (Fig. 5). Viewed in the course of greater and lesser occipital nerve, difference of pain degree in occiput between two groups is similar with that in forehead.

Third, a 26-gauge sharp-beveled needle was used in group B, which would have reduced pain at time of skin penetration as compared with the 23-gauge needle used in group A.

In conclusion, proper stereotactic frame fixation is of considerable importance for successful GKRS, but pain reduction when fixing the frame also requires consideration. In the present study, we found that our modified method could be helpful for reducing pain and discomfort during the stereotactic procedure.

Conflicts of Interest
The authors have no potential conflicts of interest.

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