Laser interstitial thermal therapy for subependymal giant cell astrocytoma: technical case report

David Y. A. Dadey, PhD,1,2 Ashwin A. Kamath, MD,1 Eric C. Leuthardt, MD,1 and Matthew D. Smyth, MD1

1Department of Neurological Surgery and 2Medical Scientist Training Program, Washington University School of Medicine, St. Louis, Missouri

Subependymal giant cell astrocytoma (SEGA) is a rare tumor occurring almost exclusively in patients with tuberous sclerosis complex. Although open resection remains the standard therapy, complication rates remain high. To minimize morbidity, less invasive approaches, such as endoscope-assisted resection, radiosurgery, and chemotherapy with mTOR pathway inhibitors, are also used to treat these lesions. Laser interstitial thermal therapy (LITT) is a relatively new modality that is increasingly used to treat a variety of intracranial lesions. In this report, the authors describe two pediatric cases of SEGA that were treated with LITT. In both patients the lesion responded well to this treatment modality, with tumor shrinkage observed on follow-up MRI. These cases highlight the potential of LITT to serve as a viable minimally invasive therapeutic approach to the management of SEGAs in the pediatric population.

http://thejns.org/doi/abs/10.3171/2016.7.FOCUS16231

KEY WORDS laser interstitial thermal therapy; LITT; SEGA; subependymal giant cell astrocytoma; ventriculostomy
at 4 years of age, and fenestration of the septum pel- 

lucidum during her first craniotomy to reduce future risk of 

hydrocephalus. Her recovery from these procedures was 

uneventful and she continued to be followed with serial 

imaging over the years. In 2015, a new enlarging mass was 

observed at her prior resection bed in her left lateral ven-

tricle. Her parents also noted new behavioral changes— 

namely, abandoning the family during outings and arrests 

in her ambulation. Her neurological examination was nor-

mal except for left ocular esotropia.

Initial Imaging

MRI showed postsurgical changes consistent with pre-

vious bilateral frontal horn SEGA resections. Multiple T2 
signal hyperintense foci were seen in the cortical and sub-

cortical matter, which were consistent with tubers of TSC. 

A 1.7 × 1.4 × 2.1–cm lobular enhancing mass was present 

in the left lateral ventricle, which had increased in size 

from 6 months previously, when it measured 1.1 × 1 × 1.2 

cm (Fig. 1).

Operation

As the mass was likely to be a recurrent SEGA, her 

family was again counseled and offered various treatment 

options. Conservative management, medical management 

with everolimus (Novartis), a third craniotomy, and the 

option of MRI-guided LITT were discussed. Ultimate-

ly the family elected LITT. On the day of surgery, the 

StealthStation MRI-guided Navigation System (Medtron-

ic) was coregistered with her preoperative MRI scans to 

allow for trajectory planning. A right frontal approach 

with a trajectory crossing through the right lateral ven-

tricle to reach the contralateral left lateral ventricle was 

chosen to allow for probe insertion into the core of the 

tumor. A stab incision was made to allow for drilling of 

a 4.5-mm bur hole and for placement of a 4.5-mm-out-

ner-and 3.2-mm-inner-diameter anchor bolt. Then, a biopsy 

needle was used to cannulate the left ventricle and create 

an entry point on the tumor surface. Using frameless ste-

reotactic guidance, a Monteris Diffusing Tip Laser Probe 

was advanced into the tumor. Once inserted to the de-

sired depth, intraoperative 1.5-T MRI (IMRIS Inc.) was 

used to track thermal ablation with real-time MR ther-

mometry. Using thermal damage threshold (TDT) lines, 

the conformity of the treatment zone was monitored to 

ensure complete coverage of the SEGA. The TDT yel-

low and blue lines are lines that indicate the region of 

tissue heated to the thermal dose equivalent of 43°C for 

2 minutes and 43°C for 10 minutes, respectively (Fig. 2). 

After completion of ablation, intraoperative MRI revealed 

expansion of the tumor consistent with ablation-induced 

swelling. We therefore performed additional ablation to 

expand the treatment zone to encompass the new tumor 

margins. Upon completion of this additional ablation, 

the laser probe and bolt were removed, and the scalp was 

closed with Monocryl 3-0 suture.

Postoperative Course

After surgery, the patient was monitored overnight in 

the pediatric intensive care unit (PICU). She did well and 

was discharged the following day. She was seen for fol-

low-up 3 months later, and her parents indicated she had 

made significant improvement in her behaviors and ex-

hibited no complications since surgery. On neurological 

examination her status was normal except for stable left 

ocular esotropia. MRI performed at the time showed in-

terval decrease in the size of the SEGA, which measured 

1.2 × 0.9 × 1.2 cm. After 9 months of ongoing observation 

since her procedure, she continues to do well.

Case 2

History and Examination

This 14-year-old girl had a history of TSC. She was di-

agnosed at the age of 2 years when she developed infantile 

spasms and refractory epilepsy, and she was found to have 

a spontaneous TSC mutation on genetic evaluation. Her 

seizures have been well controlled since she began taking 

topiramate, and though she does have the classic tubers 

and a SEGA, her disease course overall has been relatively 

benign. On routine neuroimaging, an interval enlargement 

of her SEGA was observed, along with dilation of her lat-

teral ventricles. Despite these findings, she did not indicate 

experiencing any headaches and had a normal neurologi-

cal status on examination.

Initial Imaging

MRI showed multiple subcortical T2 signal hyper-

intense foci consistent with hamartomas of TSC. In ad-

dition, a large enhancing mass was observed in her left 

lateral ventricle near the foramen of Monro, most likely
representing a SEGA. The mass measured 0.7 × 1.0 × 1.1 cm on imaging 6 months previously, and it was found to have grown to a size of 1.6 × 1.7 × 1.9 cm. Interval bilateral enlargement of the lateral ventricles suggestive of obstructive hydrocephalus was also noted (Fig. 3).

Operation

Out of concern for the marked increase in her tumor burden and radiographic evidence of hydrocephalus, we recommended therapeutic intervention. Multiple options including LITT, craniotomy, radiosurgery, and medical management were discussed with the family, but they were most interested in LITT as a definitive, minimally invasive surgical option. Due to the location of the tumor proximal to the foramen of Monro and the possibility of postablation tumor swelling, we planned for the addition of a ventriculostomy during surgery. In addition, we felt she would be a good candidate for treatment with the ROSA robotic device–assisted LITT (Medtech). ROSA is a robotic stereotactic navigation platform with an arm capable of motion with 6 degrees of freedom, allowing for accurate frameless stereotaxy. We secured the patient’s bed frame to the ROSA robot and proceeded to co-register the machine with the patient’s cranial landmarks and preoperative scans. A trajectory using a frontal approach to reach the left lateral ventricle was chosen based on preoperative identification of a pathway avoiding cortical blood vessels. We targeted the exact entry point with ROSA assistance. We made a stab incision at the entry point and drilled a 4-mm bur hole through which a Medtronic biopsy needle was inserted and used to obtain tumor tissue samples. Then, an anchor bolt was placed within the bur hole, after which ROSA was used to determine the distance between the bolt edge and the tumor. This distance was used to determine the insertion depth of the laser probe required to reach the base of the tumor. ROSA was then removed to allow for placement of the laser probe. Intraoperative MRI was used to confirm that the probe was within the lesion and was subsequently used to monitor thermal ablation. LITT was delivered to a treatment zone spanning 90%
ment effects 4 months after surgery.

and axial (B)

FIG. 3. Case 2. Preablation sagittal (A) and axial (C) T1-weighted MR images with Gd contrast showing the SEGA. Postablation sagittal (B) and axial (D) T1-weighted MR images with Gd contrast showing treatment effects 4 months after surgery.

of the tumor, as the eccentricity of the probe placement relative to the lesion did not allow for complete ablation without risk of damage to adjacent normal tissues. Upon completion of the procedure, we removed the probe and bolt and inserted and anchored a ventricular catheter prior to closure of the incision as previously performed.

Postoperative Course

The patient was admitted to the PICU after surgery, where she was monitored overnight. Over subsequent days, she did quite well and the ventriculostomy drain was eventually clamped. However, she developed progressive headaches and experienced two episodes of CSF leakage at the ventriculostomy incision site on the scalp. CT scanning revealed worsened ventriculomegaly. The drain was opened, after which her intracranial pressure normalized and her headaches resolved. Drainage continued for another 72 hours, and on a second clamp trial, repeat CT showed stable ventricular size, and the drain was successfully removed. The patient was discharged 9 days after surgery. Analysis of her biopsy samples confirmed the diagnosis of SEGA. The patient was discharged 9 days after surgery. Analysis of her biopsy samples confirmed the diagnosis of SEGA.

At 4-month follow-up, she had remained symptom free and the ventriculostomy drain was even-

where she was monitored overnight. Over subsequent days, the SEGA and overall stable ventricular size with slightly increased left ventriculomegaly.

Discussion

SEGAs are slow-growing tumors that tend to develop within the first 2 decades of life. Thus, the current management of these tumors involves frequent surveillance for developing hydrocephalus and changes in tumor growth rate. Surgery remains the standard therapy and is often considered curative if gross-total resection is achieved. However, the outcomes of surgery can vary, as one study found that excision of SEGAs greater than 3 cm in diameter was associated with 67% risk for surgery-related complications. In that study, adverse events after surgery were found to correlate with tumor burden and patient age, with patients younger than 3 years having greater risk. Furthermore, incomplete excision may be associated with tumor recurrence, necessitating repeat resection. In an effort to avoid the drawbacks of radical surgery, medical therapies for SEGAs have been a subject of significant investigation. Specifically, mTOR inhibitors have been shown to have efficacy in clinical trials, where tumor volume reductions of greater than 50% were observed in patients treated with everolimus. Though these results are promising, additional investigation is required to determine if long-term therapy is required to prevent SEGA regrowth. Another noninvasive strategy is Gamma Knife radiosurgery, but there is limited evidence to support its efficacy as a treatment for SEGA. To our knowledge, there are very few studies investigating LITT as a treatment for SEGAs. One review article mentions an unpublished series that included SEGAs among a variety of pediatric brain lesions treated with LITT. However, the outcomes were not specifically described. We therefore, herein, illustrate two cases representing our experience with LITT as a therapy for SEGAs. In both cases, LITT was tolerated well by the patients, and there were no intraoperative complications. In both cases, we observed tumor shrinkage on follow-up neuroimaging, suggesting that LITT may have therapeutic efficacy in treatment of SEGAs. Both families indicated improved symptomatology after laser ablation, and for the patient in Case 1 the family also reported reduced seizure frequency and improved overall quality of life. Seizure frequency in the patient in Case 2 did not change. This is likely due to the multifactorial nature of TSC and differences in the underlying pathologic basis of their seizures. Patient 1, at 9 months after laser ablation, continues to remain free of any postoperative complications. Patient 2, at 4 months since her procedure, is also doing well despite a somewhat prolonged postoperative hospital course due to transient obstructive hydrocephalus. We postulate that this progression was due to postablation tumor swelling and that the resulting obstruction was more pronounced because the SEGA was closer to the foramen of Monro.

Transient peritumoral edema is a known treatment effect of LITT in various tumor types. In cases of SEGAs, anticipating the downstream effects of this edema may be of higher importance given that a large proportion of these tumors are close to the foramen of Monro. Furthermore, patients may already present with hydrocephalus at the time of surgery, and it is possible that LITT may lead to transient worsening of their CSF flow obstruction. Our experience with Case 2 suggests that preemptive ventriculostomy could be beneficial when performing ablation of SEGAs. By providing a safety valve in case of aggravated hydrocephalus, ventriculostomy preserves the option of...
immediate pressure relief without necessitating repeat surgical intervention in the immediate postoperative period. Additionally, ventriculostomy is well suited to the transient nature of postablation tumor edema, as it allows for gradual weaning and may obviate the need for a more permanent shunt. The patient in Case 1, due to her prior septostomy and known patency of the foramen of Monro contralateral to her SEGA, was deemed to have a low risk of post-LITT hydrocephalus. Ventriculostomy was therefore not performed in her case.

This report is limited by its retrospective design and limited number of cases and, as such, is not amenable to meaningful statistical analysis. Also, the use of two different methods for stereotactic guidance may confound the overall efficacy observed, as the initial placement of the laser probe can impact the extent of treatment delivery. Lastly, longer follow-up is needed to comment on the long-term effect of LITT on SEGA growth.

Conclusions

The development of minimally invasive techniques for treating SEGAs is needed to avoid morbidity associated with open resection. At our institution, we have found that LITT can be used to treat SEGAs. In both patients described in this report, tumor ablation was achieved without intraoperative complication and resulted in improved symptomatology. Given the proximity of SEGAs to the foramen of Monro and the possibility of transient postablation tumor edema, we devised a method in which ventriculostomy was performed after the LITT procedure. This allowed for effective management of transiently worsened hydrocephalus during the postoperative period (Case 2). Though larger studies are needed with longer follow-up to evaluate overall treatment efficacy, our experience suggests that ventriculostomy should be considered in patients receiving LITT as a treatment for SEGA.

References


Disclosures

Dr. Leuthardt reports having a consulting relationship with Monteris Medical and ownership in Neurolutions, OsteoVantage, and Pear Therapeutics.

Author Contributions

Conception and design: Smyth. Acquisition of data: Kamath, Leuthardt, Smyth. Analysis and interpretation of data: Dadey, Kamath, Smyth. Drafting the article: Dadey. Critical revision: all authors. Approval of the final version of the manuscript: all authors. Approved the final version of the manuscript on behalf of all authors: Dadey. Administrative/technical/material support: Leuthardt, Smyth. Study supervision: Smyth.

Correspondence

David Y. A. Dadey, Department of Neurological Surgery, Washington University in St. Louis, 660 S Euclid Ave., Box 8057, St. Louis, MO 63110. email: ddadey@wustl.edu.