TOPIC REVIEW

Intraoperative MRI for Brain Tumors

Cara Marie Rogers1 · Pamela S. Jones2 · Jefrey S. Weinberg[3](http://orcid.org/0000-0001-5587-6673)

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Abstract

Introduction The use of intraoperative imaging has been a critical tool in the neurosurgeon's armamentarium and is of particular beneft during tumor surgery. This article summarizes the history of its development, implementation, clinical experience and future directions.

Methods We reviewed the literature focusing on the development and clinical experience with intraoperative MRI. Utilizing the authors' personal experience as well as evidence from the literature, we present an overview of the utility of MRI during neurosurgery.

Results In the 1990s, the frst description of using a low feld MRI in the operating room was published describing the additional beneft provided by improved resolution of MRI as compared to ultrasound. Since then, implementation has varied in magnetic feld strength and in confguration from foor mounted to ceiling mounted units as well as those that are accessible to the operating room for use during surgery and via an outpatient entrance to use for diagnostic imaging. The experience shows utility of this technique for increasing extent of resection for low and high grade tumors as well as preventing injury to important structures while incorporating techniques such as intraoperative monitoring.

Conclusion This article reviews the history of intraoperative MRI and presents a review of the literature revealing the successful implementation of this technology and benefts noted for the patient and the surgeon.

Keywords Surgery · Brain tumor · MRI · Glioma · Glioblastoma · Stereotaxy · Stereotactic biopsy · Pituitary tumor · Pediatric

Introduction

The emergence and innovation of intraoperative magnetic resonance imaging (iMRI) over the last several decades has been of particular beneft to the practice of neurosurgical oncology. While a variety of surgical guidance and imaging modalities exist, iMRI provides the highest quality evaluation of surgical execution and assessment of the dynamic changes that occur during surgery in near real time. Understanding the clinical benefts of iMRI as well as its potential limitations is important in the consideration of its utilization.

- ² Department of Neurosurgery, Massachusetts General Hospital, Harvard Medical School, Boston, MA, USA
- ³ Department of Neurosurgery, University of Texas M.D. Anderson Cancer Center, Houston, TX, USA

The evolution of iMRI systems

Intraoperative MRI was frst introduced to the neurosurgical community in the mid 1990s when General Electric and Brigham and Women's Hospital (BWH) in Boston, Massachusetts developed the Signa 0.5 Tesla Magnetic Resonance Therapy (MRT) Unit $[1-3]$ $[1-3]$. The vision for the MRT was fueled by a desire for an open-configuration MRI that would allow surgery to be performed in the scanner with continual image guidance. All MRI systems at the time were closed-confguration units that did not allow access to the patient while in the scanner. Fundamental alterations in the engineering and physics of the magnet and coils had to be made in order for this open-confguration model to come to fruition. The "double donut" magnet system was ultimately chosen in which two superconducting magnets and coils were placed in separate but communicating vertically oriented units. A vertical cleft between the coils allowed the surgeon access to the patient while acquiring concurrent images [\[1](#page-9-0), [2,](#page-9-2) [4–](#page-9-3)[6\]](#page-9-4).

 \boxtimes Jeffrey S. Weinberg jweinberg@mdanderson.org

¹ Department of Neurosurgery, Virginia Tech Carilion, Roanoke, VA, USA

The open-confguration concept was designed to provide surgical access to the patient within the scanner and allow for real-time imaging during the operation; that is to be able to perform surgery while the scan was being completed and with the patient in the same position, thereby reducing the inefficiency of moving the patient multiple times especially when many scans are needed. Maintaining the patient in a static position is thought to improve the accuracy of registration by reducing the risk of patient movement from the surgical position to the scanning position. However the spatial access to the patient in the open-confguration system is signifcantly restricted. The MRT unit provided a relatively small, 56 cm wide area of access, which created unique challenges with the fxed location of the space and this narrow corridor for the surgeon to stand. Another downside to this concept is that all surgical instruments are required to be MR compatible (non-ferromagnetic) to prevent the attraction of the instruments to the magnet and to also limit radiofrequency noise, which would result in imaging artifact. MR compatible instrumentation adds an additional cost to the implementation of this technology [[1,](#page-9-0) [5,](#page-9-5) [7–](#page-9-6)[9\]](#page-9-7).

The next iteration of iMRI technology came in the form of magnets with ultra low-feld strengths of 0.12 T and 0.15 T. This technological advancement was desired because the smaller magnet allowed for reduced size and increased portability. Medtronic introduced PoleStar in 2000 with the N-10 iMRI containing a 0.12 T magnet and later the N-15 iMRI with a 0.15 T magnet $[10]$. Benefits of open-configuration ultra low-feld systems are their compact structure, mobility, and ease of use [[4\]](#page-9-3). Medical centers can build a magnet shielding cabinet (MSC) in a conventional operating room to house the mobile unit, which is then taken out, draped, and positioned over or around the patient when an image is needed [[10\]](#page-9-8). The investment required for the early iMRI systems was prohibitive for many hospital systems, but the ultra low-feld system required much less extensive operative suite remodeling and minimal alterations to the operative equipment and instruments, allowing for a lower implementation cost. The most signifcant downside of the lower feld open-confguration systems, is that they produce a lower quality image when compared to the high feld diagnostic MRI scanners, thereby potentially limiting their operative beneft [\[2](#page-9-2)[–4](#page-9-3), [8](#page-9-9), [9](#page-9-7), [11](#page-9-10), [12\]](#page-9-11).

The limitations posed by low-feld, open-confguration units and the desire to have higher quality imaging led to the conversion to a closed-confguration system, high-feld strength MRI starting in the early 2000's. High-feld iMRI models are available in 1.5 T and 3 T strengths and provide image resolution and defnition that are equivalent to diagnostic MRI scanners and have the ability to provide advanced imaging protocols including difusion weighted, perfusion, spectroscopy, and difusion tensor imaging. Since these units are closed-confguration, there is no access to the patient during scanning, and they require a unique operating room setup in which the operative feld is separate from the scanner. These can be stationary systems in the same or nearby room to the operative suite into which the patient is transported, or ceiling mounted systems that are stored in a shielded alcove and then brought into the operating room on a rail system and over to the operative table [[13\]](#page-9-12). Siemens Sonata is a 1.5 T MRI that was frst introduced into the operating room 2002 and utilized a rotating table with a base that swiveled the table from an operative position into the bore of the magnet [\[14\]](#page-9-13). The IMRIS system which launched in 2005, is a 1.5 T magnet designed on a ceiling-mounted rail system that can move the scanner to the patient. The rail system can be tailored to allow the scanner to be stationed in a separate room for use as a diagnostic scanner, and also move between operating rooms for use in multiple concurrent surgeries $[10, 15]$ $[10, 15]$ $[10, 15]$ $[10, 15]$. Since the operative field is outside of the magnetic feld, this allows the team to use regular instruments and equipment as well as full access to the patient while in the operative position. These high-feld systems can be arranged solely as intraoperative scanners that are designed with operative workflow and efficiency in mind, or for dual functionality for surgery and diagnostics to optimize utilization and lessen the economic burden of the investment [\[1,](#page-9-0) [2,](#page-9-2) [4](#page-9-3), [9,](#page-9-7) [13](#page-9-12), [16,](#page-9-15) [17](#page-9-16)].

Throughout this evolution of iMRI technology, the consensus within the neurosurgical community became to prioritize high quality imaging with optimal spatial and temporal resolution. Therefore, many of the technological advancements in iMRI in recent years have focused on higher feld strength with advancement in image acquisition technology $[18]$ $[18]$ $[18]$. A focus has also been on customization, so that an iMRI suite can be confgured to ft the unique needs of each hospital. A system should be chosen and a suite designed to maximize utility, efficiency, and incorporate all the additional modalities desired by the surgeons. Incorporating the armamentarium of tools available in a conventional operating room was important as well. Suites can be equipped with microscopes, endoscopes, MRI-compatible stereotactic systems, robotic systems, laser technologies, and even hybrid suites with MR and X-ray capabilities have allowed integration of angiography. While the technology was created originally for cranial oncologic purposes, the technology is now being utilized in vascular, functional, and spinal procedures which makes it signifcantly more versatile and potentially cost effective for a hospital system [\[5](#page-9-5), [10](#page-9-8), [18\]](#page-9-17).

Benefts

Accounting for the brain shift phenomenon

Intraoperative MRI provides near real-time information about the dynamic changes that occur in surgery known as the brain shift phenomenon $[1, 19]$ $[1, 19]$ $[1, 19]$ $[1, 19]$. The anatomic position of structures shift within the cranial vault due to a myriad of forces. Administration of anesthetic medications, osmotic agents, changes in overall fuid volume status, and hyperventilation can alter the physiological properties of the tissue. By opening the skull and dura, the brain tissue is exposed to atmospheric pressure resulting in a loss of the forces that act on a closed system leading to gravitational shift. Size of the craniotomy, physical manipulation of the tissue, egress of cerebrospinal fuid from opening of the dura or entry into a ventricle, and removal of a space-occupying lesion can also impact the anatomic configuration of structures [[20,](#page-9-19) [21](#page-9-20)]. This cumulative distortion is nonlinear and can be more profound in patients with hydrocephalus, parenchymal atrophy, edema, or lesions producing signifcant mass efect. Gering et al. investigated the extent and direction of brain shift and found that it is a continuous dynamic process that impacts specific regions of brain tissue differently [\[22](#page-9-21)]. Nimsky and colleagues showed that shift varies greatly throughout the cranial vault with up to 24 mm of shift at the cortical surface and beyond 3 mm at the deep tumor border [\[21\]](#page-9-20). The accuracy of stereotactic navigation that utilizes preoperative imaging can therefore decrease precipitously due to these anatomic shifts during surgery. Stereotactic navigation simply becomes an inefectual tool when the surgeon recognizes the loss in accuracy due to the aforementioned factors, yet can become a dangerous instrument that misleads a surgeon if the inaccuracy is overlooked. Shahar et al. found that in tumors adjacent to the corticospinal tract, there was a mean intraoperative shift in the tumor to corticospinal tract distance of 3.18 mm [[23](#page-9-22)]. If the surgeon does not appreciate this shift, eloquent tissue can easily be violated leading to devastating consequences. Once the accuracy of navigation is efectually lost, updating the image via an intraoperative scan, allows the surgeon to restore their orientation and understanding of the spatial anatomy. Multiple studies have concluded that the only way to provide accurate intraoperative guidance is to obtain serial intraoperative imaging, which is provided in greatest detail by MRI $[21-25]$ $[21-25]$.

Patient position has also been shown to impact brain position [[26\]](#page-9-24). In a typical operating room, stereotactic navigation is registered to the patient using a preoperative diagnostic image, that was obtained with the patient in a supine position. Surgical position, dictated by the tumor location, may not be supine, thereby negatively impacting the accuracy of navigation. Therefore, utilizing iMRI may enhance the accuracy of navigation as the patient is registered to a scan performed in the operative position.

Extent of resection and survival

The success of oncologic surgeries is increasingly being defned by the extent of resection (EOR) with a goal of gross total resection (GTR) or maximal safe cytoreduction. Lacroix et al. found that the only modifable factor that impacted survival in glioblastoma patients, was the EOR. This beneft was found when 89% or more of the enhancing disease was resected, with the most signifcant survival advantage of over 4 months when 93% or more of the enhancing tumor was removed [\[27](#page-9-25)]. Sanai et al. showed a significant survival advantage even at subtotal EORs as low as of 78% in newly diagnosed glioblastomas [\[28](#page-9-26)]. Oppenlander et al. found that in recurrent glioblastoma, an 80% EOR was found to provide an improvement in overall survival (OS) [[29\]](#page-9-27). Scherer et al. reported that in low-grade gliomas, GTR was associated with significantly longer progression free survivals (PFS) and OS [[30](#page-9-28)]. Roelz et al. also showed a signifcant survival advantage in difuse low grade gliomas, with 5 year OS of 82% with initial maximal safe resection versus only 54% when biopsied [\[31](#page-9-29)]. Ultimately, maximal safe resection has been shown by numerous studies to have a signifcant prognostic beneft by improving OS and has become the foundation for a multidisciplinary approach to low- and high-grade brain tumors [\[32](#page-9-30)[–35](#page-9-31)].

The unparalleled ability of MRI to defne pathological tissue has proved superior to surgeon assessment and other imaging modalities. Hatiboglu et al. found that with gliomas, even when the surgeon feels that the goals of surgery have been achieved, iMRI demonstrated unexpected residual tumor resulting in additional resection in 47% of cases [\[36](#page-9-32)]. Golub et al. showed in a meta-analysis published in 2020, that iMRI was superior to conventional navigation in achieving GTR. Roder et al. found in 117 glioblastoma patients reviewed retrospectively, that iMRI was superior to both conventional and 5-ALA guided surgery in achieving smaller residual volumes and more frequent GTRs. Coburger et al. found that the highest sensitivity for detection of residual tumor in low grade gliomas was with iMRI when compared to linear array ultrasound and conventional ultrasound [\[37](#page-9-33)]. MRI is therefore considered the gold standard when determining the EOR $[8, 33]$ $[8, 33]$ $[8, 33]$. Utilization of iMRI has the benefit of optimizing the EOR in an oncologic surgery by providing the surgeon with a more accurate assessment of the tissue at the surgical margins than direct visualization, tactile feedback, or even ultrasound. This allows for intraoperative evaluation as to the EOR and aids the surgeon in removal of any safely accessible residual pathology. Surgeons are able to navigate with the newly obtained intraoperative images, guiding them to the residual area of interest [\[5,](#page-9-5) [36,](#page-9-32) [38,](#page-9-35) [39\]](#page-9-36).

Utilization of iMRI has been shown in numerous studies to correlate with greater EOR and improved patient PFS and even OS. Bohinski et al. and Knauth et al. both showed that iMRI identifed residual resectable tumor in 53% of cases thereby leading to greater EOR in glioma patients [\[17](#page-9-16), [40](#page-9-37)]. In a randomized, prospective study of iMRI as compared to conventional surgery, Senft et al. found that, among the

58 patients enrolled, more patients in the iMRI group had complete tumor resection (23/24 [96%]) than in the control group (17/25 [68%], $p = 0.023$) [\[41](#page-9-38)]. Senft et al. also demonstrated in 103 glioma patients that iMRI led to further resection in 30.1% of cases and that OS was superior in those with complete resections [[42](#page-9-39)]. Zhang et al. reported that patients with nonfunctioning pituitary adenomas undergoing endoscopic transphenoidal resection, use of iMRI to confrm GTR resulted in extended PFS [\[43](#page-9-40)]. In Golub's meta analysis, the improved rates of GTR seen with iMRI correlated to prolonged PFS and OS in high-grade gliomas [\[44](#page-9-41)]. This beneft can be applied to a wide range of pathologies including surgically curable tumors as well as infltrative tumors where optimal cytoreduction may provide a meaningful survival advantage or reduction in the need for toxic adjuvant therapies [[8,](#page-9-9) [16,](#page-9-15) [38,](#page-9-35) [45\]](#page-10-0).

Identifying surgical complications

The use of iMRI can potentially mitigate surgical complications because it allows for visualization beyond the surgically exposed surface. Hemorrhage deep to the resection cavity can be identifed and evacuated prior to closure [\[46](#page-10-1)]. Areas of tissue ischemia can be visualized on an intraoperative scan and medical interventions potentially initiated to prevent further infarct [\[47](#page-10-2)]. Development of hydrocephalus could be recognized and treated with external ventricular drainage if appropriate. Theoretically, complications can be identifed intraoperatively allowing for earlier interventions and thereby reducing neurological compromise and the need for potential reoperations [[16,](#page-9-15) [48,](#page-10-3) [49\]](#page-10-4).

Clinical applications (Literature summary presented in Table [1](#page-4-0))

Low grade gliomas

The utility of iMRI in the resection of low grade gliomas is arguably the most evident. As mentioned previously, gross total or maximal safe resection of low grade gliomas is associated with signifcantly longer PFS and OS. This beneft is so profound in low grade gliomas, that reoperation to remove unintended residual is oftentimes warranted [[30](#page-9-28), [31](#page-9-29), [50](#page-10-5)]. Utilization of iMRI allows the surgeon to ensure that the goals of surgery have been accomplished prior to closing, thereby eliminating the potential need for a reoperation for residual disease. Yahanda and colleagues reported that they performed further resection in almost half (49.7%) of their low grade gliomas cases due to fndings discovered on iMRI, which led to a GTR in 64% of these cases [\[50](#page-10-5)].

Resection of low grade gliomas with the use of iMRI is also extremely valuable because these tumors oftentimes have an appearance and texture similar to normal brain tissue. This characteristic can make defning the border between normal and abnormal tissue more challenging. In low grade gliomas located in ineloquent tissue, a generous resection beyond the borders typically defned by the FLAIR sequences may be a viable option to ensure maximal cytoreduction [\[51](#page-10-6)]. However, achieving GTR in tumors near eloquent regions can become difficult unless there are obvious anatomic borders such as sulci or fssures to guide the surgeon. If these borders do not exist, then a surgeon in a conventional operating room may have to choose between a more conservative approach with the potential of leaving residual tumor behind versus an aggressive approach with the potential of causing neurological injury. The use of iMRI guidance allows the surgeon to err on the side of caution, reimage when the stereotactic navigation becomes unreliable secondary to shift, utilize neurophysiologic monitoring, and identify remaining pathological tissue that may not be obvious on gross examination [\[9](#page-9-7), [33,](#page-9-34) [52\]](#page-10-7).

High grade gliomas

Intraoperative MRI can be an invaluable tool in high grade gliomas located within or adjacent to eloquent regions. The gross appearance of high grade gliomas is typically distinguishable from the surrounding parenchyma and defning the plane between normal and abnormal simpler than with low grade gliomas. However, manipulation of this plane could be fraught with danger when dealing with eloquent structures. Traction on eloquent tissue or deviating out of the plane into eloquent brain could lead to devastating neurological consequences. Any prognostic beneft that an aggressive resection offers is negated by a postoperative neurological deficit. In addition to reduced quality of life and functional status, Rahman et al. found that survival was reduced by over 5 months whether the deficit was transient or persistent [[53](#page-10-8)]. Therefore, a fne balance must be targeted between maximal resection of disease and preservation of neurological function [[35,](#page-9-31) [40,](#page-9-37) [54–](#page-10-9)[56](#page-10-10)].

Pediatric tumors

The benefts of iMRI in the resection of pediatric tumors are similar to those in low grade gliomas since complete resection when feasible is imperative in this population. Maximal safe resection with ideally GTR is the standard of care in children because it leads to signifcant improvements in PFS and OS. In pediatric low-grade gliomas, 10 year PFS is only 50% with subtotal resection versus 85% with GTR [\[57](#page-10-11)]. Multiple studies have demonstrated the utility of intraoperative MRI in aiding to achieve an increase in the EOR of pediatric brain tumors [\[58](#page-10-12), [59](#page-10-13)]. Giordano and colleagues found that iMRI led to the identifcation and removal of

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residual tumor in 27% of glioma and 18% of craniopharyngioma cases resulting in GTR that would otherwise have been subtotal. They also reported that iMRI prompted further resection of unintended residual in 21–52% of cases improving their reoperation rate of 7–14% without iMRI to 0% [[16](#page-9-15), [38](#page-9-35), [60](#page-10-22)]. Shah et al. showed that utilization of iMRI potentially reduces the need for early reoperation in pediatric lesions with 7.7% of their conventional resections requiring reoperation in the frst 2 weeks versus none when iMRI was used [\[61\]](#page-10-16).

A practical beneft of iMRI in the pediatric population is that the volumetric scan needed for registration of navigation, can be performed at the beginning of the case. This eliminates the need for an additional scan prior to surgery that may be distressing for children and may require general anesthesia, particularly in the infant population.

Stereotactic biopsy

Stereotactic biopsies are employed to obtain tissue pathology and the diagnostic yield is reported in multiple large series to be between 90 and 95%. This leaves 5–10% of patients without the necessary diagnosis to guide their treatment [\[62](#page-10-23)[–65](#page-10-24)]. Lesions that are smaller, deeper, lack enhancement, or have extensive necrosis or large cystic components have the lowest diagnostic yields. Failures to obtain pathological tissue can be secondary to faulty targeting or the result of brain shift [[62](#page-10-23)]. The use of iMRI allows the surgeon to verify placement of the cannula and adjust intraoperatively if necessary to ensure the lesion will be sampled appropriately [[25\]](#page-9-23). Studies utilizing iMRI to obtain stereotactic brain biopsies have reported diagnostic yields over 97% [[66](#page-10-20), [67](#page-10-25)]. While iMRI may not be essential to all biopsies, it can be particularly advantageous in the biopsy of small deep lesions that pose the greatest risk of nondiagnostic yield with traditional stereotactic procedures.

Pituitary and skull base tumors

A growing body of evidence suggests that certain patients with skull base tumors may beneft from the guidance of an intraoperative MRI scan. It can be useful in identifying remnants of tumor that may be difficult to visualize in narrow operative corridors unique to these surgeries. Tumors frequently distort the normal anatomy and then the shift of structures due to debulking may further disorient the surgeon [[8](#page-9-9)]. An intraoperative scan may aid the surgeon in determining whether they have accomplished their goals of surgery in terms of debulking of the mass or decompression of critical structures. Although it may not be advantageous for all skull base surgeries, one group found that up to 15.7% of their operations benefted from the guidance of an iMRI by improving EOR or avoiding gratuitous exploration [[68\]](#page-10-26). Multiple studies have established the efficacy of iMRI in transphenoidal resection of pituitary adenomas through identifcation and removal of residual tumor thereby increasing the rate of GTR, and, in some cases, showing a trend towards improved PFS with these increased resection rates [[69–](#page-10-19)[72\]](#page-10-18).

Laser interstitial thermal therapy

Laser interstitial thermal therapy (LITT) is an emerging technique that can be utilized in the treatment of a multitude of brain lesions. LITT is a form of thermal ablation that involves the stereotactic-guided placement of an MRIcompatible optical fber that delivers focused light energy into a lesion to cause thermal injury and immediate necrosis with near real time MR thermometry guidance. The procedure requires a laser, an MRI, and a linked workstation that produces thermal maps that allow the surgeon to monitor the pattern of thermal injury. There are currently two commercial platforms available in the United States including Visualase from Medtronic and NeuroBlate from Monteris Medical. The main diferences between these systems are the wavelength of their laser, cooling mechanism, and patterns of heat production and distribution. The NeuroBlate system utilizes a 1064 nm diode laser that emits light in the form of optical pulses, is cooled with CO2, and has both a difusing and side-fring tip. The Visualase system utilizes a 980 nm diode laser that emits light in the form of a continuous wave, is cooled with saline, and offers a diffusing tip.

LITT can be used in a spectrum of cranial oncologic conditions including both benign and malignant tumors, primary and metastatic tumors, recurrent tumors, and treatment efect secondary to radiation. It is a minimally invasive procedure requiring shorter hospitalizations and recovery times than open surgery. It is an excellent alternative for surgically inaccessible lesions and patients who are not ideal medical candidates for more invasive procedures [\[73](#page-10-21)[–75\]](#page-10-27). LITT is also increasingly being used in the treatment of spinal oncologic tumors and in epilepsy but discussion of these uses is not in the scope of this paper.

The convenience and efficiency of iMRI technology is truly apparent when performing LITT procedures. Utilization of an iMRI streamlines the LITT process substantially by allowing the catheter to be placed with conventional stereotactic guidance in the same room as the iMRI. However, hospital systems that do not have an iMRI suite can use a diagnostic scanner for LITT. They can create an MRIcompatible, sterile operating feld within their diagnostic MRI room and place the catheter using the ClearPoint system which utilizes a percutaneous SmartFrame mount [[76,](#page-10-28) [77](#page-10-29)]. Alternatively, the catheter can be placed in a traditional operating room followed by transporting the patient to a diagnostic MRI for ablation. However, there are several potential drawbacks of this setup. There is the theoretical risk to sterility and catheter stability with transporting a patient from an operating room to a diagnostic scanner. This setup is also less ideal if the catheter is found to be malpositioned or if multiple trajectories are desired as this may require multiple trips back to the operating room [[78,](#page-10-30) [79](#page-10-31)].

Special considerations

Surgical adjuncts

Within the intraoperative MRI suite, surgeons are able to combine all surgical adjuncts utilized in a conventional operating room. Hatiboglu et al. showed that the use electrocorticography in the iMRI suite with cortical and subcortical mapping is easily performed without adverse efects or difficulty $[80]$ $[80]$. Multiple studies have demonstrated that combining awake craniotomies for mapping of eloquent regions with iMRI guidance is a viable and safe option and aids in greater EOR and prognosis. High feld iMRI systems also allow for the integration of functional imaging such as difusion tensor images (DTI) to ensure that eloquent cortical and subcortical structures are respected throughout the procedure, protecting them from unintended damage [[56,](#page-10-10) [81,](#page-10-15) [82](#page-10-33)].

The use of 5-aminolevulinic acid (5-ALA) in the resection of high-grade gliomas for visualization of enhancing malignant tissue is gaining popularity and can be used as an adjunct to iMRI as long as the microscope has the appropriate flter [[23](#page-9-22), [32\]](#page-9-30). Given that 5-ALA is a more recently developed surgical adjunct, the available literature has not been able to elucidate a clear advantage of using iMRI and 5-ALA together. In a prospective study published by Coburger et al. in 2015, they found that the EOR in 33 glioblastoma patients was signifcantly maximized when 5-ALA was used in combination with iMRI compared to 144 patients who under-went resection with iMRI alone [[83](#page-10-14)]. However, a systematic review conducted by Coburger et al. in 2019, concluded that neither iMRI or 5-ALA is superior but a combination could potentially assist in a supra-maximal resection [\[84](#page-10-34)]. A meta-analysis recently conducted by Golub et al. concluded that iMRI and 5-ALA are both individually advantageous in achieving maximal resection and survival beneft, however, they report that the current evidence is uncertain whether they provide an additive beneft [[44\]](#page-9-41).

Operative time

The use of iMRI guidance has been shown to signifcantly increase operative time in comparison to traditional operating rooms. There is additional time needed for setup, registration, draping and redraping of the patient, transport of the patient into and out of the scanner, and intraoperative scanning times. This additional operative time must be taken into account in patient selection and in surgeon and operating room staff scheduling. While multiple studies have concluded that iMRI guided surgery invariably takes more time, no studies have shown an increase in adverse events (e.g. wound infections) [[33,](#page-9-34) [81,](#page-10-15) [85–](#page-10-35)[87\]](#page-11-0).

Image quality

Image quality can be negatively afected by artifact from a variety of factors including motion, metallic susceptibility, radiofrequency noise from inadequately shielded electronics in the operating room, the brain-air interface, and blood products [[49,](#page-10-4) [88](#page-11-1), [89\]](#page-11-2). These factors need to be reduced or eradicated when possible to ensure optimal image defnition. There are also limitations to the utility of repeated contrast enhanced T1 weighted images due to alterations in the blood brain barrier caused by surgical manipulation. Tissue not originally enhancing may demonstrate contrast enhancement once the blood brain barrier is violated or manipulated during surgery [[90\]](#page-11-3). These factors need to be accounted for when interpreting the images obtained during surgery. In our opinion, intraoperative consultation with a neuroradiologist is invaluable.

Safety considerations

The magnetic feld of a MRI is always on and must be taken into consideration in every aspect of the setup and delivery of care. The patient, as well as all personnel, must be screened and cleared for entrance into the iMRI suite. Any individual with metallic implants (e.g. pacemakers, defibrillators, cochlear implants, shunts, pumps, stimulators, shrapnel, ect), may be unsafe to enter. The magnetic feld can alter the functionality of a medical device, pull the device within the feld, or cause the implant to become hot. Safety of the patient and staff is of paramount importance so extensive training with regularly scheduled recertifcation courses should be completed by staff members to ensure an understanding and adherence to safety precautions [\[91](#page-11-4)[–93](#page-11-5)].

All equipment that will enter into the magnetic feld must be MR compatible including carts, monitors, poles, probes, lines, catheters, head fxation devices, and body warmers. Meticulous surgical counts must be preformed every time the patient is transported between the operative position and the scanner to ensure that MRI incompatible instruments are not inadvertently introduced into the magnetic feld [[86,](#page-10-36) [93](#page-11-5)]. Alternatively, with an added investment, MRI compatible instruments can be used to avoid this concern.

Given that the patient will be within the MRI scanner during a portion of their procedure, the anesthesia team must take this distance into consideration and use extended ventilator circuits and access tubing. The magnet can generate heat within electrical wires or cables and burn the patient and therefore cannot be in direct contact with skin [[86](#page-10-36)].

Economic considerations

The economic efficiency of iMRI has to be taken into consideration for the healthcare system as a whole and for a hospital system contemplating its purchase. A major hindrance to universal implementation is the high cost of purchasing a system, ranging from \$3 to \$7 million, not including the cost of renovating the operative suite [[94\]](#page-11-6). Further expenses to purchase shields and MRI compatible equipment and instruments are required. There are also continual and considerable costs required for maintenance of the system and to employ highly trained specialized personnel to keep the room running [\[95](#page-11-7)]. Multiple studies have also shown that operative time is signifcantly increased in iMRI guided cases and this results in additional costs [\[33\]](#page-9-34). The use of iMRI for tumor resection also increases operative costs due to the additional Current Procedural Terminology (CPT) code added to the standard craniotomy for tumor resection code for iMRI interpretation, a potential fnancial incentive for the hospital system but additional burden to the healthcare system.

On the other hand, there are also a multitude of potential economic advantages associated with iMRI for a hospital and for the healthcare system. A study by the University of Minnesota found that in comparison to frst time adult brain tumor resections performed in conventional operating rooms, iMRI guided surgeries resulted in 54.9% shorter lengths of stay, 12.2% lower hospital charges, and 14.4% lower total hospital costs. They also reported that while conventional surgery resulted in immediate repeat resections for residual tumor in 20% of adults and 30% of pediatric cases, there were no repeat resections required after iMRI guided surgery. The mean time to repeat resection was 11.3 months in adults and 18 months in children for iMRI guided surgery in comparison to 9.3 months in adults and 13.3 months in children with conventional surgery $[55]$ $[55]$ $[55]$. The ability to offer iMRI technology to patients has enormous marketing potential for a hospital system, especially those that want to be able to tout comprehensive and innovative oncologic treatments. Hospitals can also offset the cost by constructing a suite that can be used for diagnostic purposes when not being used for an operative case. Therefore, while there are many upfront and continuing costs to the implementation of iMRI, there may also be substantial clinical and fnancial benefts [\[55](#page-10-37)].

Future directions

For a technology that was frst conceptualized and created over 30 years ago, it has advanced remarkably since then and gained global traction. The Health Policy Institute

reported in 2016 that 12% of hospitals have access to iMRI technology in the United States [[96](#page-11-8)]. There is also a growing body of literature investigating the utility of iMRI in the treatment of brain tumors, with over 2600 results in a PubMed search [[97](#page-11-9)]. However, the current available data is not strong enough to recommend that iMRI be included into the standard of care for brain tumors. As Jenkinson and colleagues reported in their review of the literature published in 2018, the evidence to support the use of iMRI in the removal of brain tumors is considered low and very-low quality. This is predominantly because the majority of studies utilize highly selective inclusion criteria favoring younger age and higher functional status, are single-institution studies, and oftentimes conducted at specialized centers [[98](#page-11-10)]. At this time, due to the exorbitant cost associated with investment and maintenance of iMRI technology, and the lack of strong evidence to suggest its superiority over less costly intraoperative imaging and guidance modalities, universal implementation is not feasible. In order to determine whether iMRI will be considered a luxury modality versus a necessity for brain tumor removal in the future, will need to be determined by randomized controlled multi-institutional studies. Ideally, through rigorous research that focuses on neurological and functional outcomes as well as survival, we will be able to elucidate which, if any, pathologies are best approached with iMRI guided resections. As we have watched the technology advance and expand over time, the utility of iMRI is being appreciated for pathologies other than brain tumors and by clinicians other than neurosurgeons. As iMRI technology evolves, the focus should be on optimizing image quality, integration of potentially valuable surgical or intraoperative imaging adjuncts, efficiency of operative setup and workfow, fexibility to accommodate the scope of potential operative positions, efective utilization, and cost-efectiveness.

Conclusion

Intraoperative MRI is a valuable tool in a neurosurgeon's armamentarium in the treatment of brain tumors. The technology is advancing at a rapid pace and providing surgeons with a plethora of innovative techniques targeted toward greater accuracy and improved efficiency. Surgeons must consider the potential benefts of iMRI guidance in each surgical case and weigh them against the additional time and costs associated with its use. Ultimately, iMRI has been shown to be of great beneft in a variety of pathologies and patient populations in helping surgeons to achieve their surgical goals of maximal resection while minimizing morbidity.

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