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Diagnostic and prognostic role of magnetic resonance imaging in cases of moderate to severe traumatic brain injury

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ABSTRACT

Moderate to severe traumatic brain injury (TBI) remains a leading cause of death and disability worldwide. Timely diagnosis and accurate prognostication play a key role in informed clinical decision-making. Though magnetic resonance imaging (MRI) is a superior anatomical scan compared to computerized tomography (CT), the latter remains the current investigation of choice in the clinical setting of TBI due to some of the former's inherent deficiencies in imaging bone/blood, limited access, cost, etc. Nevertheless, the fact that MRI is a valuable adjunct in evaluating the TBI patients with clinical findings disproportionate to the CT scan substantiates its possible complementary/supplementary diagnostic and prognostic role in TBI. MRI scan is ideally placed on demonstrating the shear/diffuse axonal injury (DAI), non-haemorrhagic intraparenchymal lesions, and brain stem lesions poorly delineated by a CT scan. The currently available literature demonstrates that DAI and caudal brainstem lesions are indicators of poorer outcomes. However, the prognostic value of MRI, in addition to that of CT, remains an area of active investigation. We have tried to present the evidence-based use of MRI in moderate to severe TBI. Advances in newer MRI sequences like susceptibility-weighted imaging (SWI), diffusion tensor imaging (DTI), functional MRI (fMRI), and magnetic encephalography (MEG) have the potential to revolutionize the current role of MRI in TBI.

INTRODUCTION

Though computerized tomography (CT) scan of the head remains the investigation of choice for evaluation of a victim of acute traumatic

Keywords

diffusion axonal injury,
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non-haemorrhagic,
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brain injury (TBI), magnetic resonance imaging (MRI) can play a valuable complementary diagnostic/prognostic role in the same [1]. MRI is a superior anatomical scan to CT in all clinical settings, except in the acute moderate to severe TBI, due to its inherent deficiencies in imaging bone/blood. With ever-increasing accessibility seen to MRI scanners worldwide, there is an accompanying increase in the interest, among clinicians and radiologists alike, regarding the utility of MRI in diagnosing and prognosticating a variety of traumatic pathological conditions [2].

The role of MRI scanning in identifying and characterizing specific sequelae of TBI is well documented [3]. Particularly in evaluating patients whose clinical findings are disproportionate to the CT findings, MRI represents a valuable adjunct even in the acute phase. In such cases, MRI demonstrates the shear/diffuse axonal injury (DAI), non-hemorrhagic intraparenchymal lesions, and brain stem lesions poorly delineated by a CT scan. In subacute or chronic head injury, MRI is superior to CT and should be the primary imaging technique whenever possible [3]. Despite its unparalleled sensitiveness to deep intraparenchymal TBI lesions and shear injuries, the broad clinical adaptation of MRI in the trauma setting is limited, partly due to its higher costs, finite access, and more prolonged time consumption.

Early prognostication of moderate to severe TBI facilitates appropriate counseling of patients/families, aiding in sound clinical decisions. These informed clinical decisions are essential, as >50% of such patients end up with a permanent disability. Widely used CT prognostication scoring systems like Rotterdam/Marshall/Stockholm predict mortality better than TBIs' morbidity/permanent disability [1,4]. The 'centripetal model' of TBI, based on animal and postmortem studies, was proposed to better understand the role of shearing injury in clinical outcomes, and add to the existing CT prognostication [5]. It stated that the severity of grading of TBI should be directly proportional to the most caudal cerebral structure affected by the shearing injury. MRI is an ideal imaging modality to achieve the same. Nevertheless, the prognostic value of MRI, in addition to that of CT, remains an area of active investigation. We have tried to present the evidence-based use of MRI in moderate to severe TBI.

PROTOCOL OF MRI FOR TBI

The National Institute of Neurological Disorders and Stroke (NINDS) and several co-sponsoring Federal agencies have proposed common data elements (CDE) for developing data standards for clinical research. The following table shows the CDE for tier 1 evaluation of TBI using 1.5T MRI. Other advanced protocols are available at the NINDS website (Table 1).

DIAGNOSTIC ROLE OF MRI IN TBI

MRI appearance of traumatic lesions

Though CT is highly sensitive for detecting large intracranial hematomas, Gradient echo (GRE) and T2-weighted MRI are equally sensitive. T2 FLAIR (Fluid Attenuated Inversion Recovery) is more sensitive than CT to detect minor cerebral contusions. T2 FLAIR and GRE are as sensitive as CT for traumatic subarachnoid haemorrhage (SAH). T2 FLAIR and DWI (Diffusion-weighted Imaging) are more sensitive for non-hemorrhagic DAI, and GRE and SWI (Susceptibility weighted Imaging) for hemorrhagic DAI. Susceptibility weighted imaging (SWI) is a high-resolution three-dimensional imaging sequence. It is roughly six times more sensitive than the T2-weighted sequence for detecting traumatic microhemorrhages [6]. DWI is the sequence of choice for ischemia. It is essential to differentiate between hemorrhagic and non-hemorrhagic lesions as the hemorrhagic lesions are associated with a worse prognosis. The GRE and T2-weighted MRI sequences also detect signal dropout caused by iron-containing heme groups in slow-moving blood. Distributions of the microhemorrhages in areas associated with axonal injury such as the corpus callosum, brainstem, and other white matter tracts strongly suggest an imaging diagnosis of DAI. DAI microhemorrhages typically appear as punctate signal-free lesions in the white matter that 'bloom'.

Consequently, signal loss caused by punctate haemorrhages from DAI can be visualized years after injury though lesions fade over time. The density of T2-weighted lesions is associated with the severity of TBI in terms of admission Glasgow Coma Scale (GCS) and maximum intracranial pressure (ICP) during admission 3-month Glasgow Outcome Score. Another factor that needs consideration is the magnetic field strength used, as a 3 Tesla MRI demonstrates practically twice the sensitivity of a 1.5 Tesla device for microhemorrhages.

Table 1. Common data element (CDE) protocol for imaging parameters for MRI in TBI¹.

| Sequence | 3D T1W (MPRAGE, 3D IRFSPGR, 3D FFE) | T1W SE (opt if no 3D T1W) | T2W FSE | T2W FLAIR | DWI EPI | 3D SWI† | 2D GRE (FFE) (opt if no 3D SWI) |
|--|-------------------------------------|---------------------------|------------|------------|--------------|------------|---------------------------------|
| Orient | Sagittal | Sagittal | Axial* | Axial* | Axial* | Axial* | Axial* |
| TR (ms) | 9-30 | 500-600 | >3500 | >9500 | >5000 | 50 | >500 |
| TE(ms) | 3-5 | 15-20 | >90 | >110 | >100 | 40 | >20 |
| TI (ms) | 1000 | | | >2000 | | | |
| FA (degrees) | 8-10 | 90 | 90 | 150 | 90 | 15 | 15-20 |
| Freq FOV mm (Phase FOV) | 256(100%) | 230(87.5%) | 230(87.5%) | 230(87.5%) | 256(100%) | 230(87.5%) | 230(87.5%) |
| Matrix size | 256x256 | 256x192 | 256x192 | 256x192 | 128x128 | 512x192 | 256x192 |
| # Slices/ Thickness (mm) | 120/2 | 32/4 | 32/4 | 32/4 | 32/4 | 94/2 | 32/4 |
| Gap | 0 | 0-20% | 0-20% | 0-20% | 0-20% | 0 | 0-20% |
| Voxel size(mm) | 1x1x2 | 1x1x4 | 1x1x4 | 1x1x4 | 2x2x4 | 0.5x1x2 | 1x1x4 |
| NEX | 1 | 2 | 1-2 | 2 | 1-3 | 1 | 1-2 |
| Phase Enc. Dir | A to P | A to P | R to L | R to L | A to P | R to L | R to L |
| Fat suppress | no | no | no | yes | yes | no | no |
| ≅BW(Hz/Px) | 160 | 120 | 130 | 200 | ≥1200 | 80 | 80-100 |
| Flow Comp | no | slice | no | no | no | slice | slice |
| ≅ETL | | | 15-20 | 30 | 128 | | |
| b-values (sec/mm ²) (Directions) | | | | | 0/1000** (3) | | |
| ≅Time*** | 8:00 | 3:00 | 3:00 | 3:00 | 2:00 | 8:00 | 3:00 |

¹ Source-<https://www.commondataelements.ninds.nih.gov/Traumatic%20Brain%20Injury#pane-162>

*Recommend angle to AC-PC line; ** Use 800 sec/mm² for infants <1 year old; ***To reduce acquisition time, use parallel imaging, if possible, AF/CL of 2/24-32; † Can use SWAN on GE systems.

Abbreviations: IR-FSPGR - Inversion recovery, fast spoiled gradient recalled echo; MPRAGE - Magnetization prepared-rapid gradient echo; FFE - Fast field echo; FSE - Fast spin echo; SE- Spin echo; FLAIR - Fluid attenuated inversion recovery; EPI - echo-planar imaging; SWI- Susceptibility-weighted imaging; GRE- Gradient echo; TE - echo delay time; TR - repetition time; TI- Inversion time; FA - fractional anisotropy; FOV - field of view; NEX - number of acquisitions; BW - bandwidth; ETL- Echo train length; A to P - Anterior to posterior; R to L - Right to left.

Classification of Severity of DAI

Grading of DAI has been described histologically according to the anatomic distribution of injury, which correlated with the clinical outcome. The classification was proposed first by Adams in 1989 and divided DAI into three grades [5]. This is the commonly applied grading system for classifying DAI lesions seen on MRI.

- Grade I: Involves grey-white matter interfaces (Figure 1).

Most commonly: Parasagittal regions of frontal lobes, periventricular temporal lobes, Parietal and occipital lobes, internal and external capsules, and cerebellum.

- Grade II: Involves corpus callosum in addition to grade I locations (Figure 2).

Most commonly: Posterior body and splenium but advance anteriorly with increasing injury severity. Most frequently seen unilateral. It may be seen on the SWI sequence.

- Grade III: Involves brainstem in addition to grade I and II locations (Figure 3).

Most commonly: Rostral midbrain, superior cerebellar peduncles, medial lemnisci, and corticospinal tracts.

Grading of brainstem lesions was proposed by Firsching *et al.* and has been validated by others [7-10]

- Grade I: No brainstem involvement; lesions of only hemispheres.
- Grade II: Unilateral brainstem lesions at any level with or without supratentorial lesions.
- Grade III: Bilateral lesions of the mesencephalon with or without supratentorial lesions.
- Grade IV: Bilateral lesions of the pons with or without lesions of lesser grades.

Brainstem injury can be categorized as anterior or posterior, hemorrhagic or non-hemorrhagic, and unilateral or bilateral [11]. Other alternative classification of brainstem injury is primary or secondary due to herniation. The primary injury can be due to a direct impact against the tentorial free edge or as a part of DAI [12,13]. The brainstem lesions are also classified as superficial or deep [13]. The grading assigned by both the systems is

determined by the most caudal brain lesion present, though their definitions vary.

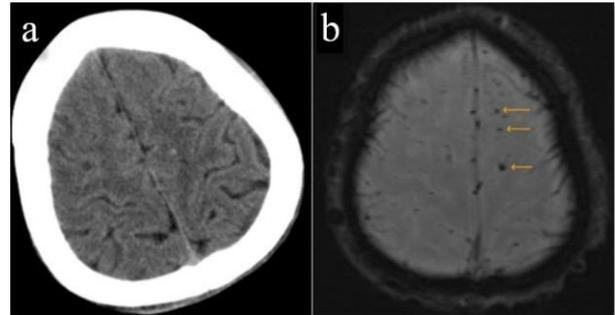


Figure 1. Computerized tomography (CT) scan (a) and magnetic resonance imaging (MRI) - susceptibility-weighted imaging (SWI) sequence (b) showing multiple haemorrhages in left parasagittal grey-white junction (yellow arrows). The presence of such lesions in absence of corpus callosal or brainstem involvement indicates grade 1 diffuse axonal injury (DAI).

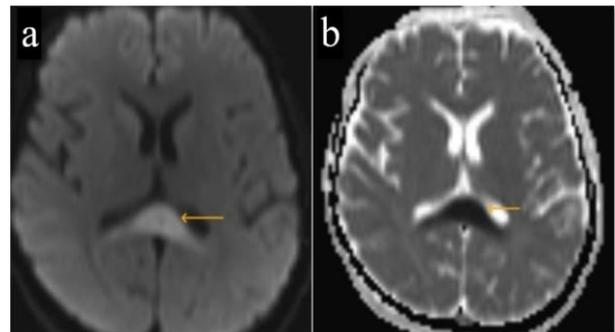


Figure 2. Magnetic resonance imaging (MRI) - diffusion-weighted imaging (DWI) sequence (a) and apparent diffusion coefficient (ADC) sequence (b) showing restricted diffusion (yellow arrows) in the splenium of corpus callosum indicating grade 2 diffuse axonal injury (DAI).

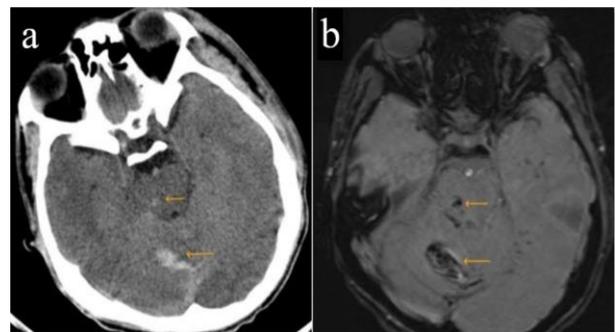


Figure 3. Computerized tomography (CT) scan (a) and magnetic resonance imaging (MRI) - susceptibility-weighted imaging (SWI) sequence (b) showing hemorrhages (yellow arrows) in right paramedian brainstem and cerebellum indicating grade 3 diffuse axonal injury (DAI).

Newer MRI sequences for TBI

Diffusion tensor imaging (DTI) is a newer MRI sequence increasingly used to diagnose DAI [14,15]. DTI is based on the principle of calculating the directional asymmetry of water diffusion, called anisotropy, in mapping white matter tracts. The injured axons in white matter tracts have lower anisotropy than normal ones. Some studies had found it extremely sensitive to axonal injuries following TBI of any severity, even when the CT and conventional MRI sequences showed no abnormalities [15,16]. DTI comes with its fair share of limitations, like its relative insensitivity to axonal injuries in complex white matter regions, due to the absence of one predominant direction of the axons and lack of spatial resolution to detect injuries in small white matter tracts. Other described unique MRI sequences for TBI include magnetic resonance spectroscopy (MRS), functional MRI (fMRI) and magnetic encephalography (MEG). MRS can map changes in the brain metabolic processes and resulting metabolite concentration changes after TBI [17]. fMRI can also reveal areas of either reduced or increased activation and altered connectivity via its BOLD (Blood Oxygenation Level-Dependent) imaging technique [18]. Some authors have reported the usefulness of MEG in detecting TBI induced abnormalities that were not identified with DTI or conventional MRI and correlated with clinical outcome [19].

Prognostic Role of MRI in TBI

There is no consensus about the optimal timing for MRI after TBI for predicting outcome. However, it should preferably be performed within the first-week post-injury for reliable prognostication [20]. The presence of brainstem involvement is correlated with outcome. Mortality increases gradually from 14% with grade I brainstem lesions to 100% with grade IV lesions [10]. Bilateral brainstem involvement is strongly associated with poor outcomes. Posterior location is associated with disability. The non-haemorrhagic, anterior lesions or unilateral injuries are associated with better outcome [11]. Superficial lesions are associated with better outcomes than deeper lesions [13]. SWI lesions in the midbrain corresponding to substantia nigra and tegmentum are independently related to poor outcomes [20].

The grading is also used as a prognostic indicator for recovery of consciousness. Park et al. reported that 14.3% of patients with cerebral white matter lesions (grade 1) did not recover their consciousness, and 50% of patients with corpus callosum lesions (grade 2), 51.6% of patients with brain stem lesion (grade 3) did not recover their consciousness [21]. The grading of DAI is well correlated with time to regain consciousness after TBI. Patients with grade 1 injury become conscious within a week after injury, while grade 2 take two weeks to recover, and patients with grade 3 take two months to regain consciousness [21]. Besides grading, the lesion volume also bears the outcome. The volume of visible DAI lesions in the corpus callosum, brainstem, and thalamus, in DWI and T2 FLAIR sequences, are independent prognostic factors in patients with severe TBI. An essential predictive MRI variable is DWI lesion volume in the corpus callosum. However, in moderate TBI cases, the number of cortical contusions is more critical for prognosis [22].

A systematic review published by Haghbayan et al. (2017), reviewing the prognostic value of MRI in moderate to severe TBI, concluded that the brainstem lesions were associated with higher mortality and unfavourable functional outcome (Glasgow Outcome Scale 1-3) at ≥ 6 months [23]. And that DAI patterns were associated with an increased risk of unfavourable functional outcomes, with depth-based MRI scores demonstrating an increased risk of unfavourable outcomes as more caudal structures were affected. Along the same lines, a recent meta-analysis by M. M. Van eijck et al. (2018), studying the prognostic value of DAI in TBI, reported that the presence of DAI resulted in three times higher risk of an unfavourable outcome, and this risk further increased three times with each increasing grade of DAI [24]. This meta-analysis reviewed data from 32 selected articles, deducing an overall unfavourable (Glasgow outcome score 1-3) functional outcome of 38% in patients with DAI in TBI.

CONCLUSION

MRI following TBI provides valuable assistance, either complementary or supplementary to CT scan, in determining diagnosis and prognosis in individual cases. The presence of DAI and caudal brainstem lesions lead to a poorer outcome. Advances in newer MRI sequences like SWI, DTI, fMRI, MEG. have the

potential to revolutionize the current role of MRI in TBI.

Abbreviations:

ADC - Apparent diffusion coefficient

CT - Computed tomography

DAI - Diffuse axonal injury

DWI - Diffusion-weighted imaging

GCS - Glasgow Coma Scale

GRE - Gradient recalled echo

ICP - Intracranial pressure

MRI - Magnetic resonance imaging

NINDS - National Institute of Neurological Disorders and Stroke

SAH - Subarachnoid hemorrhage

SDH - Subdural hematoma

SWI - Susceptibility-weighted imaging

T2 FLAIR - T2-weighted Fluid Attenuated Inversion Recovery

TBI - Traumatic brain injury

fMRI - Functional magnetic resonance imaging

BOLD - Blood oxygen level-dependent

MRS - Magnetic resonance spectroscopy

DTI - Diffusion tensor imaging

MEG - Magnetic encephalography

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