

## Deep transcranial magnetic stimulation (dTMS) for treatment of major depressive disorder (MDD) status post-surgical removal of medulloblastoma: A case report of safety



Dear Editor:

We are reporting a case with history of surgical removal of medulloblastoma, ventriculogaleal shunt in place, history of mild TBI and intake of bupropion for treatment of MDD. This patient tolerated dTMS treatment using H1-coil without incidence of any complications. (see Fig. 1).

### Background

Major depressive disorder is one of the most common mental disorders in the United States. In 2017, 7.1% of adults aged 18 or older (17.3 million adults) had at least one Major Depressive Episode (MDE) in the past year, and 4.5% of adults (11.0 million adults) had a MDE with severe impairment in the past year [1]. Depression is the leading cause of disability worldwide, and is a major contributor to the overall global burden of disease. In October 2008 transcranial magnetic stimulation (TMS) was approved by the Food and Drug Administration (FDA) for the treatment of major depressive disorder (MDD) [2]. Induction of seizures is one of the most severe acute adverse effects of TMS therapy. The occurrence of seizures as acute adverse effect of TMS therapy has been extremely rare [3]. Personal history of seizure disorder, history of Traumatic Brain Injury (TBI), brain tumor, brain lesions, congenital brain malformations, sleep deprivation, alcoholism, use of drugs that decrease seizure threshold and recent withdrawal of anticonvulsant drugs, are other risk factors for seizures in patients undergoing TMS [4]. This is a case report of safety of dTMS treatment for Major Depressive Disorder (MDD) in a patient with past history of craniotomy for removal of medulloblastoma, ventriculogaleal shunt in place, history of mild TBI and intake of bupropion.

### Case report

A 52-year-old Caucasian male with history of Major Depressive Disorder presented with worsening depression symptoms. He started seeking psychiatric services at age 47 after a Major Depressive Episode (MDE). He endorsed sad mood, anhedonia, hopelessness, hypersomnia, lack of motivation, lack of physical activity, weight gain, loss of libido, lack of functioning. These symptoms got worse in fall and winter months. Mood and energy levels improved during summer but never to the remission. He denied suicidal ideations. Screen for bipolar disorder, psychotic disorders, anxiety disorders, eating disorder was negative. There was no

previous history of psychiatric hospitalizations, self-injurious behaviors, and suicide attempts. Trials of different medications citalopram, fluoxetine, sertraline, duloxetine, bupropion, aripiprazole, risperidone did not result in remission of MDD symptoms. He attended Cognitive Behavioral Therapy (CBT) and supportive psychotherapy sessions without change in depression symptoms. Patient was taking duloxetine 60mg daily and bupropion XL 300mg daily before starting TMS treatment. His mother has history of Major Depressive Disorder. Patient was average student in school. No history of learning disability. He worked as forklift driver, left work at age 47 due to depression. Patient denied recent or remote history of drug or alcohol use. Medical history is positive for paroxysmal atrial fibrillation, HTN, GERD, Esophageal stricture.

At age 48, he had mild TBI with loss of consciousness, multiple rib fractures, pneumothorax, vertebral compression fracture and pelvic fracture after a motor vehicle accident. He recovered from these injuries without residual symptoms. At age 18, sub occipital craniotomy with excision of fourth ventricular tumor was done. Total tumor was approximately 3.5 cm in diameter and appeared to fill the entire 4th ventricle. Histology reports showed tumor was medulloblastoma. During surgery ventriculogaleal shunt was placed. He also finished radiation therapy for tumor. Patient denied any recent complaints related to this surgery.

After critical consideration of risks and benefits of Transcranial Magnetic Stimulation (TMS), treatment procedure was discussed in details with patient and his mother. He was told that risk of seizure during TMS therapy is rare but he was still at high risk due to previous surgery of brain tumor, ventriculogaleal shunt in place, traumatic brain injury and intake of bupropion [5]. Informed consent was obtained for TMS therapy. Medications were not changed before and throughout the TMS therapy, although it is well documented that bupropion can lower seizure threshold [6]. He clinically benefitted from this medication and was not ready to stop this medication. TMS was delivered using the H1-coil (Brainsway Deep TMS) to the left dorsolateral prefrontal cortex according to the FDA-approved protocol [7]: 18-Hz stimulation for 2 second per train with 20 seconds between trains, with 55 trains (1980 pulses) delivered over 20 minutes. The maximum intensity of the stimulation was 120% of Motor Threshold (MT). Initial treatment was delivered daily for 6 weeks and taper was started week 7. Three sessions delivered week 7, 2 sessions week 8 and one session week 9. TMS was discontinued after week 9. Hamilton Rating Scale for Depression (HAM-D) was done one week prior to the first session of TMS treatment and every 2 weeks after start

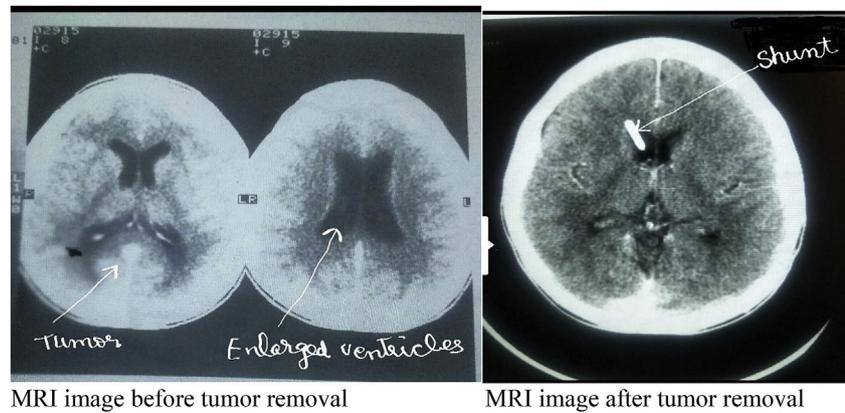


Fig. 1. MRI images before and after tumor removal.

of the treatment. HAM-D score was 22 prior to the first session of TMS and the score decreased from 22 to 5 after 4 weeks of the treatment. HAM-D score was 1 after 8 weeks of the treatment. Patient denied any side effects of TMS therapy. There were no episodes of incidental seizures throughout the treatment.

### Discussion

Carefully weighing the balance of benefits versus potential harm is always important in the practice of medicine. Strictly adhering to the warning about increased seizure risk and depriving patients like this of a treatment that can improve their quality of life so dramatically is unjust. A meta-analysis estimated the risk of a seizure from repetitive TMS in subjects with previous history of epilepsy is 1.4% [8] and no cases of status-epilepticus. The deep TMS induced seizure rate is 0.087% overall and 0.028% per instructions for use rate (3). Therefore, in carefully selected cases, if depression is severe, treatment resistant, if alternative treatments have failed or are unavailable and person's functioning is impaired, benefits of TMS therapy can outweigh the potential harm from an induced seizure. This case report is to provide some reassurance about safety to trying this modality of treatment even in patients with several risk factors for seizures.

### Declaration of interest

None.

### Conflicts of interest

None.

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