Deep brain stimulation for Alcohol Use Disorder

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- Cost-effectiveness analysis of deep brain stimulation for the treatment of alcohol use disorder and alcoholic liver disease
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- Cognitive Dysfunction in the Addictions (CDiA): A Neuron to Neighbourhood Collaborative Research Program on Executive Dysfunction and Functional Outcomes in Outpatients Seeking Treatment for Addiction
- Cost-Effectiveness Analysis of Deep Brain Stimulation for the Treatment of Alcohol Use Disorder and Alcoholic Liver Disease

DBS has been primarily used to treat movement disorders such as Parkinson's disease and essential tremor, it has also been explored as a potential treatment for other neurological and psychiatric conditions, including Alcohol Use Disorder (AUD).

Here are some key points about DBS for AUD:

Research Status: DBS for AUD is an emerging and experimental treatment. Clinical trials and research studies are ongoing to investigate its safety and effectiveness. It is not yet considered a standard or FDA-approved treatment for AUD.

Target Areas: Different research studies have targeted various brain regions with DBS for AUD. The nucleus accumbens, a part of the brain's reward system, has been a common target because of its role in addiction.

Mechanism of Action: The exact mechanism by which DBS may help individuals with AUD is not fully understood. It is believed to modulate neural circuits involved in reward processing and craving, potentially reducing the urge to consume alcohol.

Patient Selection: Patients considered for DBS for AUD are typically those with severe and treatmentresistant alcohol addiction who have not responded to other therapies. Candidates undergo thorough psychiatric and medical evaluations to determine their eligibility.

Procedure: The DBS procedure involves the surgical implantation of electrodes in the target brain region. The electrodes are connected to a pulse generator (similar to a pacemaker), which is typically implanted under the skin in the chest or abdomen. The pulse generator can be programmed to deliver electrical stimulation to the brain region as needed.

Risks and Side Effects: As with any surgical procedure, DBS carries risks, including infection, bleeding, and device-related complications. Additionally, side effects related to stimulation can occur, such as

mood changes, cognitive changes, and physical symptoms.

Outcome and Efficacy: The effectiveness of DBS for AUD varies among individuals. Some studies have shown promising results in reducing alcohol consumption and cravings, while others have been less successful. Further research is needed to better understand which patients are most likely to benefit from this treatment.

Ethical Considerations: The use of DBS for AUD raises ethical and societal questions, including issues related to patient autonomy, consent, and the potential for altering behavior through brain manipulation.

Cost and Accessibility: DBS is a complex and costly procedure that is not widely available. It is typically reserved for research settings and specialized centers.

It's important to note that DBS for AUD is still in the experimental phase, and its long-term safety and efficacy are not fully established. Individuals with AUD should discuss their treatment options with healthcare professionals, consider evidence-based approaches such as medications and behavioral therapies, and, if appropriate, explore participation in clinical trials or research studies related to DBS for AUD. Treatment decisions should be made on a case-by-case basis, taking into account individual circumstances and preferences.

The current literature suggests that DBS has a moderate effect on SUD symptoms. However, the limited number of studies and small sample size indicate that more research is needed to better understand the factors that influence its effectiveness ¹⁾.

Bach et al. reported a double-blind randomized controlled trial comparing active DBS ("DBS-EARLY ON") against sham stimulation ("DBS-LATE ON") over 6 months in n = 12 AUD inpatients. This 6-month blind phase was followed by a 12-month unblinded period in which all patients received active DBS. Continuous abstinence (primary outcome), alcohol use, alcohol craving, depressiveness, anxiety, anhedonia, and quality of life served as outcome parameters. The primary intention-to-treat analysis, comparing continuous abstinence between treatment groups, did not yield statistically significant results, most likely due to the restricted number of participants. In light of the resulting limited statistical power, there is the question of whether DBS effects on secondary outcomes can nonetheless be interpreted as indicative of a therapeutic effect. Analyses of secondary outcomes provide evidence for this, demonstrating a significantly higher proportion of abstinent days, lower alcohol cravings, and anhedonia in the DBS-EARLY ON group 6 months after randomization. Exploratory responder analyses indicated that patients with high baseline alcohol craving, depressiveness, and anhedonia responded to DBS. The results of this first randomized controlled trial are suggestive of the beneficial effects of DBS in treatment-resistant AUD and encourage replication in larger samples²⁰.

Six patients with severe, refractory AUD underwent NAc-DBS. Safety metrics and clinical outcomes were recorded. Positron emission tomography (FDG-PET) was used to measure glucose metabolism in the NAc at baseline and 6 months. Functional magnetic resonance imaging (fMRI) was used to

characterize postoperative changes in NAc functional connectivity to the rest of the brain, as well as NAc and dorsal striatal reactivity to alcoholic visual cues. This study was registered with ClinicalTrials.gov, NCT03660124. All patients experienced a reduction in craving. There was a significant reduction in alcohol consumption, alcohol-related compulsivity, and anxiety at 12 months. There was no significant change in depression. FDG-PET analysis demonstrated reduced NAc metabolism by 6 months, which correlated with improvements in compulsive drinking behaviors. Clinical improvement correlated with reduced functional connectivity between the NAc and the visual association cortex. Active DBS was associated with reduced activation of the dorsal striatum during passive viewing of alcohol-containing pictures. NAc-DBS is feasible and safe in patients with severe, otherwise refractory AUD. It is associated with a reduction in cravings and addictive behavior. A potential mechanism underlying this process is a down-regulation of the NAc, a disruption of its functional connectivity to the visual association cortex, and interference of cue-elicited dorsal striatum reactivity. Trial Registration NCT03660124 (www.clinicaltrials.gov) ³¹

Test

What is the primary status of DBS as a treatment for Alcohol Use Disorder (AUD)?

- a) It is a standard and FDA-approved treatment for AUD.
- b) It is experimental and emerging, with ongoing research and clinical trials.
- c) It is widely available and accessible.
- d) It is a widely used alternative to medications.

Which brain region has been commonly targeted in DBS for AUD due to its role in addiction?

- a) Prefrontal cortex
- b) Hippocampus
- c) Nucleus accumbens
- d) Cerebellum
- What is the believed mechanism of action of DBS for AUD?
- a) Modulating neural circuits involved in motor control
- b) Modulating neural circuits involved in memory processing
- c) Modulating neural circuits involved in reward processing and craving
- d) Modulating neural circuits involved in visual perception
- What type of patients are typically considered for DBS for AUD?
- a) Individuals with mild alcohol addiction
- b) Individuals who have not responded to other therapies but have not severe addiction

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c) Patients with severe and treatment-resistant alcohol addiction

- d) Patients with a history of addiction to other substances
- What does the DBS procedure involve?
- a) Implanting electrodes in the heart
- b) Implanting electrodes in the eyes
- c) Implanting electrodes in the target brain region connected to a pulse generator
- d) Implanting electrodes in the liver
- What are some potential risks associated with DBS for AUD?
- a) Hair loss and skin discoloration
- b) Mood changes, cognitive changes, and physical symptoms
- c) Improved memory and concentration
- d) Enhanced athletic performance
- How does the effectiveness of DBS for AUD vary among individuals?
- a) It has a consistent and high success rate in all cases.
- b) It is equally effective for all stages of AUD.
- c) It varies among individuals, with some showing promising results and others less successful.
- d) It only works for young individuals.
- What ethical questions are raised by the use of DBS for AUD?
- a) Questions about the patient's age and gender

b) Questions about the potential for altering behavior through brain manipulation, patient autonomy, and consent

- c) Questions about the cost of the procedure
- d) Questions about the potential for addiction to DBS
- What is the primary status of DBS for AUD in terms of cost and accessibility?
- a) It is widely available and affordable.
- b) It is a standard procedure covered by most insurance plans.
- c) It is complex and costly, typically reserved for research settings and specialized centers.
- d) It is free of charge for all patients.

What does FDG-PET measure in the context of DBS for AUD?

- a) Brain structure
- b) Glucose metabolism in the nucleus accumbens
- c) Alcohol content in the bloodstream
- d) Electrical activity in the brain

Answers:

b) It is experimental and emerging, with ongoing research and clinical trials. c) Nucleus accumbens c) Modulating neural circuits involved in reward processing and craving c) Patients with severe and treatment-resistant alcohol addiction c) Implanting electrodes in the target brain region connected to a pulse generator b) Mood changes, cognitive changes, and physical symptoms c) It varies among individuals, with some showing promising results and others less successful. b) Questions about the potential for altering behavior through brain manipulation, patient autonomy, and consent c) It is complex and costly, typically reserved for research settings and specialized centers. b) Glucose metabolism in the nucleus accumbens

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