



## Brain stimulation may reduce food cravings as obesity treatment EMBARGOED UNTIL 00.01 CEST MONDAY 21 MAY 2018

Stimulating the brain to alter its intrinsic reward system shows promise in the treatment of obesity, according to results presented in Barcelona at the <u>European Society of Endocrinology</u> annual meeting, <u>ECE 2018</u>. The technique has yielded positive results after just a single treatment session, revealing its potential to become a safer alternative to treat obesity, avoiding invasive surgery and drug side effects.

Obesity is a global epidemic, with approximately 650 million adults and 340 million children and adolescents currently considered obese, and the disease contributing to an estimated 2.8 million deaths per year worldwide. It has been reported that, in some obesity cases, the reward system in the brain may be altered, causing a greater reward response to food than in normal weight individuals. This can make patients more vulnerable to craving, and can lead to weight gain. This dysfunction in the reward system can also be seen in cases of addiction to substances, e.g. drugs or alcohol, or behaviours, e.g. gambling.

Deep Transcranial Magnetic Stimulation (dTMS) is a medical treatment that uses magnetic energy to stimulate neurons in specific areas of the brain. It is used to treat depression and addictive behaviours, and previous studies have suggested that dTMS could be a good option to reduce drug and food cravings. However, the potential mechanism driving these changes had not been investigated until now.

In this study, Professor Livio Luzi and colleagues, from the Istituto di Ricovero e Cura a Carattere Scientifico Policlinico San Donato, Italy, investigated the effects of dTMS on appetite and satiety in obese people. They studied the effects of a single 30 minute session of dTMS, at high or low frequency, on blood markers potentially associated with food reward in a group of 40 obese patients. They found that high frequency dTMS significantly increased blood levels of beta-endorphins - neurotransmitters involved in producing heightened feelings of reward after food ingestion - compared to low frequency dTMS or controls.

"For the first time, this study is able to suggest an explanation of how dTMS could alter food cravings in obese subjects" says Professor Luzi. "We also found that some blood markers potentially associated with food reward, for example glucose, vary according to gender, suggesting male/female differences in how vulnerable patients are to food cravings, and their ability to lose weight."

Since the current study only measures changes in blood markers, the next steps for the research group include using brain imaging studies to directly identify how high frequency dTMS changes the structure and function of the obese brain, both short and long term, and extending this treatment to a larger population of obese patients.

"Given the distressing effects of obesity in patients, and the socioeconomic burden of the condition, it is increasingly urgent to identify new strategies to counteract the current obesity trends. dTMS could present a much safer and cheaper alternative to treat obesity compared to drugs or surgery", Professor Luzi adds.





## Abstract

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DEEP TRANSCRANIAL MAGNETIC STIMULATION ACUTELY MODULATES NEURO-ENDOCRINE PATHWAYS UNDERLYING OBESITY Anna Ferrulli<sup>1</sup>, Michela Adamo<sup>1</sup>, Stefano Massarini<sup>2</sup>, <u>Livio Luzi<sup>1,2</sup></u> <sup>1</sup>IRCCS Policlinico San Donato, San Donato Milanese (MI), Italy. <sup>2</sup>University of Milan, Milan, Italy.

Deep Transcranical Magnetic Stimulation (dTMS) is a non-invasive modulation technique of cortical excitability that has shown to affect the mesolimbic and mesostriatal dopaminergic pathways. Consistent with these assets, dTMS is widely used as a therapeutic tool in neuro-psychiatric disorders associated with abnormal cortical excitability and dopaminergic activity, as addiction. Hence, considering the dysfuncional role of PFC and dopaminergic reward system in controlling appetite, dTMS was suggested to be affective in reducing food craving and in controlling body weight in obese subjects. However, the underlying physiological mechanisms of the dTMS effects are not fully know. Aim of this study was to investigate the effects of a single dTMS session on neuro-endocrine pathways in obesity.

Fourty obese patients (11 M, 29 M; age: 48.0±1.6; BMI: 36.3±0.7) were assigned to receive one session of high frequency (18 Hz, HF), low frequency (1 Hz, LF) or Sham stimulation via an H-coil dTMS. H-coil was targeted to stimulate Prefrontal Cortex and Insula, bilaterally. Metabolic and neuro-endocrine parameters were evaluated before and after a single dTMS session.

Following the 18 Hz dTMS session, a significant increase of norepinephrine was found ( $5.6\pm0.9 \text{ vs } 6.5\pm1.2 \text{ ng/mL}$ , +18.0 $\pm6.8\%$ , p=0.01; p=0.05 vs LF); a rise in  $\beta$ -endorphins levels was also shown ( $0.338\pm0.049 \text{ vs } 0.372\pm0.048 \text{ ng/mL}$ , +13.9 $\pm4.6\%$ , p=0.017; p=0.010 vs Sham; p=0.011 vs LF). In the same group, glucose levels significantly increased ( $90.2\pm4.1 \text{ vs } 96.2\pm4.2 \text{ mg/dL}$ , +7.0 $\pm1.8\%$ , p=0.002), whilst leptin levels significantly decreased ( $66.9\pm10.5 \text{ vs } 56.3\pm9.0 \text{ ng/mL}$ , -16.3 $\pm3.0\%$ , p=0.002). Furthermore, pituitary hormones significantly decreased after a single 18 Hz dTMS session, specifically TSH ( $2.71\pm0.25 \text{ vs } 2.09\pm0.18 \mu$ U/mL, -20.7 $\pm4.7\%$ , p=0.001) and prolactin ( $17.1\pm1.3 \text{ vs } 10.6\pm0.8 \text{ ng/mL}$ , -34.7 $\pm4.7\%$ , p=0.015).

These results suggest that dTMS can acutely affect or exygenic pathways and metabolic parameters mainly via modulation of the sympathetic activity and hypothalamic-pituitary-adrenal axis. The increase of  $\beta$ -endorphins could suggest a potential role of HF dTMS in inducing the dopaminergic system activation and therefore, in modulating the food reward system. Together these findings support the role of dTMS as a novel promising treatment for obesity.





## **Notes for Editors**

- 1. The study "Deep transcranial magnetic stimulation acutely modulates neuro-endocrine pathways underlying obesity" is an oral communication that will take place on 21 May 2018, at the European Congress of Endocrinology at the Centre Convencions Internacional Barcelona, Spain.
- 2. For other press enquiries please contact the ECE 2018 press office:

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- 3. The European Congress of Endocrinology is held at Centre Convencions Internacional Barcelona, Spain on the 19-22 May 2017. See the <u>full scientific programme</u>.
- 4. The <u>European Society of Endocrinology</u> was created to promote research, education and clinical practice in endocrinology by the organisation of conferences, training courses and publications, by raising public awareness, liaison with national and international legislators, and by any other appropriate means.