

Transcutaneous supraorbital nerve stimulation enhances somatosensory thalamic activity in migraine between attacks: a central mechanism of clinical efficacy?

Davide Di Lenola¹, Gianluca Coppola², Mariano Serrao¹, Cherubino Di Lorenzo³, Francesco Pierelli¹

1. "Sapienza" University of Rome Polo Pontino, Department of medico-surgical sciences and biotechnologies, Latina, Italy
2. G.B. Bietti Foundation IRCCS, Department of Neurophysiology of Vision and Neurophthalmology, Rome, Italy
3. Don Carlo Gnocchi Onlus Foundation, Milan, Italy

BACKGROUND

In a recent randomized double-blind sham-controlled study the Cefaly®, a novel transcutaneous supraorbital electrostimulation device, has been successfully used as a prophylactic treatment for episodic migraine (Schoenen et al., 2013).

The possible mechanisms of action through which the device is able to induce clinical improvement in migraine are not known.

In the present study, we investigated whether Cefaly® may act centrally at the thalamocortical/cortical level.

DESIGN & METHODS

To explore the central mechanisms of action of Cefaly®, we have recorded the somatosensory evoked potentials (SSEPs) before, and twice (intersession interval of 5 minutes) after one session of supraorbital stimulation lasting 20min in 10 migraine without aura patients between attacks.

We measured the N20-P25 amplitudes on the low-frequency-SSEP, and, after applying a band-pass filter (450-750 Hz), maximal peak-to-peak amplitudes of the *pre-synaptic*, reflecting thalamocortical activity, and *post-synaptic*, reflecting primary cortical activation, high-frequency oscillations (HFOs) (Coppola et al., 2005).

RESULTS

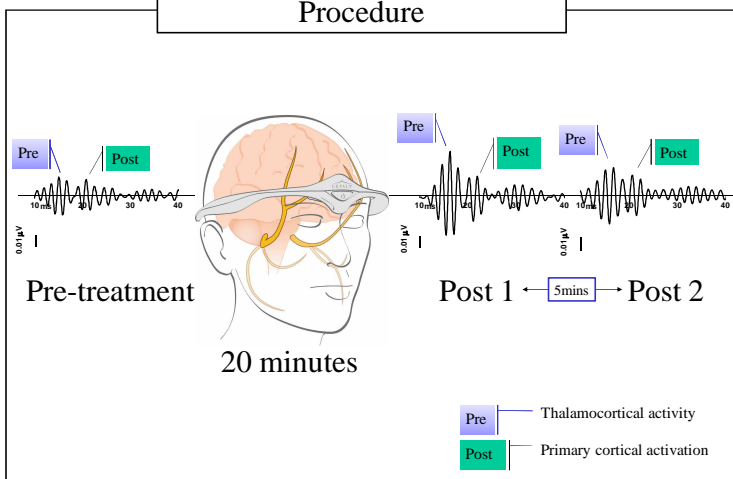
Pre-synaptic HFO amplitudes, reflecting somatosensory thalamocortical activity, significantly increased immediately after the end of the supraorbital stimulation with Cefaly® (from 0.035 μ V to 0.058 μ V, $p < 0.01$), but recovered to the pre-stimulation values as early as 5 minutes later. Both the low-frequency N20-P25 SSEP component and post-synaptic HFOs were unaffected by Cefaly®.

DISCUSSION

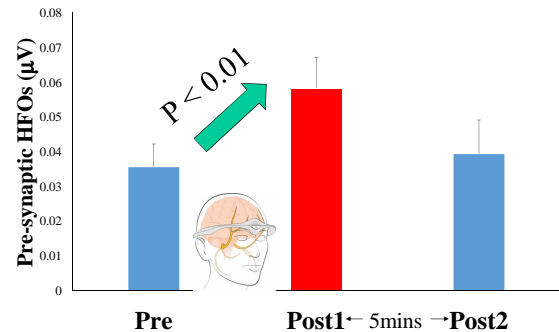
Present data might support the hypothesis that Cefaly® acts centrally through increased thalamocortical activity induced by the neurostimulation.

It is of obvious interest to verify whether these device-induced changes might persist at long-term after 3-month daily preventive stimulation, and if they follow clinical improvement.

Procedure

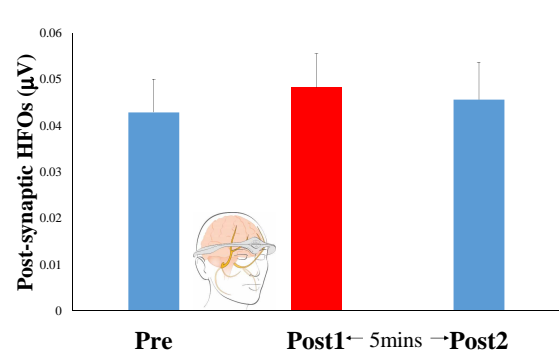


Pre-synaptic HFOs



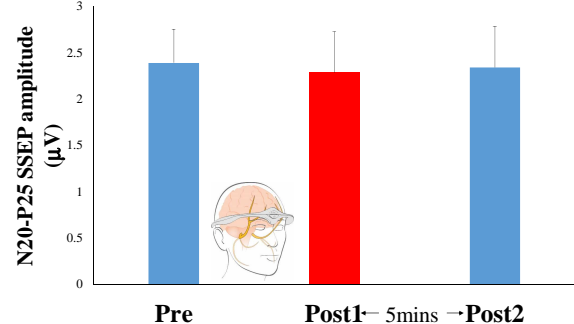
Pre-synaptic HFO amplitudes, reflecting somatosensory *thalamocortical activity*, significantly increased after the supraorbital stimulation with Cefaly®.

Post-synaptic HFOs



Post-synaptic HFOs, reflecting *primary cortical activation*, were unaffected by 20 min supraorbital stimulation with Cefaly®.

Somatosensory evoked potentials (SSEPs)



Low-frequency N20-P25 SSEP amplitude were unaffected by 20 min supraorbital stimulation with Cefaly®.

REFERENCES

- Schoenen J, Vandersmissen B, Jeanette S, Herroelen L, Vandehede M, Gérard P, Magis D. Migraine prevention with a supraorbital transcutaneous stimulator: a randomized controlled trial. *Neurology*. 2013 Feb 19;80(8):697-704.
- Coppola G, Vandehede M, Di Clemente L, Ambrosini A, Fumal A, De Pasqua V, Schoenen J. Somatosensory evoked high-frequency oscillations reflecting thalamo-cortical activity are decreased in migraine patients between attacks. *Brain*. 2005 Jan;128(Pt 1):98-103.