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LETTER TO THE EDITOR

Pupil area and photopigment spectral sensitivity are relevant to study of migraine photophobia

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Sir,

Noseda and colleagues (2016) report findings from a very informative multifaceted study aimed at exploring the effect of different light wavelengths in eliciting photophobia in patients with migraine, and probing the basis for any differential effect. They found that green light exacerbated headache significantly less than white, blue, amber or red lights. They went on to show, using electroretinography and visual evoked potentials in patients, that green light stimuli produce less activation of cone-driven pathways, discernible at the level of retinal responses. Their patients did not undergo pharmacological mydriasis to minimize discomfort, and this may be reasonable in that it more faithfully reproduces natural circumstances. However, pupil diameter affects retinal illuminance (which is calculated as the product of pupil area and luminance at the cornea). Differences in the amplitude, velocity and latency of pupil light responses occur with different wavelengths of light (Lobato-Rincon et al., 2014), and it is possible that the pupil areas, and therefore light reaching the retina differed in the different conditions. It would be helpful to know whether pupil diameters were similar during the different coloured light exposures, in which case the effect would relate directly to stimulation of cone pathways by the different wavelengths. If the pupil diameters differed significantly, then a more complex interaction of effects of light wavelengths on the pupil area (the afferent pathway of the pupil reflex is driven ultimately by photon absorption by rod and cone photopigments as well as melanopsin) and the resulting retinal stimulation would need to be considered. It is also possible that activation of the pupil light reflex itself might contribute to photophobia.

The authors were also able, using multi-unit recordings in rats, to identify thalamic neurons responding differently to different wavelengths (least responsive to green light). This is of interest and relevance, but interspecies differences bear consideration. As photopigment spectral sensitivities differ between human and rat (Jacobs *et al.*, 2001), it is not clear that patterns of relative stimulation by different wavelengths will be identical. Thus some degree of caution is advisable before directly relating findings in rodents to the human visual system.

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