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Letter to Editor

Acute psychosis in elderly: do not forget the CASPR2 spectrum as a possible cause

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Antibodies against contactin-associated protein 2 (CASPR2), a protein associated with the Voltage-Gated Potassium Channel Complex (VGKC) [1], represent an emerging cause of some old neurological manifestations; such as Morvan's and Isaacs' syndromes, as well as a variety of phenotypes encompassing limbic encephalitis, neuropathic pain, late onset epilepsy and dysautonomia [2,3].

Although the prevalence of anti-neuronal surface antibodies and their impact on clinical practice is not well-established [4-7], they represent an important etiology of autoimmune encephalitis and a curable cause of dementia and acute psychosis [8-10].

In a study by *Baumgartner*, *et al.* 34% of the patients with autoimmune encephalitis were initially admitted to the psychiatry department. In addition, psychiatry disturbances represent the second most common presentation of autoimmune encephalitis after seizures [11].

When it comes to anti-CASPR2 psychiatric phenotypes, symptoms tend to present in a progressive manner (5-7 months); mostly with no fluctuations, and a tumor is present in 19% of the patients [2,10]. It affects mostly elderly man (Figure 1) [2,12] and clinical presentation ranges from cognitive decline, sleep disorders, depressive mood, global amnesia – especially when associated with anti-Leucine-rich, glioma inactivated 1 (anti-LGI1) antibodies – and psychotic episodes¹². Other symptoms, such as seizures, peripheral nerve symptoms and dysautonomia may appear later in the disease, possibly being misdiagnosed as dementia or psychiatric disease [2].

It is important to highlight that most of the patients with anti-CASPR2 have either normal MRI image or hippocampal

atrophy, which holds and association with anti-LGI1 antibodies; and therefore the diagnosis may be challenging when facing a case with this suspected etiology [13].

Who is the anti-CASPR2 + psychiatric patient ?

Profile: Male, mean age of onset 66 [2], with subacute cognitive disfunction. Don't present symptoms fluctuations.

Diagnostic test: MRI - normal or with bilateral syperintensity on

symptoms:
Hallucinations, global amnesia, severe deficits in memory, naming and frontal lobe function, sleep disorders, behavioral disorders, psychosis.

Psychiatric

Diagnostic test: MRI normal or with bilateral hyperintensity on medial temporal lobe. EEG - normal, epileptic or slow. R

Figure 1: We summarized the profile of the patient that usually present with the anti-CASPR2 psychiatric phenotypes. Elderly man image acquired at http://smart.servier.com/.

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In conclusion, antibodies against CASPR2 should be remembered as a cause of cognitive imbalance or psychosis, especially in elderly men, even with a normal MRI image. Also, if a tumor is not yet known, it is important to make a screening. Finally, anti-CASPR2 disorders respond well to immunotherapy, but may acquire a poor prognosis, especially when treatment is delayed due a misdiagnosis [12].

Author's contributions

Performed data acquisition and data analysis as well helping with the writing in the text: André Ricardo Merkle, Letícia Caroline Breis, Marco Antônio Machado Schlindwein.

Made substantial contributions to conception and design of the study, as well as provided technical support: Marcus Vinicius Magno Gonçalves MD PhD.

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