

BACKGROUND

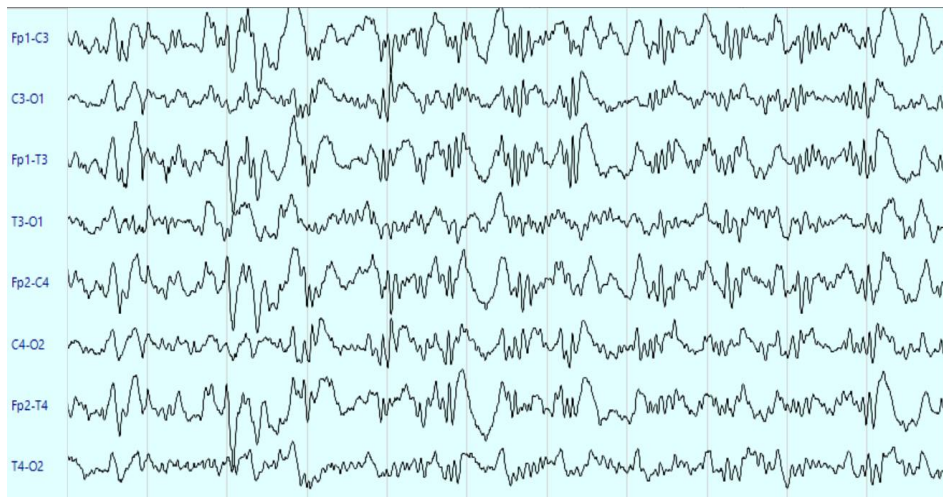
Paraneoplastic encephalitis is an autoimmune brain inflammation driven by onconeural antibodies targeting tumor-related neuronal antigens. We report a case of anti-Ma2 paraneoplastic encephalitis presenting with new-onset seizures and subacute cognitive-behavioural changes, in which initial negative tests contributed to diagnostic complexity as well as the absence of a primary tumor; a multidisciplinary approach was important to guide diagnostic investigations.

MEDICAL HISTORY

A 63-year-old male with a past medical history of hypertension and atrial fibrillation was admitted to the ICU for sudden-onset unresponsiveness and morsus sign. His wife recently reported behavioural changes, stereotyped speech and significant weight loss.

CLINICAL EVALUATION & DIAGNOSTIC TESTS

- Routine laboratory tests, brain imaging (CT, MRI), initial EEG recordings: unremarkable
- Neurological status fluctuated: episodes of **aphasia, stereotyped language, obsessive thoughts** and **confusion** during which EEG recording revealed **non-convulsive status epilepticus (NCSE)**, treated acutely with diazepam and lacosamide, later with chronic valproate and levetiracetam due to bradycardia
- CSF: negative except for **borderline positivity for anti-Ma2 antibodies**, also found in serum.

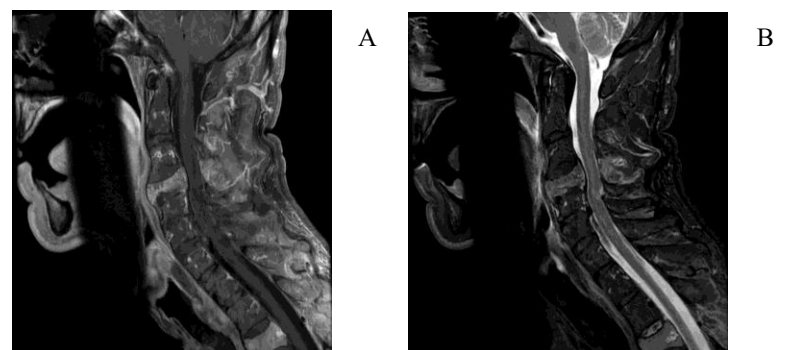


EEG excerpt: bilateral frontotemporal spike-wave and polyspike-wave complexes, with a tendency to posterior propagation, occurring periodically, consistent with non-convulsive status epilepticus (NCSE)

SUBSEQUENT DIAGNOSTIC WORK-UP

The patient was managed by a multidisciplinary team including neurologists, neurosurgeons, oncologists, urologists and radiologists.

- ❖ Total body CT → **bone alterations suggestive for metastases**: sternal manubrium, C4, D3, left femur
- ❖ PET imaging → **significant FDG uptake** in axial bones, prostate, and gastric wall.
- ❖ **Tumor markers** → PSA, AFP, chromogranin A **moderately elevated**
- ❖ Cervicothoracic spine MRI → supported the **suspicion of neoplastic etiology of the vertebral alterations**
- ❖ Bone biopsy at D3 → aspecific
- ❖ EGDS → normal
- ❖ Prostate MRI → signs of prostatitis, no tumor signs
- ❖ Testicular ultrasound → bilateral hydrocele with parenchymal calcifications in the left testis.



T1-weighted post-contrast (A) and STIR (B) sagittal MRI images of cervicothoracic vertebral alterations (C4, D3)

THERAPY & RESULTS

- ✓ Five days of **methylprednisolone** (1 g/day IV) led to **significant** clinical and electroencephalographic **improvement**
- ✓ Clinical presentation, bone alterations and antibody profile supported a **diagnosis of paraneoplastic anti-Ma2 encephalitis, with occult primary tumor, suspicious bone metastases and elevation of tumor markers**
- ✓ The patient was **discharged seizure-free on double antiepileptic therapy**, without language or cognitive-behavioral disorders, awaiting vertebral stabilization with histological diagnosis to define the primary tumor. Diagnostic work-up may also require further periodical radiological and PET checks
- ✓ **Empirical orchietomy** will be **considered** as second diagnostic option in light of the **hypothesis of intratubular germ cell neoplasia (ITGCN)**, which is an established etiology of anti-Ma2 encephalitis, even in the absence of a testicular mass. A multidisciplinary approach is essential for appropriate management.

REFERENCES

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