

## **Ventral subpial and central cord enhancement in spinal neurosarcoidosis: the reverse trident sign.**

Amir M. Jaafari<sup>1\*</sup>, Eric Newman<sup>1\*</sup>, Eoin P. Flanagan<sup>2</sup>, Sebastian Chiriboga-Lopez<sup>1</sup>, Sarosh R. Irani<sup>1</sup>

### Affiliations

1. Departments of Neurology and Neuroscience, Mayo Clinic, Jacksonville, Florida, USA
2. Department of Neurology and Laboratory Medicine and Pathology, Mayo Clinic, Rochester, Minnesota, USA

\*These authors contributed equally

Corresponding author: Professor Sarosh R Irani (Departments of Neurology and Neurosciences, 4500 San Pablo Road, Mangurian Building, 5th Floor, Jacksonville, FL 32224, USA; +19049532000: [irani.sarosh@mayo.edu](mailto:irani.sarosh@mayo.edu))

Type of article: Imaging case-report

Words and Figures: 150 words; one figure and one supplementary table

Keywords: neurosarcoidosis, spinal, trident, infliximab

A 64-year-old female presented with 18-months of progressive scalp-neck dysesthesias, left leg weakness and urinary urgency. MRI revealed a longitudinally extensive transverse T2-hyperintensity, from obex to thoracic spine, with ventral subpial and central contrast enhancement<sup>1</sup> (Fig.1). CSF revealed mild hyperproteinorrachia and oligoclonal bands. Body CT-PET, MRI brain, and malignancy-granulomatous screenings were unremarkable (Suppl.Table.1). Isolated neurosarcoidosis was diagnosed and treatment with corticosteroids, rituximab, and infliximab yielded marked clinical and radiographic improvements over 2.5 years (Fig.1). A recognized radiographic presentation of neurosarcoidosis is the trident sign, characterized by dorsal subpial and central enhancement.<sup>2</sup> In this patient, the pattern was inverted, with ventral prominence, a finding we term the “reverse trident sign”. Both appearances reflect the prominent blood-CNS barrier leakiness, with leptomeningeal predilection, of neurosarcoidosis. Considering the diagnostic challenges of isolated spinal neurosarcoidosis and its responsiveness to immunotherapy, recognizing alternative radiographic presentations is crucial for accurate diagnosis and effective management.

## References

1. Boban J, Thurnher MM. Ventral-subpial enhancement in spinal cord sarcoidosis: A braid-like sign. *Neurology* 2019;92(5):236-38. doi: 10.1212/WNL.0000000000006857 [published Online First: 20190128]
2. Zalewski NL, Krecke KN, Weinshenker BG, et al. Central canal enhancement and the trident sign in spinal cord sarcoidosis. *Neurology* 2016;87(7):743-4. doi: 10.1212/WNL.0000000000002992

**Figure 1. The reverse trident sign in neurosarcoidosis.** Sagittal T2 MRI demonstrating longitudinally extensive hyperintensity (A,C,E,G) with subpial cervical contrast enhancement (sagittal T1; B,D,F,H). Axial post-gadolinium MRI showing ventral subpial and central enhancement (the ‘reverse trident’ sign; yellow outline; M-P). T2 signal and cord swelling improved markedly with time. Drug timings shown.

**Funding.** SRI was supported by the Wellcome [104079/Z/14/Z], the Medical Research Council (MR/V007173/1), BMA Research Grants – Vera Down grant (2013), Margaret Temple (2017), and by the National Institute for Health Research (NIHR) Oxford Biomedical Research Centre. The views

expressed are those of the authors and not necessarily those of the NHS, the NIHR, or the Department of Health.

**Consent.** The patient has provided informed written consent to share anonymized data (Mayo IRB 08-007846-128)

### **Competing interests statement.**

A.M. Jaafari reports no disclosures relevant to the manuscript. E. Newman reports no disclosures relevant to the manuscript. E.P. Flanagan has served on advisory boards for Alexion, Genentech, Horizon Therapeutics and UCB. E.P. Flanagan has received research support from UCB, funding from the NIH (R01NS113828), honoraria for editing and writing articles for The Continuum Lifelong Learning in Neurology Journal which is a publication of the American Academy of Neurology and received royalties from UpToDate. E.P. Flanagan is a site principal investigator in a randomized clinical trial of Rozanolixizumab for relapsing myelin oligodendrocyte glycoprotein antibody-associated disease run by UCB, site principal investigator and a member of the steering committee for a clinical trial of satralizumab for relapsing myelin oligodendrocyte glycoprotein antibody-associated disease run by Roche/Genentech, co-Investigator on a study of ravulizumab for neuromyelitis optica spectrum disorder run by Alexion and a member of the medical advisory board of the MOG project . E.P. Flanagan has given educational talks on neuromyelitis optica spectrum disorder funded by Alexion. E.P. Flanagan is an editorial board member of Neurology, Neuroimmunology and Neuroinflammation, The Journal of the Neurological Sciences and Neuroimmunology Reports. E.P. Flanagan has submitted a patent on DACH1-IgG as a biomarker of paraneoplastic autoimmunity. S. Chiriboga-Lopez reports no disclosures relevant to the manuscript. S.R. Irani has received honoraria/research support from UCB, Immunovant, MedImmune, Roche, Janssen, Cerebral therapeutics, ADC therapeutics, Brain, CSL Behring, and ONO Pharma; licensed royalties on patent application WO/2010/046716 entitled “Neurological Autoimmune Disorders”; and has filed two other patents entitled “Diagnostic method and therapy” (WO2019211633 and US-2021-0071249-A1; PCT application WO202189788A1) and “Biomarkers” (PCT/GB2022/050614 and WO202189788A1).

