Neurons but Motor Neurons in Motor Neuron disease

Francesco Fornai¹, Paola Lenzi¹, Michela Ferrucci¹, Stefano Ruggieri, Loredana D’Este⁴, Antonio Pellegrini¹, Lorenzo Fumagalli⁵, Francesco Giannessi¹ and Antonio Paparelli¹

¹Human Anatomy Department of Translational Research and New Technologies in Medicine and Surgery, University of Pisa, 56126 Pisa, Italy
²Neuronal Sciences, Sapienza University of Rome, Rome, Italy
³I.R.C.C.S. Neuromed, Pozzilli, Italy
⁴Laboratory of Immunohistochemistry Tindaro G. Renda, Department of Anatomic, Histologic, Forensic and Locomotor Apparatus Sciences, Sapienza University of Rome, Rome, Italy
⁵Department of Anatomy, Histology, Forensic Medicine and Orthopedics, Sapienza University of Rome, Rome, Italy

The occurrence of motor neuron death is the milestone of amyotrophic lateral sclerosis (ALS). Therefore, morphological analysis along decades focussed on motor neuron loss as the sole marker to score disease severity.

Recently, non autonomous cell death took a prominent role to explain the need for additional cell types to induce motor neuron degeneration in ALS. This concept encompasses a variety of inflammatory cells including resident astroglia and microglia.

Despite tremendous clinical and basic research activity the role of inflammation remains controversial. On the other hand very recently non autonomous cell death creped back in considering neuronal cell types differing from motor neurons.

In the present communication evidence is provided demonstrating that interneurons and other neuronal cell types participate to degeneration. Remarkably, this multi-neuronal non-autonomous motor neuron degeneration extends to multiple models of motor neuron disorders. Interneurons degenerate earlier and more than motor neurons, thus suggesting a key role in the pathophysiology of motor neuron disease. Additionally, these findings portray ALS as a whole spinal cord disorder rather than a disease affecting solely motor neurons. Novel concepts of abnormal cell to cell communication are likely to underlie multi-neuronal disease spreading.

This work was supported by research grant PRIN 2010-2011 FF

Key words

Non autononomous cell death, motor neuron Ranshow cell, ALS.