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## Gut microbiota alteration in Clinically Isolated Syndrome: a pilot study

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**Abstract:** EP1495

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**Background:** Relatively little is known about what might trigger or facilitate the first episode of demyelization in Multiple Sclerosis (MS); recent data indicate that the dysregulation of the immune system that occurs in MS could be controlled by environmental factors. The composition of the gut microbiota structure in terms of species richness and distribution, as well as the functional potential of the community, can greatly impact the host immune system; an imbalance in the gut microbiome has been shown to induce a profound alteration of immune responses both in the gut-associated tissue and in the periphery and could be a risk factor for MS.

**Aims:** As the Clinically Isolated Syndrome (CIS) allows to study the disease processes closest to the biological onset of MS, the aim of this pilot project was to investigate whether alteration in the composition of the gut microbiota could be associated with CIS and its immune system alteration.

**Methods:** Stool and blood samples were collected from 20 CIS patients and 20 Healthy Volunteers (HV). DNA isolated from stools were subjected to shotgun metagenomic sequencing strategy in order to discover the microbiota composition as well as the microbial function. T helper (Th)17 and T regulatory (Treg) cells were analyzed by FACS in the peripheral blood (PB).

**Results:** Our preliminary results indicate a lower abundance of *Bacteroides* and a decrease species richness in CIS patients versus HV. In the PB, CIS patients displayed an increase of pathogenic Th17 cells expressing Toll Like Receptor 2 and a decrease of Treg cells producing Interleukin-10 and expressing CD39 compared to HV.

**Conclusions:** These findings indicate that gut microbial dysbiosis could exist at the onset of MS and could be suggestive of a pro-inflammatory milieu observed in the periphery. The analysis on metagenomic content and microbial gene identification will allow us to determine the presence/abundance of specific genes that can be correlated with CIS in order to design strategies to modulate the immune system through alteration of gut microbiome.

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