



POSTER PRESENTATION

Open Access

Expression and activation of human endogenous retroviruses of the W family in blood cells and astrocytes: implications for the pathogenesis of multiple sclerosis

Luciana Poddighe¹, Giuseppe Marnelli¹, Alessandra Mei¹, Elena Uleri¹, Caterina Serra¹, Roberto Manetti², Antonina Dolei^{1*}

From *Frontiers of Retrovirology* 2011
Amsterdam, The Netherlands. 3-5 October 2011

Background

Multiple sclerosis (MS) is a chronic inflammatory disease of the central nervous system, with demyelination and gliosis. Proposed pathogenic co-factors triggering MS pathogenesis are the Epstein Barr virus (EBV), and two elements of the W family of human endogenous retroviruses (HERV-W): MSR_V, that forms free virions, and syncytin-1, the ERVWE1env protein; both retroelements have neuropathogenic properties. In the past we studied MSR_V in MS patients in various temporal and clinical stages; in all cases, striking parallels between MS behaviour and MSR_V/HERV-W presence/load were found. By simultaneous detection of MSR_V and HHV-6, we found a direct correlation between MS and MSR_V presence/load, but not for HHV-6. MS brains over-express MSR_Venv and syncytin-1 transcripts, with respect to controls, while EBV presence was not detected.

Materials and methods

Since late EBV seroconversion is a strong risk factor for MS development, we performed *in vitro* experiments on PBMC from MS patients and MSR_V⁺ volunteers, as well as on U87-MG astrogloma cells, that were studied as such or were exposed to EBV or to recombinant EBV glycoprotein350 (EBVgp350), or to proinflamma-

tory cytokines. The levels of MSR_Venv and syncytin-1 mRNAs were evaluated by discriminatory real time RT-PCR assays. Flow cytometry was used to evaluate the HERV-Wenv protein on the plasmamembrane, as well the PBMC subsets.

Results

Basal expression of MSR_Venv and syncytin-1 occurs in astrocytes and in NK, B and monocyte cells, but not in T cells. This uneven expression is amplified in naive MS patients. Astrocyte infection by EBV and exposure to EBVgp350 stimulate the expression of HERV-W/MSR_V/syncytin-1, with requirement of the NF- κ B pathway. In EBVgp350-treated PBMC, MSR_Venv and syncytin-1 are activated in B cells and monocytes, but not in T cells, nor in the highly expressing NK cells. The latter cells, but not the T cells, are activated by proinflammatory cytokines.

Conclusions

The study demonstrates that there are interactions among the above proposed MS-cofactors. *In vivo*, a pathogenic outcome would depend on activation in abnormal situations/tissues, as it may occur in delayed EBV infection, or in the presence of particular host genetic backgrounds, or both.

Author details

¹Department of Biomedical Sciences and Center of Excellence for Biotechnology Development and Biodiversity Research. ²Department of Clinical, Experimental and Oncological Medicine, University of Sassari, 07100 Sassari, Italy.

¹Department of Biomedical Sciences and Center of Excellence for Biotechnology Development and Biodiversity Research
Full list of author information is available at the end of the article

Published: 3 October 2011

doi:10.1186/1742-4690-8-S2-P19

Cite this article as: Poddighe *et al.*: Expression and activation of human endogenous retroviruses of the W family in blood cells and astrocytes: implications for the pathogenesis of multiple sclerosis. *Retrovirology* 2011 **8**(Suppl 2):P19.

**Submit your next manuscript to BioMed Central
and take full advantage of:**

- Convenient online submission
- Thorough peer review
- No space constraints or color figure charges
- Immediate publication on acceptance
- Inclusion in PubMed, CAS, Scopus and Google Scholar
- Research which is freely available for redistribution

Submit your manuscript at
www.biomedcentral.com/submit

