Jiří Hejnar hejnar@img.cas.cz

Laboratory of Viral and Cellular Genetics

Receptors for retroviruses, retroviral vectors, endogenous retroviruses





Jiří Hejnar, PhD / Head of Laboratory Jana Blažková, PhD / Research Scienstist Josef Geryk, PhD / Research Scienstist Jiří Plachý, PhD / Research Scienstist Jan Svoboda, Prof, DSc / Research Scienstist Kateřina Trejbalová, PhD / Research Scienstist Věra Hoserová, MSc / Research Assistant Dana Kučerová, MSc / Research Assistant Markéta Reinišová. MSc / Research Assistant Jitka Dvořáková / Technician Lenka Mikušová / Technician Kamila Thunová / Technician Magda Matoušková, MSc / PhD Student Dana Průková, MSc / PhD Student Volodymyr Stepanets, MSc / PhD Student Filip Šenigl, MSc / PhD Student Miroslav Auxt / Diploma Student Petr Daniel / Diploma Student Jan Kotáb / Diploma Student Denisa Kovářová / Diploma Student Anton A Buzdin / Visiting Scientist



Retroviral vector transduces GFP reporter gene into chicken testicular cells including the spermatogonial stem cells

Research topics

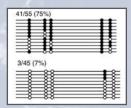
The main scientific interest of our group has been traditionally focused on the interactions of retroviruses with the host cells. Retroviruses enter their natural host cells via specific receptors, integrate into the host genome, and use the cell transcription machinery to express their structural or enzymatically active proteins. At the level of whole organism, retroviruses induce various pathologies and may even penetrate the germ lines being thereafter transmitted vertically as endogenous retroviruses. Human endogenous retroviruses do not replicate themselves but some of them are inevitable for our health and some of them may be harmful by inadvertent activation. Host cells inactivate the integrated invaders by their transcriptional silencing via DNA methylation and modifications of adjacent histones. This is, however, an obstacle in using retroviruses as vectors for gene transfer and transgenesis. In the years 2006 and 2007, we have identified a novel semiresistant variant of chicken receptor for ASLV-B and described the ASLV-induced wasting disease in chicken. We have described that insertion of a core element from CpG island into retroviral vectors improves their resistance to transcriptional silencing and ensures long-term expression of such vectors. We have successfully used a retroviral vector for transduction of reporter genes in chicken male germ line, which opens the way to efficient transgenesis in chicken. We have also characterized the CpG methylation patterns of human syncytins, endogenous retroviruses involved in differentiation of human placenta syncytiotrophoblast. Furthermore, we are intersted in porcine endogenous retroviruses as a potential risk factor in xenotransplantation of pig organs and tissues.

Current grant support

Ministry of Education, Youth and Sports (Center LC-06061), GA CR (204/05/0939, 204/07/1030, 523/07/1171, 523/07/1282), GA AS CR (A500520709), FP6 International project XENOME

Selected recent papers

- Matoušková M, Blažková J, Pajer P, Pavlíček A, Hejnar J. CpG methylation suppresses transcriptional activity of human syncytin-1 in non-placental tissues. Exp Cell Res. 2006;312:1011-1020.
- Průková D, Vernerová Z, Pilčík T, <u>Stepanets V</u>, Indrová M, <u>Geryk J</u>, <u>Plachý J</u>, <u>Hejnar J</u>, <u>Svoboda J</u>. Differences in pathogenicity among ALV strains belonging to the same subgroup. **Avian Pathol**. 2007;36:15-27.
- Kalina J, <u>Šenigl F</u>, Mičáková A, Mucksová J, <u>Blažková J</u>, Poplštein M, <u>Hejnar J</u>, Trefil P. Retrovirus-mediated in vitro gene transfer into chicken male germ line cells. **Reproduction.** 2007:134:445-453.
- 4. <u>Reinišová M, Šenigl F, Yin X, Plachý J, Geryk J, Elleder D, Svoboda J, Federspiel MJ, Heinar J.</u> A single amino acid substitution in the TvbS1 receptor results in the semi-resistant phenotype of an inbred chicken line to infection by subgroup B and D avian sarcoma and leukosis viruses. **J Virol**; in press.



Comparison of the CpG methylation within syncytin-1 LTR in HeLa (upper part) and BeWo (lower part) cells