Pavel Hozák

hozak@img.cas.cz

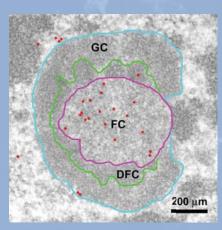


Laboratory of Biology of the Cell Nucleus

Regulation of gene transcription, nucleoskeleton, nuclear actin, myosin



Pavel Hozák, Prof, DSc / Head of Laboratory Zdeněk Hodný, MD, PhD / Research Scientist Vlada V. Philimonenko, PhD / Research Scientist Michal Kahle, MD, PhD / Research Scientist Michael Murtagh. PhD / Research Scientist Lenka Jarolímová, PhD / Research Assistant Ivana Nováková / Technician Iva Jelínková / Technician Vladimíra Bayerová / Technician Pavel Kříž / Technician Zora Nováková, MSc / PhD Student Lenka Rossmeislová, MSc / PhD Student Jana Vlasáková. MSc / PhD Student Helena Fulková, MSc / PhD Student Jana Hofmannová, MSc / PhD Student Rastislav Dzijak, MSc / PhD Student Anatolij Philimonenko, MSc / PhD Student Miloslava Maninová, MSc / PhD Student Sukriya Yildirim, MSc / PhD Student Jakub Kukla / Diploma Student Tomáš Chum / Diploma Student Lenka Pišlová / Secretary



Ultrastructural detection of actin molecules (red dots) in the nucleolus of resting human lymphocyte

Research topics

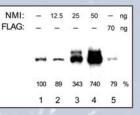
In diploid mammalian cells, some 6x10⁹ base pairs of DNA fold as a nucleoprotein complex (i.e. chromatin) into higher-order arrays so as to fit in a nucleus measuring only 10 µm. The nucleus also contains machineries for transcription of genes and processing of RNA products, and for precise DNA replication, repair and recombination. Nuclear interior is therefore functionally highly compartmentalized, and the recent evidence points strongly to structure-related regulation of nuclear functions – however, the mechanisms forming the 3D-structure of the nucleus are still mostly obscure. We therefore employ a multi-disciplinary approach in order to study nuclear functions in relation to the higher-order nuclear structures, e.g. nuclear bodies, the nucleolus, and the nucleoskeleton. Our research concentrates on: (1) the relationship between nuclear compartmentalization and regulation of gene expression, (2) structure, dynamics, and function of the nucleoskeleton which might direct nuclear compartmentalization, (3) functions of nuclear myosin I and actin in transcription and gene expression, (4) development of new microscopy methods for ultrastructural studies.

Current grant support

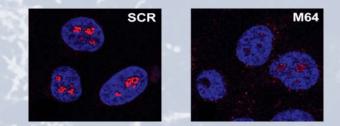
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Selected recent papers

- <u>Philimonenko AA, Hodny Z</u>, Jackson DA, <u>Hozak P</u>. The microarchitecture of DNA replication domains. Histochem Cell Biol. 2006;125:103-17.
- Janderova-Rossmeislova L, Novakova Z, Vlasakova J, Philimonenko V, Hozak P, Hodny Z. PML protein association with specific nucleolar structures differs in normal, tumor and senescent human cells. J Struct Biol. 2007;159:56-70.
- <u>Vlasakova J, Novakova Z, Rossmeislova L, Kahle M, Hozak P, Hodny Z</u>. Histone deacetylase inhibitors suppress IFNalpha-induced up-regulation of promyelocytic leukemia protein. Blood. 2007;109:1373-80.
- <u>Kahle M</u>, <u>Pridalova J</u>, <u>Spacek M</u>, <u>Dzijak R</u>, <u>Hozak P</u>. Nuclear myosin is ubiquitously expressed and evolutionary conserved In vertebrates. Histochem Cell Biol. 2007;127:139-48.



Addition of nuclear myosin I accelerates transcription of ribosomal genes in vitro (Northern blot, visualization of rRNA transcripts)



Depletion of NMI by siRNAs reduces rDNA transcription (red – transcripts, blue – DNA)