

Georges Chalepakis - CV

Georges Chalepakis
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Education – Academic appointments

Education:

- 1978-1984 Technical University Darmstadt, Germany, study of Chemistry
- 1983 Diploma, final examination
- 1983-1984 Preparation of diploma thesis “Identification of α -chains of collagen I in gelatine” (supervisor: Prof. E. Heidemann) Institute of Macromolecular Chemistry, Department of Protein Chemistry, Darmstadt
- 1985-1988 Preparation of doctoral thesis “Interaction of Progesterone- and Glucocorticoid-Receptor with the regulatory elements of the Mouse Mammary Tumor Virus (MMTV)” (supervisor: Prof. M. Beato) Institute of Molecular Biology and Tumor Research, Marburg, Germany
- 1988 Doctorate, Marburg University, Germany
- 1988-1990 Postdoctoral assistant at the Institute of Molecular Biology and Tumor Research, Marburg, Germany. Research field: Gene regulation by steroid hormone receptors
- 1990-1991 Postdoctoral fellow (Stipend of Boehringer-Ingelheim), Max-Planck-Institute of Biophysical Chemistry, Goettingen, Germany, Department of Molecular Cell biology in the group of Prof. P. Gruss
- 1991-1996 Senior scientist in Max-Planck-Institute of Biophysical Chemistry, Goettingen
Research topics: Functional analysis of Pax-gene products and identification of neuronal specific genes using the gene-trap method in mouse

Positions:

- 1996-2008 Associate Professor of Cell Biology at the Department of Biology, University of Crete
- 2009 Professor of Cell Biology at the Department of Biology, University of Crete
- 2005-today Director of the laboratory of Electron Microscopy
G. Chalepakis is the director of the laboratory of Electron Microscopy 'Vassilis Galanopoulos' which belongs to the Faculty of Sciences and Technology of the University of Crete. The facility offers its services not only to the scientific community of the University of Crete but to the whole of Greece. The activities of the laboratory are not confined only to basic research, as electron microscopy is a useful tool for dermatologists, ophthalmologists and nephrologists for the diagnosis of specific pathologies.
The laboratory is well equipped with two Transmission Electron Microscopes (TEM), a JEOL-100C and a high resolution JEM-2100 microscope (which provide solutions for a wide range of problems in the fields of materials, nanoelectronics and biological sciences). Two Scanning Electron Microscopes, a JSM 840 and a JSM-6390LV complement the facility of electron microscopy. In addition, the laboratory is equipped with a low energy ion milling for sample

preparation for nanoanalytical electron microscopy, a sputtering SCD 050 sample coater for SEM, a critical point dryer CPD 030, a cryo-preparation Leica EM AFS, an ultramicrotome LKB 2088 and knife maker LKB 7800B. Furthermore, the facility is equipped with a confocal Microscope SP1 LEICA, an optical microscope NIKON eclipse E800 and an Axiovert D1 Time-lapse microscopy system. The microscopes are fully equipped for routine activities such as simple observation of biological tissue preparations and nanomaterials.

Research Interests

During his career G. Chalepakis has been involved in different projects including functional characterization of gene products as well as developmental genetics using gene manipulation techniques in mouse. His major interest over the time between 1990 and 1996 was to elucidate the molecular and cellular function of proteins (Pax-proteins) which have been associated with mouse mutants and human syndromes (mouse mutants: undulated, splotch, small eye; Human syndromes: Waardenburg, Aniridia). G. Chalepakis has also been involved in a large scale gene-trap project in the Group of Dr P. Gruss at the Max-Planck-Institute in Goettingen (Germany) with the aim to create as many random mutations in the mouse genome as possible. In September 1996, G. Chalepakis took his present position at the University of Crete. His group has isolated and characterized the *Fras1* gene in mouse and in collaboration with Dr P. Scambler (UK) has shown that mutations in the human counterpart *FRAS1*, are responsible for the Fraser syndrome. Fraser syndrome is a rare genetic disorder with autosomal recessive inheritance pattern, primarily characterized by cryptophthalmos, syndactyly and renal agenesis. The group of G. Chalepakis has generated the *Fras1* deficient mice which serve as model to investigate the molecular pathology of the human Fraser syndrome. In addition to *Fras1*, the family of *Fras1*/*Frem* proteins comprises three additional members, *Frem1*, *Frem2* and *Frem3* which interact with each other to form an interdependent macromolecular protein complex within the basement membrane surrounding the embryonic epithelia. The major focus of the research group is to identify the proteins which interact with *Fras1*/*Frem*, aiming to explore the composition, the assembly and the protein-protein interactions of the extracellular matrix components that underlie epithelia and confer the structural cohesiveness as well as the functional interaction between epithelia and mesenchyme in mammals.

Publications

- 1) Chalepakis, G., Tanay, I. and Heidemann, E. (1985). Wie spezifisch ist der Kollagenabbau bei der Gelatineherstellung? **Das Leder** 36, 2-10.
- 2) Beato, M., Arnemann, J., Chalepakis, G., Slater, E. and Willmann, T. (1987). Gene regulation by steroid hormones. **J. Steroid Biochem.** 27, 9-14.
- 3) Chalepakis, G., Arnemann, J., Slater, E., Bruller, H.-J., Gross, B. and Beato, M. (1988). Differential gene activation by glucocorticoids and progestins through the hormone regulatory element of mouse mammary tumor virus. **Cell** 53, 371- 382.
- 4) Chalepakis, G., Postma, J.P.M. and Beato, M. (1988). A model for hormone receptor binding to the mouse mammary tumor virus regulatory element based on hydroxyl radical footprinting. **Nucl. Acids Res.** 16, 10237-10247.
- 5) Chalepakis, G. and Beato, M. (1989). Hydroxyl radical interference: a new method for the study of protein-DNA interactions. **Nucl. Acids Res.** 17, 1783.

- 6) Schauer, M., Chalepakis, G., Willmann, T. and Beato, M. (1989). Binding of hormone accelerates the kinetics of glucocorticoid and progesterone receptor binding to DNA. **Proc. Natl. Acad. Sci. USA** 86, 1123-1127.
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- 8) Chalepakis, G., Schauer, M., Cao, X.A. and Beato, M. (1989). Efficient binding of the glucocorticoid receptor to its responsive element requires two monomers and DNA flanking sequences. **DNA Cell. Biol.** 9, 355-368.
- 9) Truss, M., Chalepakis, G. and Beato, M. (1989). Contacts between steroid hormone receptors and thymines in DNA: An interference method. **Proc. Natl. Acad. Sci. USA** 87, 7180-7184.
- 10) Beato, M., Chalepakis, G., Schauer, M. and Slater, E. (1989). DNA regulatory elements for steroid hormones. **J. Steroid Biochem.** 32, 737-748.
- 11) Pina, B., Hache, R.J.G., Arnemann, J., Chalepakis, G., Slater, E.P. and Beato, M. (1990). Hormonal induction of transfected genes depends on DNA topology. **Mol. Cell. Biol.** 10, 625-633.
- 12) Goulding, M.D., Chalepakis, G., Deutsch, U., Erselius, J.R. and Gruss, P. (1991). Pax-3, a novel murine DNA binding protein expressed during early neurogenesis. **EMBO J.** 10, 1135-1147.
- 13) Truss, M.*, Chalepakis, G.*, Slater, E.P., Mader, S. and Beato, M. (1991). Functional interaction of hybrid response elements with wild-type and mutant steroid hormone receptors. **Mol. Cell. Biol.** 11, 3247-3258.
* οι δύο συγγραφείς έχουν συνεισφέρει ισάξια (equal first authors)
- 14) Chalepakis, G., Fritsch, R., Deutsch, U., Fickenscher, H., Goulding, M. and Gruss, P. (1991). The molecular basis of the undulated / Pax-1 mutation. **Cell** 66, 873-884.
- 15) Jones, F.S., Chalepakis, G., Gruss, P. and Edelman G.M. (1992). Activation of the cytotoxin promoter by the homeobox-containing gene *Evx-1*. **Proc. Natl. Acad. Sci. USA** 89, 2091-2095.
- 16) Truss, M., Bartsch, J., Chalepakis, G. and Beato, M. (1992). Artificial steroid hormone response element generated by dam-methylation. **Nucl. Acids Res.** 20, 1483-1486.
- 17) Truss, M., Chalepakis, G., Pina, B., Baretino, D., Bruggemeier, U., Kalff, M., Slater, E.P. and Beato, M. Transcriptional control by steroid hormones. Review (1992). **J. Steroid Biochem. Mol. Biol.** 41, 241-248.
- 18) Chalepakis, G., Tremblay, P. and Gruss, P. (1992). Pax genes, mutants and molecular function. **J. Cell Sci.** 16 (Suppl.), 61-76.
- 19) Truss, M., Chalepakis, G. and Beato, M. (1992). Interplay of steroid hormone receptors and transcription factors on the mouse mammary tumor virus promoter. **J. Steroid Biochem. Mol. Biol.** 43, 365-378.

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- 21) Chalepakis, G., Stoykova, A., Wijnholds, J., Tremblay, P. and Gruss, P. (1993). Pax: gene regulators in the developing nervous system. **J. Neurobiol.** 24, 1367-1384.
- 22) Chalepakis, G., Goulding, M., Read, A., Strachan, T. and Gruss, P. (1994). The molecular basis of *spotch* and *Waardenburg* Pax-3 mutations. **Proc. Natl. Acad. Sci. USA** 91, 3685-3689.
- 23) Chalepakis, G., Wijnholds, J., Giese, P., Schachner, M. and Gruss, P. (1994). Characterization of Pax-6 and Hoxa-1 binding to the promoter region of the neural cell adhesion molecule L1. **DNA Cell. Biol.** 13, 891-900.
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- 27) Song, D-Li, Chalepakis, G., Gruss, P. and Joyner, A.L. (1996). Two Pax-binding sites are required for early embryonic brain expression of an *Engrailed-2* transgene. **Development** 122, 627-635.
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- 30) McGregor, L., Makela, V., Darling, S.M., Vrontou, S., Chalepakis, G., Roberts, C., Smart, N., Rutland, P., Prescott, N., Hopkins, J., Bentley, E., Shaw, A., Roberts, E., Mueller, R., Jadeja, S., Philip, N., Nelson, J., Francannet, C., Perez-Aytes, A., Megarbane, A., Kerr, B., Wainwright, B., Woolf, A.S., Winter, R.M. and Scambler, P.J. (2003). Fraser syndrome and mouse blebbed phenotype caused by mutations in *FRAS1/Fras1* encoding a putative extracellular matrix protein. **Nat. Genet.** 34, 203-208.
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- 35) Petrou, P., Chiotaki, R., Dalezios, Y. and Chalepakis, G. (2007). Overlapping and divergent localization of Frem1 and Fras1 and its functional implications during mouse embryonic development. **Exp. Cell Res.** 313, 910-920.
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- 39) Petrou, P., Makrygiannis, A. and Chalepakis, G. (2008). The Fras1/Frem family of extracellular matrix proteins: Structure, function and association with Fraser syndrome and the mouse bleb phenotype. **Connect. Tissue Res.** 49, 277-282.
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