

MARIE SKŁODOWSKA-CURIE POSTDOCTORAL FELLOWSHIPS 2021
EXPRESSION OF INTEREST FOR HOSTING MARIE CURIE FELLOWS

HOST INSTITUTION

NOVA University Lisbon | NOVA Medical School

RESEARCH GROUP AND URL

Proliferation and fate regulation of stem cells.
<http://cedoc.unl.pt/proliferation-and-fate-regulation-of-stem-cells-3/>

SUPERVISOR (NAME AND E-MAIL)

Catarina Homem
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SHORT CV OF THE SUPERVISOR

EDUCATION AND PROFESSIONAL EXPERIENCE

2016-present: Group leader, Proliferation and fate regulation of stem cells. Chronic Diseases Research Center (CEDOC), NOVA Medical School (NMS), Lisbon, Portugal.

11/2015-06/2016 – Maternity leave

2009-2015: PostDoc at the Institute of Molecular Biotechnology of Austria (IMBA), Vienna, Austria.

2004-2008: PhD in Biology, University of North Carolina at Chapel Hill-USA

2004-2009: PhD Student, Laboratory of Dr. Mark Peifer, Cell adhesion and signal transduction in *Drosophila*, University of North Carolina at Chapel Hill (UNC-Chapel Hill)

2002-2003: PhD candidate – 1st year of classes. 3rd Gulbenkian PhD Programme in Biomedicine (PGDB3), Gulbenkian Foundation/Luso-American Foundation (FLAD).

1996-2000: BSc in Biochemistry, Porto University, Portugal.

PUBLICATIONS

(*Publications cited more than 50x)

Oliveira AC, Rebelo AR, Homem CC (2021) Integrating animal development: How hormones and metabolism regulate developmental transitions and brain formation. *Dev Biol* 4;S0012-1606(21)00029-4

Garcez M, Branco-Santos J, Gracio PC, Homem CC (2021) Mitochondrial Dynamics in the *Drosophila* Ovary Regulates Germ Stem Cell Number, Cell Fate, and Female Fertility. *Front Cell Dev Biol* 28;8:596819.

Rebelo AR, Garcez M, Homem CC. (2020) Tumor start-up: mitochondrial fusion makes it happen. *EMBO J* 1;39(23):e106927.

*Homem CC, Repic M, Knoblich JA (2015) Proliferation control in neural stem and progenitor cells. *Nat Rev Neurosci*. 16, 647-59.

*Homem CC, Steinmann V, Burkard TR, Jais A, Esterbauer H, Knoblich JA (2014) Changes in energy



metabolism triggered by Ecdysone and Mediator end proliferation in *Drosophila* neural stem cells. *Cell* 158, 874-888. Preview in *Cell Stem Cell* 15, 262-264, 2014

*Eroglu E, Burkard TR, Jiang Y, Saini N, Homem CC, Reichert H, Knoblich JA (2014) SWI/SNF complex prevents lineage reversion and induces temporal patterning in *Drosophila* neural stem cell lineages. *Cell* 156, 1259-1273.

Homem CC, Reichardt I, Berger C, Lendl T, Knoblich JA (2013) Long-Term Live Cell Imaging and Automated 4D Analysis of *Drosophila* Neuroblast Lineages. *PLoS ONE* 8(11): e79588. doi:10.1371/journal.pone.0079588.

*Homem CC, Knoblich JA (2012) *Drosophila* neuroblasts: a model for stem cell biology. *Development* 139, 4297-4310.

*Homem CC, Peifer M (2009) Exploring the roles of diaphanous and enabled activity in shaping the balance between filopodia and lamellipodia. *Mol Biol Cell* 20, 5138-5155.

Gates J, Nowotarski SH, Yin H, Mahaffey JP, Bridges T, Herrera C, Homem CC, Janody F, Montell DJ and Peifer M (2009) Enabled and Capping protein play important roles in shaping cell behavior during *Drosophila* Oogenesis. *Dev Biol* 333, 90-107.

*Homem CC, Peifer M (2008) Diaphanous regulates myosin and adherens junctions to control cell contractility and protrusive behavior during morphogenesis. *Development* 135, 1005-1018. Journal highlight: <http://dev.biologists.org/cgi/reprint/135/6/e601>

(Highly Recommended by Faculty of 1000)

Stevens TL, Rogers EM, Koontz LM, Fox DT, Homem CC, Nowotarski SH, Artabazon NB, and Peifer M (2008) Using Bcr-Abl to examine mechanisms by which Abl kinase regulates morphogenesis in *Drosophila*. *Mol Bio Cell* 19, 378-393.

*Fox DT, Homem CC, Myster SH, Wang F, Bain EE, Peifer M (2005) Rho1 Regulates *Drosophila* Adherens Junctions Independently of p120ctn. *Development* 132, 4819-4831.

AWARDS AND FELLOWSHIPS

2018: ERC Starting Grant (European Research Council)

2017: HHMI and Wellcome Trust International Scholar (Howard Hughes Medical Institute)

2016: EMBO Installation Grant

2015: Starting FCT Investigator Grant

2015: Exploratory Project FCT

2009-2011: EMBO long term Postdoctoral fellowship

2002-2007: 5-year PhD studentship in Biomedicine awarded by Foundation Calouste Gulbenkian and Foundation for Science and Technology (FCT)

REVIEWER AND EDITORIAL ACTIVITIES

2021-present: Review editor for Stem Cell Research (specialty section of *Frontiers in Cell and Developmental Biology*, *Frontiers in Genetics*, *Frontiers in Oncology* and *Frontiers in Bioengineering and Biotechnology*)

2019-present: Member of Editorial Board of *Plos Biology*

2017-present: Reviewer for *E-Life* (Awarded a recognition gift)

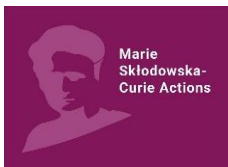
2017-present: Reviewer for *Developmental Biology* (Awarded a Certificate of Outstanding contribution)

2017-present: Reviewer for *EMBO Journal*

2013-2015: Co-Reviewer for journals as *Cell*, *Cell Stem Cell* or *Science*

5 SELECTED PUBLICATIONS

- Oliveira AC, Rebelo AR, Homem CC (2021) Integrating animal development: How hormones and metabolism regulate developmental transitions and brain formation. *Dev Biol* 4;S0012-1606(21)00029-4



- Garcez M, Branco-Santos J, Gracio PC, Homem CC (2021) Mitochondrial Dynamics in the Drosophila Ovary Regulates Germ Stem Cell Number, Cell Fate, and Female Fertility. *Front Cell Dev Biol* 28;8:596819.
- Rebelo AR, Garcez M, Homem CC. (2020) Tumor start-up: mitochondrial fusion makes it happen. *EMBO J* 1;39(23):e106927.
- *Homem CC, Repic M, Knoblich JA (2015) Proliferation control in neural stem and progenitor cells. *Nat Rev Neurosci.* 16, 647-59
- *Homem CC, Knoblich JA (2012) Drosophila neuroblasts: a model for stem cell biology. *Development* 139, 4297-4310.

PROJECT TITLE AND SHORT DESCRIPTION

Metabolic control of Stem Cell fate: Epigenetic regulation by Acetyl-CoA levels.

Stem Cells (SC) are responsible for the development and the adult regenerative processes, giving rise to specialized cell types. Dysregulation of SC differentiation may not only cause developmental defects but can also cause serious diseases such as cancer. So, a full understanding of the mechanisms controlling stem cell fate regulation is crucial.

Metabolism has recently been proposed as one of the main elements involved in cell differentiation control. Several studies point out that the cellular metabolic state conveys to the epigenome through post-translational modifications of histones[1]. The epigenetic changes alter chromatin dynamics, strongly implicated in cell fate commitment by transcriptome regulation[2]. The metabolic intermediate Acetyl-CoA is the substrate of histone acetylation reactions and has been suggested to play a relevant role in the metabolic regulation of stemness[3]. Despite the increasing evidence of its importance, it is still unknown how Acetyl-CoA levels impact the epigenome and the fate transcriptional programs. The goal of this proposal is to identify the mechanisms by which Acetyl-CoA regulates epigenome and hence the transcriptome to control cell fate.

To address this objective, we will use Drosophila Neural Stem Cells (NSCs) –neuroblasts (NB) – as a model. We will generate several tools to manipulate the levels of Acetyl-CoA by overexpression or downregulation of the Acetyl-CoA generating and degradation pathways in NBs in vivo. We will quantitatively investigate the consequence of Acetyl-CoA abundance in levels of chromatin acetylation by mass spectrometry. We will more specifically ask if Acetyl-CoA levels trigger region and gene-specific changes by analyzing the acetylation patterns upon different Acetyl-CoA levels by Chip-Seq. We will further look for interactions between Acetyl-CoA synthesis enzymes and histone’s acetylation complexes as a mechanism to confer histone modification region specificity. Ultimately, we will inquire about the transcriptional regulation of fate and stemness genes exerted by the epigenetics changes found.

Altogether, we seek to elucidate the role of Acetyl-CoA as an important regulator of epigenetics thus connecting metabolism and cell fate.

SCIENTIFIC AREA WHERE THE PROJECT FITS BEST*

Life Sciences (LIF)