# **TIAGO FLEMING OUTEIRO**

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Work

Instituto de Medicina Molecular Cell and Molecular Neuroscience Unit Av. Professor Egas Moniz 1649-028 Lisboa Portugal

### **EDUCATION**

4/99 - 7/04	Ph.D. in Molecular and Cell Biology
	Whitehead Institute for Biomedical Research, MIT, Cambridge, USA
	University of Chicago (UC), Chicago, USA
7/98-9/98	Teaching Training ("Curso de Formação de Formadores")
	Employment Institute (IEFP), Portugal
9/94-9/98	B.S. in Biochemistry
	Faculty of Sciences – University of Porto, Porto, Portugal

## RESEARCH AND PROFESSIONAL EXPERIENCE

12/07 – present 5/07 – 7/08	Auxiliar Professor, Instituto de Fisiologia, Faculdade de Medicina da Universidade de Lisboa, Portugal Visiting Scientist, Massachusetts General Hospital, Harvard Medical School, Boston, USA.
5/07 –	<b>Principal Investigator</b> , Institute of Molecular Medicine, University of Lisbon,
present	Portugal
11/05	<b>Co-Founder,</b> BioEPI Clinical and Translational Research Center, Taguspark, Portugal
6/04-5/07	Postdoctoral Research Fellow; advisor Dr. Brad Hyman, MGH – Harvard
0/04/0/04	University, USA
6/04-9/04	Consultant and Research Scientist, FoldRx Pharmaceuticals, Inc, Cambridge,
	USA
	Ph.D. work was transferred to the start up company FoldRx Pharmaceuticals, Inc.
4/99 – 8/04	Graduate Research; advisor Dr. Susan Lindquist
	The Whitehead Institute for Biomedical Research, MIT, Cambridge, USA
	University of Chicago, Chicago, USA
11/98-4/99	Graduate Research; advisor Prof. M. J. Saraiva
	University of Porto/IBMC, Porto, Portugal
2/98-9/98	Undergraduate Research; advisor Prof. John Findlay
_,	University of Leeds, Leeds, UK
9/97-2/98	Research Assistant;
5,5. 2,66	Centro de Estudos da Paramiloidose, Hospital Geral de Santo Antonio, Porto, Portugal

## HONORS AND AWARDS

10/10	Research Grant from the Michael J. Fox Foundation, USA.
3/09	Junior Faculty Award, AD/PD Meeting, Prague, Czech Republic
12/08	EMBO Installation Grant, EMBO
7/08	Aging Research Prize, Portuguese Society for Neuroscience and Pfizer
3/08	Best Scientific Article in 2007, Portuguese Society for Neuroscience
8/07	Marie Curie International Reintegration Grant, European Commission
7/07	Research Talent Award from the Portuguese Government
7/07	Science Talent Award from the Portuguese Government
6/07	Research Grant from the Michael J. Fox Foundation, USA.
6/07	Research Award from the Calouste Gulbenkian Foundation (FCG)
7/05	Massachusetts Biomedical Research Corporation, Tosteson Award,
	Postdoctoral Fellowship
1/03	Scholarship from the Calouste Gulbenkian Foundation (FCG)
1/99 - 03	PhD Scholarship from the Foundation for Science and Technology (Portugal)

2/98	Erasmus Scholarship from the European Union
5/03	<b>Young Investigator Award</b> , Gordon Research Conference on Triplet Repeat Disorders
5/01	<b>Young Investigator Award</b> , Gordon Research Conference on Triplet Repeat Disorders
11/00	<b>Prize from ICBAS</b> , 1st Place award, 94-98 class at the Biomedical Institute, Porto, Portugal
9/00	<b>Prize from the Engineer Antonio de Almeida Foundation</b> , 1 <sup>st</sup> Place award 94-98 class in Biochemistry, Porto, Portugal

### **PROFESSIONAL ACTIVITIES**

May 2009 –	Member of the Scientific Advisory Board for the European Huntington's
present	Disease Network
July 2008 –	Contributing Editor, European Journal of Neuroscience
present	
2007 - present	International Forum of Portuguese Researchers (FIIP) – Vice-President
2007	Expert evaluator – European Commission, Framework Program 7
2007	Organizer of the PENS Summer School on: "Molecular Mechanisms in
	Neurodegeneration", Ofir, Portugal.
2006	Expert evaluator – European Commission, Framework Program 6
2006	Organizer of FENS Symposium: "CAG Triplet Repeat Disorders", Vienna,
	Austria.
2004	Organizer of FEBS Course: "Neurodegeneration and Disease", Ofir, Portugal.
9/02 - 9/03	Organizer of the Boston Area Yeast Meeting, Boston, USA
1/02 -present	Portuguese American Post-Graduate Association (PAPS) –
	President/Chairman
	Co-organizer of IV, V, VI and VII Forum of PAPS.
9/02	Organizer and Participant University of Aveiro Center for Cell Biology
	Symposium - Novel Therapeutic Opportunities in Neurodegenerative Disease
10/02 - 04	Whitehead Institute Scientist – Partnership for Education with teachers from
	Massachusetts
1/02	Organizer - MIT European Career Fair (over 2500 participants)
12/01	Organizer and Participant - Science for the Non-scientist - MIT Museum

#### **PUBLICATIONS**

da Costa G, Guerreiro A, Correia CF, Gomes RJ, Freire A, Monteiro E, Barroso E, Coelho AV, Outeiro TF, Freire AP, Cordeiro C.	A non-invasive method based on saliva to characterize transthyretin in familial amyloidotic polyneuropathy patients using FT-ICR high-resolution MS. Proteomics Clin Appl. 2010 Jul;4(6-7):674-8. PubMed PMID: 21179887
Büttner S, Delay C, Franssens V,	Synphilin-1 enhances α-synuclein aggregation in yeast and
Bammens T, Ruli D, Zaunschirm	contributes to cellular stress and cell death in a Sir2-
S, de Oliveira	dependent manner. PLoS One. 2010 Oct 27;5(10):e13700.
RM, Outeiro TF, Madeo F, Buée	PubMed
L, Galas MC, Winderickx J.	PMID: 21060871
Tauber E, Miller-Fleming L,	Functional gene expression profiling in yeast implicates
Mason RP, Kwan W, Clapp J,	translational dysfunction in mutant huntingtin toxicity. J Biol
Butler NJ, Outeiro TF,	Chem. 2010 Nov 2. [Epub ahead of print] PubMed PMID:
Muchowski PJ, Giorgini F	21044956.
Diógenes MJ, Outeiro TF	Neurotrophic Factors as a Protective Strategy in Parkinson's

m. 2010 Nov 2. [Epub ahead of print] PubMed PMID: 44956. Neurotrophic Factors as a Protective Strategy in Parkinson's Disease. CNS Neurol Disord Drug Targets. 2010 Nov 12.

[Epub ahead of print] PubMed PMID: 20942787. Drug Discovery for CNS Disorders: From Bench to Bedside.

CNS Neurol Disord Drug Targets. 2010 Nov 12. [Epub ahead of print]

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de Oliveira RM, Pais TF, Outeiro TF

Outeiro TF, Kazantsev AG

Sirtuins: common targets in aging and in neurodegeneration. Curr Drug Targets. 2010 Oct;11(10):1270-80. PubMed PMID: Marques SC, Oliveira CR, Pereira CM, Outeiro TF.

de Oliveira RM, Pais TF, Outeiro TF.

Tenreiro S, Outeiro TF.

Gonçalves SA, Matos JE, Outeiro TF.

Marques SC, Oliveira CR, Outeiro TF, Pereira CM.

Nakhjavani M, Morteza A, Khajeali L, Esteghamati A, Khalilzadeh O, Asgarani F, Outeiro TF. Miranda, HV, and Outeiro, TF

Outeiro TF, Su LJ, Auluck PK, Yeger-Lotem E, Kritzer JA, Tardiff DF, Strathearn KE, Liu F, Cao S, Hamamichi S, Hill KJ, Caldwell KA, Bell GW, Fraenkel E, Cooper AA, Caldwell GA, McCaffery JM, Rochet JC, Lindquist S. Marques, O., Oliveira, RM., Moita, LF, and Outeiro, TF.

da Costa G, Gomes R, Correia CF, Freire A, Monteiro E, Martins A, Barroso E, Coelho AV, Outeiro TF, Ponces Freire A, Cordeiro C. Ferreira. J. and Outeiro. TF

Outeiro, TF, Klucken, J, Bercury, K, Tetzlaff, J, Putcha, P, Oliveira, LM, Quintas, A, McLean, PJ, and Hyman, BT Outeiro, TF

Mollenhauer B, Cullen V, Kahn I, Krastins B, Outeiro TF, Pepivani I, Ng J, Schulz-Schaeffer W, Kretzschmar HA, McLean PJ, Trenkwalder C, Sarracino DA, Vonsattel JP, Locascio JJ, El-Agnaf OM, Schlossmacher MG. Tetzlaff JE, Putcha P, Outeiro TF, Ivanov A, Berezovska O, Hyman BT, McLean PJ.

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Epigenetics in neurodegeneration: A new layer of complexity. Prog Neuropsychopharmacol Biol Psychiatry. 2010 Aug 22. [Epub ahead of print] PubMed PMID: 20736041.

Sirtuins: Common Targets in Aging and in Neurodegeneration. Curr Drug Targets. 2010 Jul 1. [Epub ahead of print] PubMed PMID: 20594174.

Simple is good: yeast models of neurodegeneration. FEMS Yeast Res. 2010 May 20. [Epub ahead of print] PubMed PMID: 20579105.

Zooming into protein oligomerization in neurodegeneration using BiFC. Trends Biochem Sci. 2010 Jun 17. [Epub ahead of print] PubMed PMID: 20561791.

Alzheimer's Disease: The Quest to Understand Complexity. J Alzheimers Dis. 2010 Jun 16. [Epub ahead of print] PubMed PMID: 20555132.

Increased serum HSP70 levels are associated with the duration of diabetes. Cell Stress Chaperones. 2010 May 23. [Epub ahead of print] PubMed PMID: 20496051.

The sour side of neurodegenerative disorders: the effects of protein glycation. J Pathol. 2010 May;221(1):13-25. Review. PubMed PMID: 20186922.

Compounds from an unbiased chemical screen reverse both ER-to-Golgi trafficking defects and mitochondrial dysfunction in Parkinson's disease models. Dis Model Mech. 2009 Dec 28.

Sirtuins as a link between ageing and neurodegeneration, Neurodegenerative Diseases, 2009.

Identification and quantitative analysis of human transthyretin variants in human serum by Fourier transform ion-cyclotron resonance mass spectrometry. Amyloid. 2009 Dec;16(4):201-207.

Current and Future Therapeutic Strategies for Parkinson's Disease, Curr Pharm Des. 2009 Sep 15.
Dopamine-Induced Conformational Changes in Alpha-Synuclein, PLoS One. 2009 Sep 4;4(9):e6906.

From Mad Cows to Neurotic Yeast: Novel Molecular Approaches to Understand Neurodegeneration.

Microsc Microanal. 2008 Sep;14 Suppl 3:105-6.

Direct quantification of CSF alpha-synuclein by ELISA and first cross-sectional study in patients with neurodegeneration.

Exp Neurol. 2008 Jun 14. [Epub ahead of print]

CHIP Targets Toxic {alpha}-Synuclein Oligomers for Degradation.

J Biol Chem. 2008 Jun 27;283(26):17962-8. Epub 2008 Apr 24.

Formation of toxic oligomeric a-synuclein species in living cells, PloS ONE, 2008 Apr 2;3(4):e1867.

Drug Targeting of  $\alpha$ -Synuclein Oligomerization in Synucleinopathies, Perspectives in Medicinal Chemistry

Fleming, L., Giorgini, F. and Outeiro, T.F. \*Grammatopoulos TN, \*Outeiro TF, Hyman BT, Standaert DG.

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Outeiro, T.F., Grammatopoulos, TN, Altmann, S., Amore, A., Standaert, D.G., Hyman, B.T., Kazantsev, A.G. Outeiro, T.F. and Tetzlaff, J.

St. Martin, J, Klucken, J., Outeiro, T.F., Nguyen, P., Keller-McGandy, C., Cantuti-Castelvetri, I., Grammatopoulos, T., Standaert, D., McLean, P.J. and Hyman, B.T. Outeiro TF, Klucken J, Strathearn KE, Liu F, Nguyen P, Rochet JC, Hyman BT, McLean PJ. Klucken, J., Outeiro, T.F., Nguyen, P., McLean, P.J. and Hyman, B.T. Outeiro, T.F.

Bodner, R., Outeiro, T.F., Altman, S., Maxwell, M.M., Cho, S. H., Hyman, B.T., McLean, P.J., Young, A.B., Housman, D. E. and Kazantsev, A. G.

Outeiro, T.F. and Giorgini, F.

Outeiro, T.F. and Muchowski, P.J.

Rochet JC, Outeiro TF, Conway KA, Ding TT, Volles MJ, Lashuel HA, Bieganski RM, Lindquist SL, Lansbury PT

Derkatch, I., Uptain, S., Outeiro, T.F., Liebman, S. and Lindquist, S.

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Willingham, S, Outeiro, T.F., Devit, MJ, Lindquist, S and Muchowski, PJ Resende, CG, Outeiro, T.F., Sands, L, Lindquist, S and Tuite, M 2008:2 41-49

Yeast as a model to study human neurodegenerative disorders, Biotechnology J., 2008. Mar;3(3):325-38. Angiotensin II protects against alpha-synuclein toxicity and reduces protein aggregation in vitro. Biochem Biophys Res Commun. 2007 Sep 21; [Epub ahead of print] Sirtuin 2 inhibition protects against a-synuclein toxicity in Parkinson's disease models, Science. 2007 Jul 7;317(5837):516-9.

Pharmacological Inhibition of PARP-1 Reduces Alpha-Synuclein- and MPP+-Induced Cytotoxicity in Parkinson's Disease In Vitro Models, BBRC, 2007, Jun 8;357(3):596-602.

Mechanisms of Disease II – Cellular Protein Quality Control, Seminars in Pediatric Neurology, 2007 Mar;14(1):15-25. Selevtive Dopaminergic Neuron Loss and Upregulation of Chaperone Protein mRNA in an Adeno-Associated Viral Vector Model of Parkinson's Disease, J. Neurochem, 2007 Mar;100(6):1449-57.

Small heat shock proteins protect against alpha-synucleininduced toxicity and aggregation. Biochem Biophys Res Commun. 2006 Dec 22;351(3):631-8.

Detection of novel intracellular a-synuclein oligomeric species by fluorescence lifetime imaging, FASEB J., FASEB J. 2006 Oct;20(12):2050-7

A bridge from Portugal to the States. Nature. 2006 Sep 7;443(7107):118.

Pharmacological promotion of inclusion formation: A therapeutic approach for Huntington's and Parkinson's disease, PNAS, 2006; 103 (11): 4246-4251

Yeast as a drug discovery platform in Huntington's and Parkinson's diseases, Biotecnol. J, 2006: 1(3): 258-269 Molecular genetics approaches in yeast to study amyloid diseases. J Mol Neurosci. 2004;23(1-2):49-60 Interactions Among alpha-Synuclein, Dopamine, and Biomembranes: Some Clues for Understanding Neurodegeneration in Parkinson's Disease. J Mol Neurosci. 2004;23(1-2):23-34

Effects of Q/N, polyQ and non-polyQ amyloids on the *de novo* formation of the [*PSI*<sup>†</sup>] prion in yeast and aggregation of Sup35 *in vitro*, PNAS, 2004; 101(35) 12934-9.

Yeast cells provide insight into alpha-synuclein biology and pathobiology. Science. 2003 Dec 5;302(5651):1772-5

Yeast genes that enhance the toxicity of a mutant huntingtin fragment or alpha-synuclein. Science. 2003 Dec 5;302(5651):1769-72

Prion protein gene polymorphisms in Saccharomyces cerevisiae. Mol Microbiol. 2003 Aug;49(4):1005-17

#### **BOOKS**

Vaqueiro-Lopes, L. and Outeiro, T.F.

Marques, S.C. F., Pereira, C.M.F., Outeiro, T.F. Gitler, A. and Outeiro, T.F.

Outeiro, T.F., editor

Synaptic dysfunction in Parkinson's disease: from protein misfolding to functional alterations, 2010.

Epigenetics and Neurodegeneration: A Connection Overlooked, Novascience, in press.

Unravelling the Molecular Basis of Parkinson's Disease Using Yeast Models, 2010.

Protein Misfolding in Biology and Disease, Research Signpost, Novascience 2009.

Outeiro, T.F. and Kazantsev, A. Therapeutic Intervention in the Neurotoxicty of Misfolded

Proteins, 2009.

Outeiro, T.F. and Hyman, B.T. | Protein Aggregation Disorders, in Neurobiology of Disease,

2006.

Singer, M., Outeiro, T.F. and

Lindquist, S.

Thermotolerance, Metabolism and Development:

The Many Flavors of Trehalose, in Food Biotechnology, 2005.

#### **PATENTS**

Outeiro, T.F., Lindquist, S., Labaudiniere, INHIBITION OF A-SYNUCLEIN TOXICITY. U.S.

Fleming, J., R., Bulawa, C. 60/787,113

Outeiro, T. F., and Lindquist, S.

YEAST ECTOPICALLY EXPRESSING
ABNORMALLY PROCESSED PROTEINS AND

USES THEREFOR

Outeiro, T.F., Krobitsch, S. and Lindquist, YEAST AS A MODEL SYSTEM FOR

NEURODEGENERATIVE DISEASE

#### SELECTED INVITED TALKS

January 2011
October 2010
September 2010
August 2010
February 2010
July 15, 2009

Lund University, Sweden
University of Granada, Spain
University of Aarhus, Denmark
Istanbul Technical University, Turkey
University of Gottingen, Germany
University of Ulm, Germany

July 15, 2009 University of Office Neuroches

July 13, 2009 European Society for Neurochemistry, Leipzig, Germany

July 11, 2009
June 4, 2009
May 12, 2009
March, 2009
December 5, 2008

Kopfklinic, Erlangen, Germany
Amsterdam, The Netherlands
Bilkent University, Turkey
University of Leuven, Belgium
University of Leicester, UK

November 2008 Instituto de Tecnologia Química e Biológica, Lisbon, Portugal

September 2008 EPFL, Lausanne, Switzerland

Febryary 27, 2008 From Mad Cows to Neurotic Yeast: Novel Strategies to Understand

Neurodegeneration. Faculdade de Ciências de Lisboa, Portugal

February 20, 2008 Mad Cows, Neurotic Yeast, and Back to the Future. Fundação Calouste

Gulbenkian

December 13, 2007 Unraveling the molecular mechanisms of neurodegeneration: From cells to

drugs, Uppsala University, Sweden

November 22, 2007 | Top Models: What are they teaching us about Parkinson's Disease.

Neurodegenerative diseases: Science and the Mind. Faculty of Pharmacy,

Lisbon, Portugal.

November 19, 2007 | From Academia to Biotech: US and Portuguese Examples. Pharmaceutical

Innovation: A New R&D Strategy in the EU. Viseu, Portugal.

October 26, 2007 Applied Biology Forum. University of Minho, Braga, Portugal.

October 16, 2007 | Science and Technology in Portugal. Fórum Novas Fronteiras, Centro

Cultural de Belém, Lisboa, Portugal. Promoted by and with the participation

of the Prime Minister of Portugal.

#### SELECTED POSTGRADUATE TEACHING

January 2011 Neurodegenerative Diseases Course, Gulbenkian PhD Program for Medical

Doctors.

December 2010 Master's/PhD in Neurosciences, Faculty of Medicine, Lisbon

June 2010 PhD Program, University of Coimbra

February 2010 Neurodegenerative Diseases Course, Gulbenkian PhD Program for Medical

Doctors.

July 2009 PENS Summer School, Gunzburg, Germany

March 2009 ITQB PhD Program

February 2009 Neurodegenerative Diseases Course, Gulbenkian PhD Program for Medical

Doctors.

January 2009 Protein Misfolding Course, GABBA PhD Program.

November 2008	Protein Misfolding in Biology and Disease, Master's Program in	
	Neurosciences, Faculty of Medicine, Lisbon, Portugal.	
February 2008	Secrets of the genome and biobanks of secrets, Center for Biomedical Law,	
•	University of Coimbra	
January 2008	PhD Program, University of Coimbra	
December 2007	Protein Misfolding in Synucleinopathies, Faculty of Sciences and	
	Technology, Monte da Caparica, Portugal.	
December 2007	Protein Misfolding in Biology and Disease, Master's Program in	
	Neurosciences, Faculty of Medicine, Lisbon, Portugal.	
July 2007	PENS Summer School, Ofir, Portugal.	
July 2004	FEBS Summer School, Ofir, Portugal.	