CURRICULUM VITAE **EUROPEAN FORMAT**

PERSONAL INFORMATION

Name, Surname

NICOLA GIANGREGORIO

Address

CNR INSTITUTE OF BIOMEMBRANES, BIOENERGETICS AND MOLECULAR BIOTECHNOLOGIES

C/O DEPARTMENT OF BIOSCIENCES, BIOTECHNOLOGIES AND BIOFARMACEUTICS

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n.giangregorio@ibiom.cnr.it

Nationality

Italian

Date of birth

05, AUGUST, 1966

WORK EXPERIENCE

• Dates (from - to)

19/03/2001 - to date

· Name and address of employer

CNR (National Council of Research) "Centre for the study of mitochondria and energetic metabolism", now become IBIOM (Institute of Biomembranes, Bioenergetics and Molecular Biotechnologies), Via Amendola 165 A, 70126 Bari (Italy)

• Type of business or sector

Life Sciences - Biochemistry and Molecular biology

· Occupation or position held

CNR Researcher

· Main activities and responsibilities

Scientific research

• Dates (from - to)

01/12/1995 - 18/03/2001

Name and address of employer

Department of Pharmaco-Biology, University of Bari, Via Orabona 4, 70125 Bari (Italy).

Type of business or sector

Life Sciences - Biochemistry and Molecular biology

Occupation or position held

Graduate laboratory technician

· Main activities and responsibilities

Scientific research

Dates (from – to)

1995

· Name and address of employer

CNR "Centre for the study of mitochondria and energetic metabolism", Via Amendola 165/A, 70126 Bari (Italy)

Type of business or sector

Life Sciences - Biochemistry and Molecular biology

· Occupation or position held

Research grant: "Finalized Project of Biotechnologies and Bio-Instrumentation"

· Main activities and responsibilities Scientific research

EDUCATION AND TRAINING

· Title of qualification awarded

Dates (from – to)

2018, March 28

National Scientific Habilitation for Associate Professor

 Dates (from – to) 1992-1995

 Name and type of organisation providing education and training Department of Pharmaco-Biology, University of Bari, Via Orabona 4, 70125 Bari (Italy)

 Principal subjects/occupational skills covered Cellular Biochemistry and Cellular Pharmacology

• Title of qualification awarded

PhD in Biochemistry

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Dates (from – to)

skills covered

 Name and type of organisation providing education and training

providing education and training
• Principal subjects/occupational

• Title of qualification awarded

1990

University of Bari, (Italy)

Biochemistry: "Purification of the mitochondrial carnitine carrier from rat liver".

Degree in Pharmacy

RESEARCH ACTIVITIES

Research Sectors and Scientific Activities

Expertise in biochemical and molecular biology techniques with particular regard in:

- identification, isolation and purification of membrane proteins;
- their reconstitution in artificial membranes (liposomes);
- functional and kinetic studies of transport proteins;
- identification of primary and secondary structure;
- cloning and protein over-expression in bacteria;
- site-directed mutagenesis;
- genetic pathologies correlated to mutations in mitochondrial carriers:
- protein chemical and functional modifications (xenobiotics compounds such as toxic molecules, drugs or newly synthesized compounds);
- identification of posttranslational modifications (GSH, H₂S, NO, acylations, etc.) and their regulator role of the protein function;
- use of bioinformatics for the comparative structural analysis of membrane proteins or docking;
- polyclonal antibodies production of membrane proteins and immunochemical detection

Pubblications

Scopus Citations Report (January 2020): 643; H-index = 14 Google Scholar Citations Report (January 2020): 803; H-index = 16

- 1. Bolognino I, **Giangregorio N**, Pisani L, de Candia M, Purgatorio R, Tonazzi A, Altomare CD, Cellamare S, Catto M, A Prospective Repurposing of Dantrolene as a Multitarget Agent for Alzheimer's Disease, Molecules (2019) 25, 24(23)
- Giangregorio N, Tonazzi A, Console L, Pistillo M, Scalera V, Indiveri C, Tryptophan 224 of the rat mitochondrial carnitine/acylcarnitine carrier is crucial for the antiport mechanism. Biochim Biophys Acta Bioenerg. (2019) 1860, 708-716
- 3. Scalise M, Console L, Galluccio M, Pochini L, Tonazzi A, **Giangregorio N**, Indiveri C. Exploiting Cysteine Residues of SLC Membrane Transporters as Targets for Drugs. SLAS Discov. (2019) 24, 867-881
- Console L, Giangregorio N, Cellamare S, Bolognino I, Palasciano M, Indiveri C, Incampo G, Campana S, Tonazzi A, Human mitochondrial carnitine acylcarnitine carrier: Molecular target of dietary bioactive polyphenols from sweet cherry (Prunus avium L.), Chemico-biological interactions (2019), 307, 179-185
- Giangregorio N, Tonazzi A, Console L, Galluccio M, Porcelli V, Indiveri C, Structure/function relationships of the human mitochondrial ornithine/citrulline carrier by Cys site-directed mutagenesis. Relevance to mercury toxicity, Int J Biol Macromol. (2018), 120, 93-99
- 6. Scalera V, **Giangregorio N**, De Leonardis S, Console L, Carulli ES, Tonazzi A, Characterization of a novel Mitochondrial Ascorbate Transporter from rat liver and potato mitochondria, Front Mol Biosci (2018), 5, 1-9
- 7. Console L, Scalise M, Tonazzi A, **Giangregorio N**, Indiveri C, Characterization of Exosomal SLC22A5 (OCTN2) carnitine transporter, Sci Rep (2018), 8, 3758
- 8. Pisani L, De Palma A, **Giangregorio N**, Miniero DV, Pesce P, Nicolotti O, Campagna F, Altomare CD, Catto M, Mannich base approach to 5-methoxyisatin 3-(4-isopropylphenyl)hydrazone: A water-soluble prodrug for a multitarget inhibition of cholinesterases, beta-amyloid fibrillization and oligomer-induced cytotoxicity, Eu J Pharm Sci (2017), 109, 381-388
- Scalise M, Galluccio M, Pochini L, Console L, Barile M, Giangregorio N, Tonazzi A, Indiveri C, Studying interactions of drugs with cell membrane nutrient transporters: new frontiers of proteoliposome nanotechnology, Curr Pharm Des (2017) 23, 3871-3883
- 10. Tonazzi A, Giangregorio N, Console L, De Palma A, Indiveri C, Nitric oxide inhibits

- the mitochondrial carnitine/acylcarnitine carrier through reversible S-nitrosylation of cysteine 136, Biochimica et Biophysica Acta Bioenergetics, (2017) 1858, 475-482
- 11. **Giangregorio N**, Tonazzi A, Console L, Indiveri C, Post-translational modification by acetylation regulates the mitochondrial carnitine/acylcarnitine transport protein, Mol Cell Biochem (2017), 426, 65-73
- 12. **Giangregorio N**, Tonazzi A, Console L, Lorusso I, De Palma A, Indiveri C, The mitochondrial carnitine/acylcarnitine carrier is regulated by hydrogen sulfide via interaction with C136 and C155, Biochimica et Biophysica Acta (BBA)-General Subjects (2016) 1860, 20-27
- Tonazzi A, Giangregorio N, Console L, Scalise M, La Russa D, Notaristefano C, Brunelli E, Barca D, Indiveri C, Mitochondrial Carnitine/Acylcarnitine Transporter, a Novel Target of Mercury Toxicity, Chemical research in toxicology (2015) 28, 1015-1022
- 14. Tonazzi A, **Giangregorio N**, Console L, Indiveri C, Mitochondrial Carnitine/Acylcarnitine Translocase: Insights in Structure/Function Relationships, Basis for Drug Therapy and Side Effects Prediction, Mini reviews in medicinal chemistry (2015) 15, 396-405
- 15. Console L, **Giangregorio N**, Indiveri C, Tonazzi A, Carnitine/acylcarnitine translocase and carnitine palmitoyltransferase 2 form a complex in the inner mitochondrial membrane, Molecular and cellular biochemistry (2014) 394, 307-314
- Giangregorio N, Console L, Tonazzi A, Palmieri F, Indiveri C, Identification of Amino Acid Residues Underlying the Antiport Mechanism of the Mitochondrial Carnitine/Acylcarnitine Carrier by Site-Directed Mutagenesis and Chemical Labeling, Biochemistry (2014) 53, 6924-6933
- 17. Scalise M, Pochini L, **Giangregorio N**, Tonazzi A, Indiveri C, Proteoliposomes as tool for assaying membrane transporter functions and interactions with xenobiotics, Pharmaceutics (2013) 5, 472-497
- 18. **Giangregorio N**, Palmieri F, Indiveri C, Glutathione controls the redox state of the mitochondrial carnitine/acylcarnitine carrier Cys residues by glutathionylation, Biochimica et Biophysica Acta (BBA)-General Subjects (2013) 1830, 5299-5304
- 19. Tonazzi A, Console L, **Giangregorio N**, Indiveri C, Palmieri F, Identification by site-directed mutagenesis of a hydrophobic binding site of the mitochondrial carnitine/acylcarnitine carrier involved in the interaction with acyl groups, Biochim Biophys Acta Bioenergetics (2012) 1817, 697-704
- Indiveri C, Iacobazzi V, Tonazzi A, Giangregorio N, Infantino V, Convertini P, Console L, Palmieri F. The mitochondrial carnitine/acylcarnitine carrier: Function, structure and physiopathology. Mol Aspects Med. 32 (2011) 223-33
- 21. **Giangregorio N**, Tonazzi A, Console L, Indiveri C, Palmieri F, Site-directed mutagenesis of charged amino acids of the human mitochondrial carnitine/acylcarnitine carrier: Insight into the molecular mechanism of transport, Biochim Biophys Acta 1797 (2010) 839-45
- 22. Tonazzi A, **Giangregorio N**, Indiveri C and Palmieri F, Site directed mutagenesis of the His residue of the rat mitochondria carnitine/acylcarntine carrier: implications for the role of His-29 in the transport pathway, Biochim Biophys Acta 1787 (2009) 1009-15
- 23. De Lucas JR, Indiveri C, Tonazzi A, Perez P, **Giangregorio N**, Iacobazzi V, Palmieri F, Functional characterization of residues within the carnitine/acylcarnitine translocase RX2PANAAXF distinct motif, Mol Membr Biol. 25 (2008) 152-63
- 24. Pochini L, Galluccio M, Scumaci D, **Giangregorio N**, Tonazzi A, Palmieri F, Indiveri C, Interaction of beta-lactam antibiotics with the mitochondrial carnitine/acylcarnitine transporter, Chem Biol Interact. 173 (2008) 187-94
- 25. **Giangregorio N**, Tonazzi A, Indiveri C, Palmieri F. Conformation-dependent accessibility of Cys-136 and Cys-155 of the mitochondrial rat carnitine/acylcarnitine carrier to membrane-impermeable SH reagents. Biochim Biophys Acta. 1767 (2007) 1331-9
- 26. Tonazzi A, **Giangregorio N**, Palmieri F and Indiveri C, Relationships of Cysteine and Lysine residues with the substrate binding site of the mitochondrial ornithine/citrulline carrier: an inhibition kinetic approach combined with the analysis of the homology structural model, Biochim Biophys. Acta 1718 (2005) 53-60
- 27. Tonazzi A, **Giangregorio N**, Indiveri C and Palmieri F, Identification by site-directed mutagenesis and chemical modification of three vicinal cysteine residues in rat mitochondrial carnitine/acylcarnitine transporter, J. Biol. Chem. 280 (2005) 19607-12
- 28. Indiveri C, **Giangregorio N**, Tonazzi A and Palmieri F. (2004). Site-directed mutagenesis of the mitochondrial carnitine/acylcarnitine carrier: identification of four

- vicinal cysteine residues. Biochim Biophys. Acta Bioenergetics, vol. 1658, ISSN: 0005-2728
- 29. Indiveri C, **Giangregorio N**, lacobazzi V, Palmieri F, Site-directed mutagenesis and chemical modification of the six native cysteine residues of the rat mitochondrial carnitine carrier: implications for the role of cysteine-136, Biochemistry 41 (2002) 8649-56
- Indiveri C, Iacobazzi V, Giangregorio N, Palmieri F, Bacterial overexpression, purification, and reconstitution of the carnitine/acylcarnitine carrier from rat liver mitochondria, Biochem Biophys Res Commun. 249 (1998) 589-94
- Indiveri C, Iacobazzi V, Giangregorio N, Palmieri F, The mitochondrial carnitine carrier protein: cDNA cloning, primary structure and comparison with other mitochondrial transport proteins, Biochem J. 321 (1997) 713-9
- 32. Indiveri C, Tonazzi A, **Giangregorio N**, Palmieri F, Probing the active site of the reconstituted carnitine carrier from rat liver mitochondria with sulfhydryl reagents. A cysteine residue is localized in or near the substrate binding site, Eur J Biochem. 228 (1995) 271-8

ADDITIONAL INFORMATION

Recent Funded Research Projects

03/10/2011 - 31/05/2015

PON 2007-2013 Progetto cod. 01-00937 - Modelli sperimentali biotecnologici integrati per lo sviluppo e la selezione di molecole di interesse per la salute dell'uomo

Management

2009 to April 2016

Member of the Institute Committee of IBBE (Institute of Biomembrane and Bioenergetics)-CNR

Member of the organizing committee of the Italian Group of Biomembranes and Bioenergetics (GIBB) meeting, Matera, May 29-31, 2014

Academic appointments

Lecturer in charge at the University of Bari in the following specified academic years and course:

- (2019/2020), (2018/2019), (2017/2018) "Animal and Vegetable Biology" (100 hours course of frontal lessons, Faculty of Pharmacy, degree course in Pharmacy)
- (2016/2017), (2015/16), (2014/15) "Animal and Vegetable Biology" (80 hours course of frontal lessons, Faculty of Pharmacy, degree course in Pharmacy)
- (2013/14), (2012/13), (2011/12), (2010/11) "Animal and Vegetable Biology" (60 hours course of frontal lessons, Faculty of Pharmacy, degree course in Pharmacy)
- 2012 to 2018: Board member of the PhD School in "Genomica e Proteomica Funzionale Applicata", Department of Biosciences, Biotechnologies and Biopharmaceutics, University of Bari A. Moro
- 2012 to 2018: Board member of the PhD School in "Scienze Biochimiche e Farmacologiche", Department of Pharmacy Pharmaceutical Sciences, University of Bari A. Moro

Lecturer in charge at the University of Calabria in the following specified academic years and course:

- (2011/12), (2010/11), (2009/10) "Biochemistry and molecular biology applicated" (32 hours course of frontal lessons, Faculty of SMFN, degree course in Biological Science)
- (2011/12), (2010/11), (2009/10), (2008/09), (2007/08), (2006/07), (2005/06) "Biochemistry of the cellular membrane" (32 hours course of frontal lessons, Faculty of SMFN, degree course in Biological Science)
- (2008/09), (2007/08), (2006/07), (2005/06), (2004/05), (2003/04), "Applicated Biology II" (32 hours course of frontal lessons, Faculty of SMFN, degree course in Biological Science)
- (2004/05), (2003/04) "Cellular Biochemistry" (16 hours course of frontal lessons, Faculty of SMFN, degree course in Biological Science)
- (2004/05) " **Biochemistry of the cellular membranes**" (16 hours course of frontal lessons, Faculty of SMFN, degree course in Biological Science)

Teacher in Master courses:

(2009), (2007), (2005): "Protein Engineering Course" in "Master of Biotechnology", University of Calabria

Training

- Formation course in " Confocal microscopy". Bari, 2012
- Formation course in "Cytofluorimetry". Bari, 11-12 May 2010
- Formation course in " Mass spectrometry for the study of proteomic and genomic". Bari, 9-10 December 2003
- Visiting Ph.D. student at "Institut fur Biotechnologie" of Julich (Germany), headed by Prof. Reinhard Krämer. from January to April 1993, studying the transport mechanism of the mitochondrial ADP/ATP carrier

Collaborations

Prof. Cesare Indiveri, Università degli Studi della Calabria, Department of Biology, Ecology and Earth Sciences (DIBEST), Cosenza (Italy)

Dr. Matteo Gelardi, M.D., Departments of Otolaryngology and Oftalmology, University School of Medicine, Bari (Italy)

Prof. Saverio Cellamare, Università degli Studi di Bari A. Moro, Department of Pharmacy – Drug Sciences, Bari (Italy)

Prof. Cosimo Altomare, Università degli Studi di Bari A. Moro, Department of Pharmacy – Drug Sciences, Bari (Italy)

Prof. Marco Catto, Università degli Studi di Bari A. Moro, Department of Pharmacy - Drug Sciences, Bari (Italy)