

### **Rosario Rizzuto**

Dept. Biomedical Sciences

University of Padua

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Web: <http://www.biomed.unipd.it/people/rizzuto-rosario/>

### **Personal data**

Born in Rome (Italy) on April 15<sup>th</sup>, 1962

Living in Padua, Italy

Married, 3 children

### **Current position**

Professor of General Pathology, University of Padua, Italy

President of the National Center on Gene Therapy and RNA-based therapies

### **Work experience**

2021-date

Cassa di Risparmio di Padova e Rovigo

**Member of the General Council**

2019-2021

Coimbra Group Universities

**Member of the Rectors' Advisory Group**

2019-2021

European University Association (EUA)

**Member of the Research Policy Working Group**

2019-2021

Conference of the Rectors of Italian Universities (CRUI), Rome, Italy

**Member of the steering committee ("Giunta") and delegate for Research and Innovation**

**Coordinator of the Research Committee**

2015-2021

University of Padua, Padua, Italy

**Rector**

2009-2015

Department of Biomedical Sciences  
University of Padua, Padua, Italy

**Head of the Department**

2008-date  
Department of Biomedical Sciences  
University of Padua, Padua, Italy  
**Full Professor of General Pathology**

2007-2008  
School of Pharmacy  
University of Ferrara, Ferrara, Italy  
**Dean of the School**

2002-2008  
School of Pharmacy  
University of Ferrara, Ferrara, Italy  
**Full Professor of General Pathology**

1998-2002  
School of Pharmacy  
University of Ferrara, Ferrara, Italy  
**Associate Professor of General Pathology**

1992-1998  
School of Medicine  
University of Padua, Padua, Italy  
**Assistant Professor of General Pathology**

1991-1992  
Department of Experimental Biomedical Sciences  
University of Padua, Padua, Italy  
**Research Associate**

**Education**

1991  
University of Padua, Padua, Italy  
Department of Experimental Biomedical Sciences  
**Ph.D. in Molecular and Cellular Biology and Pathology**

1987-88  
Columbia University, New York, USA  
H. Merritt Center for the Study of Neuromuscular Disorders  
**2-year research stage**

1986

University of Padua, Padua, Italy  
School of Medicine  
**Medical degree, summa cum laude**

### **Honours, Awards and Memberships**

2020: Commander of the order of merit of the Italian Republic

2014: Antonio Feltrinelli Award of the Accademia dei Lincei

2004: Theodore Bucher Medal

2002: Chiara D'Onofrio Prize

2001: Biotec Award

2017-date: Fellow of the *Istituto Veneto di Scienze, Lettere ed Arti*

2015-date: Fellow of the *Accademia Galileiana di Scienze Lettere ed Arti in Padova*

2013-date: Member of the European Molecular Biology Organization (EMBO)

2008-date: Member of the Academia Europaea, Section of Physiology and Medicine

### **SCIENTIFIC ACTIVITY**

Coordinator of the Research Group on MITOCHONDRIAL CALCIUM SIGNALLING at the Department of Biomedical Sciences, University of Padua (Italy): <https://www.biomed.unipd.it/ricerca/aree-tematiche/mitochondrial-pathophysiology/mitochondrial-calcium-signalling>

### **Research keywords and abstract**

#### **5 keywords**

cell signalling

calcium homeostasis

mitochondria

apoptosis/autophagy

muscle physiology

#### **Abstract**

The research interest of Prof. Rizzuto has been centered on the study of cellular signalling, with special focus on intracellular calcium homeostasis. He pioneered the use of molecularly engineered recombinant luminescent and fluorescent proteins for studying calcium homeostasis at the subcellular level. Subcellular targeting first of the luminescent protein aequorin of *Aequorea victoria*, and then of luciferase of *Photinus pyralis* and of the continuously expanding group of fluorescent probes based on green fluorescent protein (GFP) of *Aequorea victoria*, allowed major advancements in the study of calcium signalling, cellular metabolism and organelle morphology. With this methodological

breakthrough, new biological concepts have been acquired, which include i) the participation of mitochondria in cellular  $\text{Ca}^{2+}$  homeostasis and their role in translating calcium signals in effects as diverse as stimulation of metabolism and induction of cell death, ii) the occurrence and significance of signalling microdomains in the proximity of mitochondria, iii) the identification of the Golgi apparatus as an agonist-sensitive  $\text{Ca}^{2+}$  store, and iv) the major  $\text{Ca}^{2+}$  rises occurring under the plasma membrane upon cell stimulation, to cite a few. In 2011, using a combination of experimental approaches (in silico search, reconstitution in planar lipid bilayers and electrophysiological characterization, expression and silencing in cultured cells, site-specific mutagenesis) he identified the mitochondrial calcium uniporter (MCU), the only fundamental component of the cellular calcium signalling machinery yet to be discovered. This result has opened, with an explosive pace, the molecular era of mitochondrial  $\text{Ca}^{2+}$  homeostasis, that combines the molecular insight into a multi-subunit protein complex, radically different from all other cellular channels, to the possibility of clarifying the role of mitochondria calcium homeostasis in the physiological regulation of tissues and in the pathogenesis of highly prevalent human diseases (neurodegenerative disorders, ischemic heart disease, cancer). Recent data on the role of MCU on muscle trophism and inflammasome control have provided a solid background to the possibility of targeting MCU with traditional and RNA-based drugs for treating pathological conditions such as age- and disease-related sarcopenia and inflammation-based diseases, such as inflammatory bowel diseases, lung fibrosis and atherosclerosis. Finally, with an approach similar to that employed for the discovery of MCU, also the long-sought and debated mitochondrial  $\text{K}_{\text{ATP}}$  channel, proposed to be a primary regulator of ischemic pre-conditioning, has been identified by Prof. Rizzuto's research team in 2019 and shown to control the volume of the mitochondrial matrix and the activity of respiratory complexes.

### Research Projects - Principal Investigator

The research activity of prof. Rizzuto has been supported through the year by grants from the European Research Council (ERC Ideas Advanced grant - MitoCALCIUM: Mitochondrial calcium signalling: molecules, roles and pharmacological targeting), the EU FP programs, the National Institute of Health, the Italian Association for Cancer Research (AIRC), Telethon- Italy, the Italian Education and Health Ministries and the Cariparo and Cariplo bank foundations.

### Bibliometry

**ORCID:** <https://orcid.org/0000-0001-7044-5097>

According to **Google Scholar** (User: OW0gQfUAAAAJ&hl): Total Citations: 71861.  
H-Index: 121.

According to **Scopus** (Author ID 7005289262): Total Citations: 46156.  
H-Index: 105.

According to **WoS** (Researcher ID B-6312-2008): Total number of citations: 43444.  
H-index: 103.

Ranked 42<sup>nd</sup> in Top Italian Scientists (Biomedical Sciences)

[https://www.topitalianscientists.org/TIS\\_HTML/Top\\_Italian\\_Scientists\\_Biomedical\\_Sciences.htm](https://www.topitalianscientists.org/TIS_HTML/Top_Italian_Scientists_Biomedical_Sciences.htm)

## Publications

306 full articles cited in Pubmed (<https://pubmed.ncbi.nlm.nih.gov/>)

<https://scholar.google.com/citations?user=OW0gQfUAAAAJ&hl=it&oi=ao>

### Selected publications

5 publications (last 10 years)

1. S. Feno, F. Munari, D.V. Reane, R. Gissi, D.H. Hoang, A. Castegna, B. Chazaud, A. Viola\*, R. Rizzuto\*, A. Raffaello\* (2021). The dominant negative mitochondrial calcium uniporter subunit MCUb drives macrophage polarization during skeletal muscle regeneration. *Sci. Signal.* 14(707):eabf3838.
2. A. Paggio, V. Checchetto, A. Campo, R. Menabò, G. Di Marco, F. Di Lisa, I. Szabo, R. Rizzuto\*, D. De Stefani\* (2019). Identification of an ATP-sensitive potassium channel in mitochondria. *Nature* 572:609-613.
3. M. Patron, V. Checchetto, A. Raffaello, E. Teardo, D. Vecellio Reane, M. Mantoan, V. Granatiero, I. Szabò, D. De Stefani\* and R. Rizzuto\* (2014) MICU1 and MICU2 finely tune the mitochondrial  $\text{Ca}^{2+}$  uniporter by exerting opposite effects on MCU activity. *Mol. Cell* 53:726-737.
4. A. Raffaello, D. De Stefani, D. Sabbadin, E. Teardo, G. Merli, A. Picard, V. Checchetto, S. Moro, I. Szabò, R. Rizzuto\* (2013) The mitochondrial calcium uniporter is a multimer that can include a dominant-negative pore-forming subunit. *EMBO J.* 32:2362-76.
5. R. Rizzuto\*, D. De Stefani, A. Raffaello, C. Mammucari (2012) Mitochondria as sensors and regulators of calcium signalling. *Nat. Rev. Mol. Cell Biol.* 9, 566-578.

\*corresponding author

5 publications (whole career)

1. D. De Stefani, A. Raffaello, E. Teardo, I. Szabo, R. Rizzuto\* (2011) A forty-kilodalton protein of the inner membrane is the mitochondrial calcium uniporter. *Nature* 476:336-340.
2. P. Pinton, A. Rimessi, S. Marchi, F. Orsini, E. Migliaccio, M. Giorgio, C. Contursi, S. Minucci, F. Mantovani, M. R. Wieckowski, G. Del Sal, P. G. Pelicci, R. Rizzuto\* (2007) Protein kinase C beta and prolyl isomerase 1 regulate mitochondrial effects of the life-span determinant p66Shc. *Science* 315:659-663.
3. R. Rizzuto\*, P. Pinton, W. Carrington, F. S. Fay, K. E. Fogarty, L. M. Lifshitz, R. A. Tuft, T. Pozzan\* (1998) Close contacts with the endoplasmic reticulum as determinants of mitochondrial  $\text{Ca}^{2+}$  responses. *Science* 280:1763-1766.
4. R. Rizzuto\*, M. Brini, M. Murgia, T. Pozzan\* (1993) Microdomains with high  $\text{Ca}^{2+}$  close to  $\text{IP}_3$ -sensitive channels that are sensed by neighboring mitochondria. *Science* 262:744-747.
5. R. Rizzuto\*, A. W. Simpson, M. Brini, T. Pozzan\* (1992) Rapid changes of mitochondrial  $\text{Ca}^{2+}$  revealed by specifically targeted recombinant aequorin. *Nature* 358:325-327.

\*corresponding author

### **Editorial Activity**

Prof. Rizzuto routinely acts as reviewer for international scientific journals (Nature, Science, Cell, J. Cell Biol., EMBO J., J. Biol. Chem., Trends Cell Biol., etc.) and granting agencies (he is currently member of European Research Council and EMBO review panels).

### **Program Chairs/Organization Committee**

- Gordon Research Conference on Calcium Signalling, Renaissance Tuscany Il Ciocco, Lucca, Italy, June 2013 (Chair: R. Rizzuto; vice-chair: D.I. Yule)
- Gordon Research Conference on Calcium Signalling, Colby Sawyer College- New London- NH, USA, June 2011 (chair. K. Foskett; vice-chair: R. Rizzuto)
- EMBO Workshop on "Calcium signaling and diseases", Capri, Italy, September 2004 (organizers: E. Carafoli, R. Rizzuto)
- 12th International Symposium on Calcium Binding Proteins and Calcium Function in Health and Disease - Cavalese, Italy, February 2002 (organizers: E. Carafoli, R. Rizzuto, T. Pozzan)

### **Keynote speeches and lectures**

Prof. Rizzuto gave >300 lectures at national and international meetings (including >20 plenary lectures) and seminars in Universities and Research Institutes.

A handwritten signature in black ink, appearing to read 'R. Rizzuto'.