

Cecilia Gelfi

Education

- 1980: Faculty of Pharmacy, degree in Pharmaceutical Science and Technologies
- 1980: fellowship at Mario Negri Pharmacological Research Institute, enzymology department
- 1981: fellowship at The Haemophilia and Thrombosis Center, Bianchi Bonomi
- 1982: fellowship from the National Council of Research, faculty of Medicine, Thrombosis and Haemostasis department
- 1983: fellowship from CNR, Faculty of Medicine, Separation Science Department
- 1989: Specialist on Morphological, Chemical and Biological Sciences
- 1990: permanent position at Institute of Advanced Biomedical Technologies
- 1998: head of the department of Molecular Physiology and Biotechnology
- 2001: Bioimages and Molecular Physiology, head of proteomic and separation science unit.

Current Position

Full Professor of Clinical Biochemistry and Molecular Biology at the Faculty of Medicine of the University of Milan

Employment and research experience

- **2006:** Associate Professor of Clinical Biochemistry and Molecular Biology, UNIMI
- **2001:** Coordinator of the Clinical Functional Proteomic research section of IBFM-CNR
- **1990:** Position as scientist at ITBA-CNR, from 1998: Head of the Unit of Molecular Physiology and Biotechnology.
- **1980-1989:** Fellowships at Mario Negri Pharmacological Research Institute, Bianchi Bonomi Foundation, National Research Council.
- **1980:** Degree in Pharmaceutical Science and Technologies, UNIMI

Awards

- **2018:** Member of the open source committee University of Milano
- **2017:** Member of the committee of the “Omics” technological platform for proteomic and metabolomic of the university of Milano
- **2017:** Full Member of the European Federation of Laboratory Medicine Working Group on test evaluation
- **2012:** Member of the Board of Directors of the Foundation Toscana Gabriele Monasterio for Medical Research and Public Health, based in Pisa
- **2009:** Coordinator of the Italian Human Proteome Net
- **2006:** Vice president of the Italian Proteome Association (ItPA)

Research activity

The scientific activity of Cecilia Gelfi has been focused on:

- the development of methodologies for the separation and identification of biomolecules. She contributed to the development of electrophoretic techniques transforming them into a physico-chemical tool for probing the structure and conformation of macromolecules;
- the investigation of functional and morphological changes occurring in muscle in a number of physiological and pathological states. Since 1999 has contributed to the muscle proteome signature under physiological, parapathological and pathological conditions generating a database of differential changes in muscle proteomes both in humans and in animal models. These studies allowed: i) to define the interindividual variability of proteins expressed in specific muscles, ii) to identify proteomics profiles of different muscles (e.g. vastus lateralis, deltoideus, soleus, tibialis) associating them to muscle phenotypes; iii) to identify for the first time the relationship among structural changes of the extracellular composition (ECM) and muscle atrophy identifying specific changes associated to homeostasis disruption of the ECM structure and cellular regeneration, iv) identification of the role of ECM protein dysregulation in inflammatory bowel disease, v) identification of biomarkers associated to aging process.
- the development of i) clinical methods for the monitoring of drugs; ii) protocols for serological profiles for the clinical diagnosis of pancreatic and prostate cancer; iii) MALDI imaging methodologies for monitoring nano particles; iv) HPTLC-MALDI methodologies for recognition and quantitation of gangliosides and globosides in tissue extracts.
- The matrisome proteome composition of tendon.

Achieved grants

- - ASI project approved by European Space Agency: Studio della protezione da stress ossidativo del muscolo scheletrico di soggetti sottoposti ad un protocollo di immobilizzazione forzata (bed rest) trattati con un cocktail di antiossidanti nella dieta. ole: principal investigator.
- - Cariplò 2017 Ricerca malattie invecchiamento (approved): Increased sarcopenia in a cohort of elderly carrying SNAP25 polymorphisms: mechanistic insights from the SNAP25
- Prin 2015FBNB5Y (2015): Targeting autophagy in myogenic cells to counteract muscle aging and neuromuscular diseases: novel bioengineering tools for validating pharmacological candidates.
- Ministero della salute GR-2010-2317029 (2012-2015): Integrated "OMIC" approach to explore molecular pathogenesis and clinical heterogeneity in facio-scapulo-humeral muscular dystrophy. Role: partner.
- Progetto Regione Lombardia "Dote Ricerca Applicata" (2012-2013). Role: Principal investigator. (two year fellowship:
- Istituto Italiano di Tecnologia (IIT) Project Seed (2010-2013): "Isoelectric focusing chip-array for protein profiling of biological samples with MALDI detection". Role: partner.
- Banca San Paolo di Torino, Programma Neuroscienze (2010-2012): "Moving again! New approaches arising from molecular neuroscience for the treatment of movement impairment ". Role: partner.
- Sigma Tau 2010: Differential proteomics analysis. Role: Principal investigator.
- UE VII framework program "BIO-NMD" (2009-2012): Identifying and validating pre-clinical biomarkers for diagnostics and therapeutics of Neuromuscular Disorders. Role: partner.
- Telethon GGP11082 (2010-2012): "Toward a mitochondrial therapy of collagen VI muscular dystrophies". Role: partner.
- Telethon GGP04113 (2008-2010): "Toward a mitochondrial therapy of collagen VI muscular dystrophies". Role: partner.
- FIRB RBRN07BMCT (2007-2012): Rete Nazionale per lo studio della Proteomica Umana (Italian Human ProteomeNet). Role: Principal investigator.
- WADA-World Antidoping Agency (2007-2009) WAD/06/021: Molecular signatures of IGF-1 gene doping after AAV-mediated gene transfer in animal models. Role: partner
- ASI 1/007/06/0ASI Osteoporosi ed Atrofia Muscolare (OSMA) (2006-2009): Muscular atrophy investigated by proteomic approach. Role: partner.
- ESA (2006-2008) BBR 2-2: Vibration exercise in space – Optimization of muscle training Bed rest study. Role: subcontractor.

- Ministero della salute (2005-2006): Doping genetico e tessuto muscolare. Ricerca di indicatori o marker tissutali e/o serici per evidenziare interventi genetici sul tessuto muscolare. Role: partner.
- Telethon GGP02271 (2004-2005): "Pathophysiology of Facioscapulohumeral Muscular Dystrophy: transcription from the 4q35 locus, gene expression profiling and proteome analysis in muscles derived from normal and FHSD patients". Role: partner.
- ASI 1/R/295/02 (2004): Deterioration of human skeletal muscle in microgravity: structural, contractile, energetic and proteomic features. Role: partner.
- Cariplò 2002.2267 (2004-2005): Studio proteomico del muscolo scheletrico nell'uomo: regolazione della contrattilità e del metabolismo nell'età evolutiva ed in alcune miopatie. Role: Principal investigator.
- FIRB RBAU01JS5C (2003-2005): Nuove materie prime da semi di leguminose per la preparazione di alimenti funzionali per soggetti ipercolestemici. Role: partner.

Editorial Activity

Official H Index: 43

source: Scopus

ORCID research ID: 0000000229966912

Author of 250 papers published on peer review scientific journals.

2017

Capitanio D, Moriggi M, De Palma S, Bizzotto D, Molon S, Torretta E, Fania C, Bonaldo P, Gelfi C, Braghetta P.

Collagen VI Null Mice as a Model for Early Onset Muscle Decline in Aging. *Front Mol Neurosci.* 2017 Oct 24;10:337. doi: 10.3389/fnmol.2017.00337. eCollection 2017. PMID: 29114203 Free PMC Article

Moriggi M, Pastorelli L, Torretta E, Tontini GE, Capitanio D, Bogetto SF, Vecchi M, Gelfi C. Contribution of Extracellular Matrix and Signal Mechanotransduction to Epithelial Cell Damage in Inflammatory Bowel Disease Patients: A Proteomic Study. *Proteomics.* 2017 Oct 13. doi: 10.1002/pmic.201700164. [Epub ahead of print] PMID: 29027377

Gelfi C, Capitanio D. DIGE Analysis of Human Tissues. *Methods Mol Biol.* 2018;1664:117-136. doi: 10.1007/978-1-4939-7268-5_11. PMID: 29019130

Al-Daghri NM, Torretta E, Capitanio D, Fania C, Guerini FR, Sabico SB, Clerici M, Gelfi C. Intermediate and low abundant protein analysis of vitamin D deficient obese and non-obese subjects by MALDI-profiling.

Sci Rep. 2017 Oct 3;7(1):12633. doi: 10.1038/s41598-017-13020-z. PMID: 28974732 Free PMC Article

Capitanio D, Fania C, Torretta E, Viganò A, Moriggi M, Bravatà V, Caretti A, Levett DZH, Grocott MPW, Samaja M, Cerretelli P, Gelfi C. TCA cycle rewiring fosters metabolic adaptation to oxygen restriction in skeletal muscle from rodents and humans. *Sci Rep.* 2017 Aug 29;7(1):9723. doi: 10.1038/s41598-017-10097-4. PMID: 28852047 Free PMC Article

Capitanio D, Moriggi M, Gelfi C. Mapping the human skeletal muscle proteome: progress and potential.

Expert Rev Proteomics. 2017 Sep;14(9):825-839. doi: 10.1080/14789450.2017.1364996. Epub 2017 Aug 14.

PMID: 28780899

Fania C, Arosio B, Capitanio D, Torretta E, Gussago C, Ferri E, Mari D, Gelfi C. Protein signature in cerebrospinal fluid and serum of Alzheimer's disease patients: The case of apolipoprotein A-1 proteoforms.

PLoS One. 2017 Jun 19;12(6):e0179280. doi: 10.1371/journal.pone.0179280. eCollection 2017.

PMID: 28628634 Free PMC Article

Caretti A, Vasso M, Bonezzi FT, Gallina A, Trinchera M, Rossi A, Adami R, Casas J, Falleni M, Tosi D, Bragonzi A, Ghidoni R, Gelfi C, Signorelli P. Myriocin treatment of CF lung infection and inflammation: complex analyses for enigmatic lipids. Naunyn Schmiedebergs Arch Pharmacol. 2017 Aug;390(8):775-790. doi: 10.1007/s00210-017-1373-4. Epub 2017 Apr 24. PMID: 28439630

2016

Torretta E, Fania C, Vasso M, Gelfi C. HPTLC-MALDI MS for (glyco)sphingolipid multiplexing in tissues and blood: A promising strategy for biomarker discovery and clinical applications.

Electrophoresis. 2016 Jul;37(14):2036-49. doi: 10.1002/elps.201600094. Epub 2016 Jun 1.

PMID: 27162164

Cirillo F, Ghiraldi A, Fania C, Piccoli M, Torretta E, Tettamanti G, Gelfi C, Anastasia L. NEU3 Sialidase Protein Interactors in the Plasma Membrane and in the Endosomes. J Biol Chem. 2016 May 13;291(20):10615-24. doi: 10.1074/jbc.M116.719518. Epub 2016 Mar 17. PMID: 26987901 Free PMC Article

Palorini R, Votta G, Pirola Y, De Vitto H, De Palma S, Airolidi C, Vasso M, Ricciardiello F, Lombardi PP, Cirulli C, Rizzi R, Nicotra F, Hiller K, Gelfi C, Alberghina L, Chiaradonna F. Protein Kinase A Activation Promotes Cancer Cell Resistance to Glucose Starvation and Anoikis. PLoS Genet. 2016 Mar 15;12(3):e1005931. doi: 10.1371/journal.pgen.1005931. eCollection 2016 Mar. PMID: 26978032 Free PMC Article

Scotton C, Bovolenta M, Schwartz E, Falzarano MS, Martoni E, Passarelli C, Armaroli A, Osman H, Rodolico C, Messina S, Pegoraro E, D'Amico A, Bertini E, Gualandi F, Neri M, Selvatici R, Boffi P, Maioli MA, Lochmüller H, Straub V, Bushby K, Castrignanò T, Pesole G, Sabatelli P, Merlini L, Braghetta P, Bonaldo P, Bernardi P, Foley R, Cirak S, Zaharieva I, Muntoni F, Capitanio D, Gelfi C, Kotelnikova E, Yuryev A, Lebowitz M, Zhang X, Hodge BA, Esser KA, Ferlini A. Deep RNA profiling identified CLOCK and molecular clock genes as pathophysiological signatures in collagen VI myopathy. J Cell Sci. 2016 Apr 15;129(8):1671-84. doi: 10.1242/jcs.175927. Epub 2016 Mar 4. PMID: 26945058 Free PMC Article

Capitanio D, Vasso M, De Palma S, Fania C, Torretta E, Cammarata FP, Magnaghi V, Procacci P, Gelfi C. Specific protein changes contribute to the differential muscle mass loss during ageing. Proteomics. 2016 Feb;16(4):645-56. doi: 10.1002/pmic.201500395. PMID: 26698593

2015

Vibration mechanosignals superimposed to resistive exercise result in baseline skeletal muscle transcriptome profiles following chronic disuse in bed rest. Salanova M, Gambara G, Moriggi M, Vasso M, Ungethuem U, Belavý DL, Felsenberg D, Cerretelli P, Gelfi C, Blottner D. *Sci Rep.* 2015 Nov 24;5:17027. doi: 10.1038/srep17027. PMID: 26596638 Free PMC Article

Di Blasi C, Sansanelli S, Ruggieri A, Moriggi M, Vasso M, D'Adamo AP, Blasevich F, Zanotti S, Paolini C, Protasi F, Tezzon F, Gelfi C, Morandi L, Pessia M, Mora M. (2015). A CASQ1 founder mutation in three Italian families with protein aggregate myopathy and hyperCKaemia. *J MED GENET.* Sep;52(9):617-26. doi: 10.1136/jmedgenet-2014-102882.

Leone R, Giussani P, De Palma S, Fania C, Capitanio D, Vasso M, Brioschi L, Riboni L, Viani P, Gelfi C. (2015). Proteomic analysis of human glioblastoma cell lines differently resistant to a nitric oxide releasing agent. *MOL BIOSYST.* Jun;11(6):1612-21. doi: 10.1039/c4mb00725e.

Levett DZ, Viganò A, Capitanio D, Vasso M, De Palma S, Moriggi M, Martin DS, Murray AJ, Cerretelli P, Grocott MP, Gelfi C. (2015). Changes in muscle proteomics in the course of the Caudwell Research Expedition to Mt. Everest. *Proteomics.* Jan;15(1):160-71. doi: 10.1002/pmic.201400306.

Fania C, Sogno I, Vasso M, Torretta E, Leone R, Bruno A, Consonni P, Albini A, Gelfi C. (2015). A PSA-guided approach for a better diagnosis of prostatic adenocarcinoma based on MALDI profiling and peptide identification. *Clin Chim Acta.* Jan 15;439:42-9. doi: 10.1016/j.cca.2014.10.003.

2014

De Palma S, Capitanio D, Vasso M, Braghetta P, Scotton C, Bonaldo P, Lochmüller H, Muntoni F, Ferlini A, Gelfi C. (2014). Muscle proteomics reveals novel insights into the pathophysiological mechanisms of collagen VI myopathies. *J PROTEOME RES.* Nov 7;13(11):5022-30. doi: 10.1021/pr500675e.

Salanova M, Gelfi C, Moriggi M, Vasso M, Viganò A, Minafra L, Bonifacio G, Schiff G, Gutsmann M, Felsenberg D, Cerretelli P, Blottner D. (2014). Disuse deterioration of human skeletal muscle challenged by resistive exercise superimposed with vibration: evidence from structural and proteomic analysis. *FASEB J.* Nov;28(11):4748-63. doi: 10.1096/fj.14-252825.

Zuccotti P, Colombrita C, Moncini S, Barbieri A, Lunghi M, Gelfi C, De Palma S, Nicolin A, Ratti A, Venturin M, Riva P. (2014). hnRNPA2/B1 and nELAV proteins bind to a specific U-rich element in CDK5R1 3'-UTR and oppositely regulate its expression. *BIOCHIM BIOPHYS ACTA.* Jun;1839(6):506-16. doi: 10.1016/j.bbagr.2014.04.018.

Bravatà V, Minafra L, Callari G, Gelfi C, Edoardo Grimaldi LM. (2014). Analysis of thiamine transporter genes in sporadic beriberi. *Nutrition.* Apr;30(4):485-8. doi: 10.1016/j.nut.2013.10.008.

Bergante S, Torretta E, Creo P, Sessarego N, Papini N, Piccoli M, Fania C, Cirillo F, Conforti E, Ghiraldi A, Tringali C, Venerando B, Ibatici A, Gelfi C, Tettamanti G, Anastasia L. (2014). Gangliosides as a potential new class of stem cell markers: the case of GD1a in human bone marrow mesenchymal stem cells. *J LIPID RES.* Mar;55(3):549-60. doi: 10.1194/jlr.M046672.

Torretta E, Vasso M, Fania C, Capitanio D, Bergante S, Piccoli M, Tettamanti G, Anastasia L, Gelfi C. (2014). Application of direct HPTLC-MALDI for the qualitative and quantitative profiling of neutral and acidic glycosphingolipids: the case of NEU3 overexpressing C2C12 murine myoblasts. *ELECTROPHORESIS.* May;35(9):1319-28. doi: 10.1002/elps.201300474.

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Torretta E, Vasso M, Fania C, Capitanio D, Bergante S, Piccoli M, Tettamanti G, Anastasia L, Gelfi C. Application of direct HPTLC-MALDI for the qualitative and quantitative profiling of neutral and acidic glycosphingolipids: The case of NEU3 overexpressing C2C12 murine myoblasts. *Electrophoresis*. 2013 Dec 23.

Urbani A, De Canio M, Palmieri F, Sechi S, Bini L, Castagnola M, Fasano M, Modesti A, Roncada P, Timperio AM, Bonizzi L, Brunori M, Cutruzzolà F, De Pinto V, Di Ilio C, Federici G, Folli F, Foti S, Gelfi C, Lauro D, Lucacchini A, Magni F, Messana I, Pandolfi PP, Papa S, Pucci P, Sacchetta P; Italian Mt-Hpp Study Group-Italian Proteomics Association (www.itpa.it). The mitochondrial Italian Human Proteome Project initiative (mt-HPP). *Mol Biosyst*. 2013 Aug;9(8):1984-92.

Bravatà V, Stefano A, Cammarata FP, Minafra L, Russo G, Nicolosi S, Pulizzi S, Gelfi C, Gilardi MC, Messa C. Genotyping analysis and ¹⁸FDG uptake in breast cancer patients: a preliminary research. *J Exp Clin Cancer Res*. 2013 Apr 30

De Palma S, Leone R, Grumati P, Vasso M, Polishchuk R, Capitanio D, Braghetta P, Bernardi P, Bonaldo P, Gelfi C. Changes in muscle cell metabolism and mechanotransduction are associated with myopathic phenotype in a mouse model of collagen VI deficiency. *PLoS One*. 2013;8(2):e5671

Scaringi R, Piccoli M, Papini N, Cirillo F, Conforti E, Bergante S, Tringali C, Garatti A, Gelfi C, Venerando B, Menicanti L, Tettamanti G, Anastasia L. NEU3 sialidase is activated under hypoxia and protects skeletal muscle cells from apoptosis through the activation of the epidermal growth factor receptor signaling pathway and the hypoxia-inducible factor (HIF)-1α. *J Biol Chem*. 2013 Feb 1;288(5):3153-62.

2012

Alberghina L, Gaglio D, Gelfi C, Moresco RM, Mauri G, Bertolazzi P, Messa C, Gilardi MC, Chiaradonna F, Vanoni M. Cancer cell growth and survival as a system-level property sustained by enhanced glycolysis and mitochondrial metabolic remodeling. *Front Physiol*. 2012 Sep 12;3:362

Minafra L, Norata R, Bravatà V, Viola M, Lupo C, Gelfi C, Messa C. Unmasking epithelial-mesenchymal transition in a breast cancer primary culture: a study report. *BMC Res Notes*. 2012 Jul 3;5:343

Bagnato S, Minafra L, Bravata V, Boccagni C, Sant'Angelo A, Castiglione A, Andriolo M, Lucca LF, De Tanti A, Pistarini C, Formisano R, Dolce G, Gelfi C, Galardi G (2012). Brain-derived neurotrophic factor (Val66Met) polymorphism does not influence recovery from a post-traumatic vegetative state: A blinded retrospective multi-centric study . *JOURNAL OF NEUROTRAUMA*, vol. 29, p. 2050-2059, ISSN: 0897-7151

Macedo A, Moriggi M, Vasso M, De Palma S, Sturnega M, Friso G, Gelfi C, Giacca M, Zacchigna S (2012). Enhanced Athletic Performance on Multisite AAV-IGF1 Gene Transfer Coincides with Massive Modification of the Muscle Proteome. *HUMAN GENE THERAPY*, vol. 23, p. 146-157, ISSN: 1043-0342, doi: 10.1089/hum.2011.157

D. Capitanio, M. Vasso, A. Ratti, G. Grignaschi, M. Volta, M. Moriggi, C. Daleno, C. Bendotti, V. Silani, C. Gelfi (2012). Molecular Signatures of Amyotrophic Lateral Sclerosis Disease Progression in Hind and Forelimb Muscles of an SOD1 (G93A) Mouse Model. *ANTIOXIDANTS & REDOX SIGNALING*, vol. 17, p. 1333-1350, ISSN: 1523-0864, doi: 10.1089/ars.2012.4524

M, Piccoli, G. Palazzolo, E. Conforti, G. Lamorte, N. Papini, P. Creo, C. Fania, R. Scaringi, S. Bergante, C. Tringali, L. Roncoroni, S. Mazzoleni, L. Doneda, R. Galli, B. Venerando, G.

Tettamanti, C. Gelfi, L. Anastasia (2012). The synthetic purine reversine selectively induces cell death of cancer cells. JOURNAL OF CELLULAR BIOCHEMISTRY, ISSN: 0730-2312, doi: 10.1002/jcb.24197

Minafra L, Norata R, Bravata V, Viola M, Lupo C, Gelfi C, Messa C (2012). Unmasking epithelial-mesenchymal transition in a breast cancer primary culture: A study report . BMC RESEARCH NOTES, ISSN: 1756-0500

2011

C. Gelfi, M. Vasso, P. Cerretelli (2011). Diversity of human skeletal muscle in health and disease: contribution of proteomics. JOURNAL OF PROTEOMICS, vol. 74, p. 774-795, ISSN: 1874-3919, doi: 10.1016/j.jprot.2011.02.028

P. Cerretelli, C. Gelfi (2011). Energy metabolism in hypoxia: reinterpreting some features of muscle physiology on molecular grounds. EUROPEAN JOURNAL OF APPLIED PHYSIOLOGY, vol. 111, p. 421-432, ISSN: 1439-6319, doi: 10.1007/s00421-010-1399-5

Moriggi M, Vasso M, Fania C, Capitanio D, Bonifacio G, Salanova M, Blottner D, Rittweger J, Felsenberg D, Cerretelli P, Gelfi C (2011). Long term bed rest with and without vibration exercise countermeasures: Effects on human muscle protein dysregulation. PROTEOMICS. CLINICAL APPLICATIONS, vol. 5, p. 198, ISSN: 1862-8346, doi: 10.1002/pmic.200900817

A. Viganò, M. Vasso, A. Caretti, V. Bravatà, L. Terraneo, C. Fania, D. Capitanio, M. Samaja, C. Gelfi (2011). Protein modulation in mouse heart under acute and chronic hypoxia. PROTEOMICS, vol. 11, p. 4202-4217, ISSN: 1615-9853, doi: 10.1002/pmic.201000804

C. Fania, M. Vasso, E. Torretta, P. Robach, G. Cairo, C. Lundby, C. Gelfi (2011). Setup for human sera MALDI profiling: The case of rhEPO treatment. ELECTROPHORESIS, vol. 32, p. 1715-1727, ISSN: 0173-0835, doi: 10.1002/elps.201100134

2010

M. Moriggi, M. Vasso, C. Fania, D. Capitanio, G. Bonifacio, M. Salanova, D. Blottner, J. Rittweger, D. Felsenberg, P. Cerretelli, C. Gelfi (2010). Long term bed rest with and without vibration exercise countermeasures: effects on human muscle protein dysregulation. PROTEOMICS, vol. 10, p. 3756-3774, ISSN: 1615-9853, doi: 10.1002/pmic.200900817

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2009

P. Robach, S. Recalcati, D. Girelli, C. Gelfi, N.J. Aachmann-Andersen, J.J. Thomsen, A.M. Norgaard, A. Alberghini, N. Campostrini, A. Castagna, A. Viganò, P. Santambrogio, T. Kempf, K.C. Wollert, S. Moutereau, C. Lundby, G. Cairo (2009). Alterations of systemic and muscle iron metabolism in human subjects treated with low-dose recombinant erythropoietin. BLOOD, vol. 113, p. 6707-6715, ISSN: 0006-4971

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Editorial activity

Cecilia Gelfi published **280** scientific articles on international peer-reviewed journals (**Scopus h-index 2016: 42**).

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