

## Laura SILVESTRI-Biographical Sketch



NAME Laura SILVESTRI		POSITION TITLE Tenured <i>Ospedale San Raffaele</i> researcher	
EDUCATION/TRAINING (Begin with baccalaureate or other initial professional education, such as nursing, and include postdoctoral training).			
INSTITUTION AND LOCATION	DEGREE (IF APPLICABLE)	YEAR(S)	FIELD OF STUDY
Università degli Studi di Milano, Italy	M.Sc.	1996	Pharmaceutical Chemistry and Technology
Università degli Studi di Milano, Italy	PhD	2000	Biotechnology
Università di Ferrara, Ferrara, Italy	Master	2005	Science communication
San Raffaele Scientific Institute (OSR), Milan, Italy	Postdoctoral training	2000-2005	Molecular mechanisms in neurodegenerative disorders

RESEARCH AND PROFESSIONAL EXPERIENCE: Concluding with present position, list in chronological order, previous employment, experience, and honors. Include present membership on any Federal Government public advisory committee. List in chronological order the titles, all authors, and complete references to all publications during the past 3 years and to representative earlier publications pertinent to this application. If the list of publications in the last 3 years exceeds 2 pages, select the most pertinent publications. PAGE LIMITATIONS APPLY. DO NOT EXCEED 5 PAGES FOR THE ENTIRE BIOGRAPHICAL SKETCH PER INDIVIDUAL.

**Positions and Employment**

- 1996-2000** Fellow, Division of Biochemistry and Genetic, Carlo Besta Neurological Institute, and Human Molecular Genetic Unit, San Raffaele Scientific Institute, OSR, Milan, Italy
- 2000-2005** Junior Post-Doc, Human Molecular Genetic Unit, San Raffaele Scientific Institute, OSR, Milan, Italy
- 2005-2010** Senior Post-Doc, Regulation of Iron Metabolism Unit, San Raffaele Scientific Institute, OSR, Milan, Italy
- 2010-2011** Visiting Professor (short stages). Department of Pathology, School of Medicine, University of Utah, Salt Lake City, UT, USA
- 2010-2015** Research Associate, Regulation of Iron Metabolism Unit, San Raffaele Scientific Institute, OSR, Milan, Italy
- 2016-present:** Project Leader, San Raffaele Scientific Institute, OSR, Milan, Italy.
- 2007-present:** Adjunct Professor in Molecular Biology, Vita-Salute University, Milan, Italy

## **Honors**

- 2008** Workshop on Iron Overload: Mechanisms, Measurement and Management. National Institute of Diabetes and Digestive and Kidney Diseases (NIDDK). Annapolis, MD, USA. October 27-28, 2008. Travel award
- 2009** XV Telethon Convention. Palazzo dei Congressi. Riva del Garda, Italy. March 9-11, 2009. Best poster award
- 2009** Heme Oxygenases in biology and medicine. 6th International Congress. Miami, FL, USA. September 30-October 4, 2009. Travel award.
- 2013** IBIS International BioIron Society, London, UK. April 16-18, 2013. IBIS Gunshin-Levy award for dedication, commitment and contributions to iron homeostasis
- 2014-2020** Italian Ministry of University and Research: “Abilitazione Scientifica Nazionale” as Professor of Pathology and Molecular and Applied Biology

## **Other Experience and Professional Memberships**

- 2007-** Reviewer for Blood, Journal of Clinical Investigation, Haematologica, American Journal of Hematology, Scientific Reports, PlosOne, Biochemie, American Journal of Physiology-Cell Physiology, Molecular Genetics and Metabolism, Experimental Hematology, Science Reports, Journal of Hepatology
- 2016** Reviewer for the Grant Agency Kidney Research UK
- 2020** Reviewer for the Polish Ministry of Science and Higher Education
- 2012** Abstract Reviewer, “The 54th American Society of Hematology (ASH) Annual Meeting and Exposition”, Atlanta, GA.
- 2014** Abstract reviewer and member of the scientific committee, European Iron Club Meeting, Verona, Italy.
- 2017** Abstract Reviewer, “The 59th American Society of Hematology (ASH) Annual Meeting and Exposition”, Atlanta, GA.
- 2018** Abstract reviewer, European Iron Club Meeting, Zurich, Switzerland
- 2019** Abstract reviewer, BioIron Meeting, Heidelberg, Germany
- 2019-2020** Abstract reviewer, EHA Meeting, Amsterdam, The Netherlands
- 2009-** Member, International BioIron Society (IBIS)
- 2014-** Member, American Society of Hematology (ASH)
- 2018-** Member, European Hematology Association (EHA)
- 2015-2019** Member of the Board of Directors, International BioIron Society (IBIS)
- 2018-2019** Member of the Scientific Program Committee-Advisory Board, European Hematology Association (EHA)
- 2019-2023** Member of the Scientific Committee on Iron and Heme, American Society of Hematology (ASH)

## **PUBLICATIONS**

### **Last 5 years publications**

#### **Original articles**

1. Nai A, Lorè NI, Pagani A, De Lorenzo R, Di Modica S, Saliu F, Cirillo DM, Rovere-Querini P,

- Manfredi AA, **Silvestri L.** Hpcidin levels predict Covid-19 severity and mortality in a cohort of hospitalized Italian patients. *Am J Hematol.* 2021 Jan;96(1):E32-E35. doi: 10.1002/ajh.26027.
2. Nai A, Lidonnici MR, Federico G, Pettinato M, Olivari V, Carrillo F, Geninatti C, Ferrari G, Camaschella C, **Silvestri L\***, Carlomagno F\*. NCOA4-mediated ferritinophagy in macrophages is crucial to sustain erythropoiesis in mice. *Haematologica.* 2020 Feb 27. doi: 10.3324/haematol.2019.241232. \*: co-last and co-corresponding authors
3. Pagani A., Pettinato M., Colucci S., Dulja A., Rauner M., Nai A., Camaschella C., Altamura S., Muckenthaler M., **Silvestri L.**, Hemochromatosis proteins are dispensable for the acute hepcidin response to BMP2. *Haematologica.* 2020. Oct 1;105(10):e493. DOI: 10.3324/haematol.2019.241984.
4. Nai A., Pettinato M. Federico G., Carlomagno F. and Silvestri L. Tamoxifen erythroid toxicity revealed by studying the role of Nuclear Receptor Co-Activator 4 in erythropoiesis. *Haematologica.* (2019). *Haematologica.* 2019 Aug;104(8):e383-e384. doi: 10.3324/haematol.2019.224857.
5. Crippa S, Rossella V, Aprile A, Silvestri L, Rivis S, Scaramuzza S, Pirroni S, Avanzini MA, Basso-Ricci L, Hernandez RJ, Zecca M, Markt S, Ciceri F, Aiuti A, Ferrari G, Bernardo ME. Bone marrow stromal cells from  $\beta$ -thalassemia patients have impaired hematopoietic supportive capacity. *J Clin Invest.* 2019 Feb 25;129(4):1566-1580. doi: 10.1172/JCI123191.
6. Artuso I, Pettinato M, Nai A, Pagani A, Sardo U, Billoré B, Lidonnici MR, Bennett C, Mandelli G, Pasricha SR, Ferrari G, Camaschella C, Kautz L, Silvestri L. Transient decrease of serum iron after acute erythropoietin treatment contributes to hepcidin inhibition by ERFE in mice. *Haematologica.* 2018 Sep 28. pii: haematol.2018.199810. doi: 10.3324/haematol.2018.199810.
7. Artuso I, Lidonnici MR, Altamura S, Mandelli G, Pettinato M, Muckenthaler MU, **Silvestri L**, Ferrari G, Camaschella C, Nai A. Transferrin receptor 2 is a potential novel therapeutic target for  $\beta$ -thalassemia: evidence from a murine model. *Blood.* 2018 Nov 22;132(21):2286-2297. doi: 10.1182/blood-2018-05-852277.
8. Ravasi G, Pelucchi S, Mariani R, **Silvestri L**, Camaschella C, Piperno A. A severe hemojuvelin mutation leading to late onset of HFE2-hemochromatosis. *Dig Liver Dis.* 2018 Apr 27. pii: S1590-8658(18)30715-1. doi: 10.1016/j.dld.2018.04.018.
9. Colucci S., Pagani A., Pettinato M., Artuso I., Nai A., Camaschella C., **Silvestri L.** The immunophilin FKBP12 inhibits hepcidin expression by binding the BMP type I receptor ALK2 in hepatocytes. (2017) *Blood.* 2017 doi: 10.1182/blood-2017-04-780692.
- Highlighted in Blood.** Parrow NL, Fleming RE. Releasing the FKBP12 brake on hepcidin. *Blood.* 2017 Nov 9;130(19):2049-2050. doi: 10.1182/blood-2017-09-805390.
10. Pagani, A., Colucci, S., Bocciardi, R., Bertamino, M., Dufour, C., Ravazzolo, R., **Silvestri, L.\***, Camaschella, C\*. A new form of IRIDA due to combined heterozygous mutations of TMPRSS6 and ACVR1A encoding the BMP receptor ALK2 (2017) *Blood*, 129 (25), pp. 3392-3395. DOI: 10.1182/blood-2017-03-773481 \*: co-last authors
11. Latour, C., Besson-Fournier, C., Meynard, D., **Silvestri, L.**, Gourbeyre, O., Aguilar-Martinez, P., Schmidt, P.J., Fleming, M.D., Roth, M.-P., Coppin, H. Differing impact of the deletion of hemochromatosis-associated molecules HFE and transferrin receptor-2 on the iron phenotype of mice lacking bone morphogenetic protein 6 or hemojuvelin (2016) *Hepatology*, 63 (1), pp. 126-137. DOI: 10.1002/hep.28254
12. Nai, A., Rubio, A., Campanella, A., Gourbeyre, O., Artuso, I., Bordini, J., Gineste, A., Latour, C., Besson-Fournier, C., Lin, H.Y., Coppin, H., Roth, M.-P., Camaschella, C., **Silvestri, L.\***, Meynard, D\*. Limiting hepatic Bmp-Smad signaling by matriptase-2 is required for erythropoietin-mediated hepcidin suppression in mice (2016) *Blood*, 127 (19), pp. 2327-2336. DOI: 10.1182/blood-2015-11-681494

\*: co-last and co-corresponding authors

**Highlighted in Blood.** Bartnikas T. Matriptase-2 links erythropoietin to iron. *Blood* 2016 127:2270-2271; doi: <https://doi.org/10.1182/blood-2016-02-694919>

**13.** Rausa, M., Pagani, A., Nai, A., Campanella, A., Gilberti, M.E., Apostoli, P., Camaschella, C., **Silvestri, L.** Bmp6 expression in murine liver non parenchymal cells: A mechanism to control their high iron exporter activity and protect hepatocytes from iron overload? (2015) *PLoS ONE*, 10 (4), art. no. e0122696, . DOI: 10.1371/journal.pone.0122696

**14.** Nai, A., Lidonnici, M.R., Rausa, M., Mandelli, G., Pagani, A., **Silvestri, L.**, Ferrari, G., Camaschella, C. The second transferrin receptor regulates red blood cell production in mice (2015) *Blood*, 125 (7), pp. 1170-1179. DOI: 10.1182/blood-2014-08-596254

**Highlighted in Blood.** Pantopoulos K. TfR2 links iron metabolism and erythropoiesis. *Blood* 125:1055-6. doi: 10.1182/blood-2014-12-617571.

**15.** Rausa, M., Ghitti, M., Pagani, A., Nai, A., Campanella, A., Musco, G., Camaschella, C., **Silvestri, L.** Identification of TMPRSS6 cleavage sites of hemojuvelin (2015) *Journal of Cellular and Molecular Medicine*, 19 (4), pp. 879-888. DOI: 10.1111/jcmm.12462

**16.** Pagani, A., Vieillevoys, M., Nai, A., Rausa, M., Ladli, M., Lacombe, C., Mayeux, P., Verdier, F., Camaschella, C., **Silvestri, L.** Regulation of cell surface transferrin receptor-2 by iron-dependent cleavage and release of a soluble form (2015) *Haematologica*, 100 (4), pp. 458-465. DOI: 10.3324/haematol.2014.118521

**17.** Ravasi, G., Rausa, M., Pelucchi, S., Arosio, C., Greni, F., Mariani, R., Pelloni, I., **Silvestri, L.**, Pineda, P., Camaschella, C., Piperno, A. Transferrin receptor 2 mutations in patients with juvenile hemochromatosis phenotype (2015) *American Journal of Hematology*, 90 (12), pp. E226-E227. DOI: 10.1002/ajh.24202

#### **Reviews:**

**1.** Mleczko-Sanecka K, **Silvestri L.** Cell-type-specific insights into iron regulatory processes. *Am J Hematol.* 2021 Jan;96(1):110-127. doi: 10.1002/ajh.26001.

**2.** Camaschella C, Nai A, **Silvestri L.** Iron metabolism and iron disorders revisited in the hepcidin era. *Haematologica.* 2020 Jan 31;105(2):260-272. doi: 10.3324/haematol.2019.232124.

**3.** Pagani A, Nai A, **Silvestri L**, Camaschella C. Hepcidin and Anemia: A Tight Relationship. *Front Physiol.* 2019 Oct 9;10:1294. doi: 10.3389/fphys.2019.01294.

**4.** Camaschella, C., Pagani, A., Nai, A., **Silvestri, L.** The mutual control of iron and erythropoiesis (2016) *International Journal of Laboratory Hematology*, 38, pp. 20-26. DOI: 10.1111/ijlh.12505.

#### **Book chapters:**

**1.** **Silvestri L**, Nai A, Dulja A, Pagani A. Hepcidin and the BMP-SMAD pathway: An unexpected liaison. *Vitam Horm.* 2019;110:71-99.

**2.** **Silvestri, L.** Iron Metabolism in Aging. (2016) *Molecular Basis of Nutrition and Aging: A Volume in the Molecular Nutrition Series*, pp. 523-536. DOI: 10.1016/B978-0-12-801816-3.00037-6

#### **Complete List of Published Work in My Bibliography:**

<https://www.scopus.com/authid/detail.uri?authorId=7004424357>

#### **Patents**

"Inhibitor of TFR2 and/or inhibitor of FKBP12 and uses thereof". Inventors: **Silvestri L.**, Camaschella C., Nai A., Pagani A., Colucci S. (2017). US Provisional n. 62/483,17

