Marta Kovatcheva, Ph.D.

Group Leader (Assistant Professor equivalent)

Laboratory of Cell Plasticity & Aging

IFOM, Milan, IT

POSITIONS HELD:

March 2024-present Group Leader (Tenure Track) IFOM, Milan, Italy

EDUCATION & RESEARCH TRAINING:

2017-2024 **Postdoctoral Research Fellow** IRB Barcelona, Barcelona, Spain

Laboratory of Manuel Serrano

2010-2017 **Ph.D.** in Cancer Biology MSKCC, New York, NY, USA

Gerstner Sloan Kettering Graduate

School of Biomedical Sciences
Laboratory of **Andrew Koff**

2006-2010 **Hon. B.Sc.** with High Distinction in Molecular Genetics

University of Toronto, Toronto, ON,

Canada

Undergraduate thesis Laboratory of

Daniel Durocher

PUBLICATIONS:

Preprints:

Miro-Blanch J, Capellades J, Balvay C, Maudet M, <u>Kovatcheva M</u>, Raineri S, Rabot S, Mellor J, Serrano M, and Yanes O. A targeted metabobolomic method to detect epigenetically relevant metabolites. *bioRxiv; submitted to Molecular Metabolism*. doi: 10.1101/2023.11.30.569455. Original research article.

Published Works:

Fissoun C and <u>Kovatcheva M</u>.(2025) Vitamin B12 in cell plasticity and repair. *DNA Cell Biol*, doi: 10.1089/dna.2025.0037. Review article.

Aggarwal SS, Andreani C, Deng Z, Frede J, Kameneva P, **Kovatcheva M**, Micevic G, Nicolson F, Puschhof J, Roberts M, Wang G, Watson DC. (2024) The 2024 Generation. *Nature Cancer*, doi: 10.1038/s43018-024-00866-2. Review article.

Alcon C, <u>Kovatcheva M</u>, Morales-Sanchez P, López-Polo V, Torres T, Puig S, Lu A, Samitier J, Enrich C, Serrano M, and Montero J. (2024) HRK downregulation and augmented BCL-xL binding to BAK confer

apoptotic protection to therapy-induced senescent melanoma cells. *Cell Death and Differentiation,* doi: 10.1038/s41418-024-01417-z. Original research article.

Ramponi V, Richart L, **Kovatcheva M**, Stephan-Otto Attolini C, Capellades J, Lord AE, Yanes O, Ficz G, and Serrano M. (2024) H4K20me3-Mediated Repression of Inflammatory Genes is a Characteristic and Targetable Vulnerability of Persister Cancer Cells. *Cancer Research*, doi: https://doi.org/10.1158/0008-5472.CAN-24-0529. Original research article.

Protasoni M, López-Polo V, Stephan-Otto Attolini C, Brandariz J, Herranz N, Mateo J, Ruiz S, Fernandez-Capetillo O, <u>Kovatcheva M*</u>, and Serrano M*. (2024). Cyclophilin D plays a critical role in the survival of senescent cells. *EMBO J*, doi: 10.1038/s44318-024-00259-2. *co-corresponding authors. Original research article.

López-Polo V, Maus M, Zacharioudakis E, Lafarga M, Stephan-Otto Attolini C, DM Marques F, **Kovatcheva M**, Gavathiotis E, and Serrano M. (2024) Release of mitochondrial dsRNA promotes the inflammatory phenotype of senescent cells. *Nature Communications*, doi: 10.1038/s41467-024-51363-0. Original research article.

<u>Kovatcheva M*,</u> Melendez E, Chondronasiou D, Pietrocola F, Bernad R, Caballe A, Junza A, Capellades J, Holguin-Horcajo A, Prats N, Durand S, Rovira M, Yanes O, Stephan-Otto Attolini C, Kroemer G, and Serrano M*. (2023) Vitamin B12 is a limiting factor for cellular plasticity and tissue repair. *Nature Metabolism* doi: 10.1038/s42255-023-00916-6. *co-corresponding authors. Original research article.

Dominguez-Lopez I*, **Kovatcheva M***, Casas R, Toledo E, Fito M, Ros E, Estruch R, Serrano M, and Lamuela-Raventos RM. (2023) Higher circulating vitamin B12 is associated with lower levels of inflammatory markers in individuals at high cardiovascular risk and in naturally aged mice. *J Sci Food Agric*. doi:10.1002/jsfa.12976. Original research article. *authors contributed equally. Original research article.

Gleason CE, Dickson MA, Klein Dooley ME, Antonescu CR, Gularte-Mérida R, Benitez M, Delgado JI, Kataru RP, Tan MWY, Bradic M, Adamson TE, Seier K, Richards AL, Palafox M, Chan E, D'Angelo SP, Gounder MM, Keohan ML, Kelly CM, Chi P, Movva S, Landa J, Crago AM, Donoghue MTA, Qin LX, Serra V, Turkekul M, Barlas A, Firester DM, Manova-Todorova K, Mehrara BJ, Kovatcheva M, Tan NS, Singer S, Tap WD, Koff A. (2023) Therapy-Induced Senescence Contributes to the Efficacy of Abemaciclib in Patients with Dedifferentiated Liposarcoma. *Clinical Cancer Research*. doi: 10.1158/1078-0432.CCR-23-2378. Original research article.

Marin, I, Boix O, Garcia-Garijo A, Sirois I, Caballe A, Zarzuela E, Ruano I, Stephan-Otto Attolini C, Prats N, Lopez-Dominguez JA, **Kovatcheva M**, Garralda E, Munoz J, Caron E, Abad M, Gros A, Pietrocola F, and Serrano M. (2023) Cellular senescence is immunogenic and promotes anti-tumor immunity. *Cancer Discovery*. doi: 10.1158/2159-8290.CD-22-0523 Original research article.

Chondronasiou D, Martinez J, Melendez E, Lynch C, del Pozo N, **Kovatcheva M**, Aguilera M, Prats N, Real FX, and Manuel Serrano. (2022) Deciphering the roadmap of in vivo reprogramming toward pluripotency. *Stem Cell Reports*, 17: 2501-2517. doi: 10.1016/j.stemcr.2022.09.009 Original research article.

Meyer K, López-Domínguez JA, Maus M, <u>Kovatcheva M,</u> and Serrano M. (2022) Senescence as a therapeutic target: current state and future challenges Cellular Senescence in Disease. *Elsevier*. pp.425-442. ISBN 978-0-12-822514-1. <u>Book chapter</u>

Klein ME, **Kovatcheva M**, Davies LE, Tap WD, and Koff A. (2018). CDK4/6 inhibitors: The mechanism of action may not be as simple as once thought. *Cancer Cell*, 34(1):9-20. doi: 10.1016/j.ccell.2018.03.023 Review article.

Kovatcheva M, Klein ME, Tap WD, and Koff A. (2017). Mechanistic understanding of the role of ATRX in senescence provides new insight for combinatorial therapies with CDK4 inhibitors. *Molecular and Cellular Oncology*, 5(1):e1384882. doi: 10.1080/23723556.2017.1384882 Review article.

<u>Kovatcheva M</u>, Liao W, Klein ME, Robine, N, Geiger H, Crago AM, Dickson MA, Tap WD, Singer S, and Koff A. (2017). ATRX is a regulator of therapy induced senescence in human cells. *Nature Communications*, 8(1):386. doi: 10.1038/s41467-017-00540-5. Original research article.

<u>Kovatcheva M.</u>, Liu DD, Dickson MA, Klein ME, O'Connor R, Wilder FO, Socci ND, Tap WD, Schwartz GK, Singer S, Crago AM, and Koff A. (2015). MDM2 turnover and expression of ATRX determine the choice between quiescence and senescence in response to CDK4 inhibition. *Oncotarget*, 6(10): 8226-43. doi: 10.18632/oncotarget.3364 Original research article.

FUNDING:

Grants obtained as PI:

2025-2027	European Association for Cancer Research, Rising Star Award (106 000€)
2024-2026	Impetus Foundation, Tracing the fate of partially reprogrammed cells in vivo (\$220 000)

Independent funding prior to PI position:

2018-2022 2020	Asociación Española Contra el Cáncer (AECC) Postdoctoral Fellowship (170 000€) La Caixa & IRB Barcelona BiomedTec Grant for Early Stage Technologies (14 500€)
2018-2020	European Molecular Biology Organization (EMBO) Postdoctoral Long-Term Non- Stipendiary Fellow (renounced financial support for alternate fellowship) (75 000€)
2018	Barcelona Institute of Science and Technology (BIST) EU Horizon 2020 Postdoctoral Fellowship (terminated early to accept alternate fellowship) (108 000€)
2018	Juan de la Cierva Formación Postdoctoral Fellowship (renounced for alternate fellowship) (50 000€)
2012-2013	Grayer Fellowship, Grayer Foundation, New York, NY (\$36 000)
2011-2012	Postgraduate Scholarship-Master's, Natural Sciences and Engineering Research Council (NSERC) of Canada (\$15 000)

SCIENTIFIC PRESENTATIONS:

Invited seminars:

2025	2 nd International conference of the Spanish society of cellular senescence (SENCEL),
	Barcelona, Spain
2025	Spanish Society of Cell Biology (SEBC) Annual Meeting, Santiago de Compostela, Spain
2025	Autumn Lecture Series, University of Rzeszów, Rzeszów Poland
2025	International Cell Senescence Association (ICSA) 10 th Annual Meeting, Rome, Italy
2025	European Association for Cancer Research (EACR) Annual Conference, Lisbon,
	Portugal Invited Speaker for 3 individual talks/sessions
2025	American Aging Association (AGE) 53 rd Annual Meeting, Alaska, USA

Marta Kovatcheva, PhD

2025	Genome Integrity Italian Network (GiiN), Online
2025	The Center for Research in Molecular Medicine and Chronic Diseases (CiMUS), Santiago
	de Compostela, Spain
2024	Institut de genetique et de biologie molecular et cellular (IGBMC), Strasbourg, France
2024	FASEB B-Vitamins and One-Carbon Metabolism, Niagara Falls, USA
2024	The Young International Cell Senescence Association Oxford Symposium, Oxford,
	United Kingdom. Keynote Speaker

Abstract selected for oral presentation:

Abstract selected for oral presentation:		
2024	Dynamic BH3 profiling for improved senolysis strategies. ICSA Conference 2024, Puerto Varas, Chile.	
2022	Vitamin B12 is a limiting factor for cellular reprogramming and tissue repair. UNIA Cellular Plasticity in Homeostasis and Disease, Baeza, Spain	
2022	A CRISPR/Cas9-based screen to identify novel senolytic targets. SENESTHERAPY-III. Spanish Network on Cell Senescence, Valencia, Spain	
2022	Vitamin B12 is a limiting factor for cellular reprogramming and tissue repair. FASEB Transcription, Chromatin, and Epigenetics in Aging Conference, Catania, Italy	
2022	Vitamin B12 is a limiting factor for cellular reprogramming and tissue repair. EMBO Developmental Circuits in Aging, Heraklion, Greece	
2022	Vitamin B12 is a limiting factor for cellular reprogramming and tissue repair. CNIO-CaixaResearch Frontiers Meeting: Molecular, Cellular, and Organismal Drivers of Aging, Madrid, Spain	
2021	BH3 profiling for tissue-specific senolytic strategies. 6th Annual Meeting of the Spanish Network on Cell Senescence, SENESTHERAPY-II. Spanish Network on Cell Senescence, Santiago de Compostela, Spain	
2016	ATRX: A new regulator of senescence, ASCB NY Graduate Student Symposium on Cell & Cancer Biology, New York, NY, USA	
2015	ATRX: A new regulator of senescence, ICSA Cell Senescence: From Physiology to Pathology, Santiago de Compostela, Spain	
2013	Bedside-to-bench: Clinical response to CDK4 inhibition uncovers a novel MDM2dependent, p53-independent senescence pathway, AACR Frontiers in Basic Cancer Research, National Harbor, MD, USA	
2012	Bedside-to-bench: Clinical response to CDK4 inhibition uncovers a novel MDM2 dependent, p53-independent senescence pathway, Cell Senescence in Cancer and Ageing, Wellcome Trust Genome Campus, Hinxton, Cambridge, UK	
2012	MDM2 is a mediator of senescence associated with CDK4 inhibition, The Molecular Genetics of Aging, Cold Spring Harbor Laboratory, Cold Spring Harbor, NY, USA	

Poster presentations:

2024	Vitamin B12 or transient Yamanaka Factor Expression enhances cell plasticity during
	recovery from ulcerative colitis and accelerates tissue repair, FUSION Stem Cell-Niche
	Interactions in the Gastrointestinal Tract, Cancun, Mexico
2016	ATRX is a new regulator of senescence, ICSA Cell Senescence. From Mechanisms to

Marta Kovatcheva, PhD

	Therapeutic Opportunities, Rehovot, Israel
2014	Using clinical insight to drive biologic discovery: MDM2 impinges on a novel, p53-
	independent pathway to senescence, EMBO CONNECT Conference:
	Ubiquitin and ubiquitin-like proteins: At the crossroads from chromatin to protein, Buenos
	Aires, Argentina
2013	Bedside-to-bench: Clinical response to CDK4 inhibition uncovers a novel
	MDM2dependent, p53-independent senescence pathway, AACR Frontiers in Basic
	Cancer Research, National Harbor, MD, USA
2012	Dissecting senescence induced by CDK4 inhibition in liposarcoma, Howard Hughes
	Medical Investigator Med into Grad Programs Northeast Peer Cluster Student Meeting,
	New York, NY, USA
2012	MDM2 is a mediator of senescence associated with CDK4 inhibition, The EMBO Meeting:
	Advancing the Life Sciences, Nice, France

PATENTS:

Enigma and CDH18 as companion diagnostics for CDK4 inhibitors, US1192116 **Granted**: March 5, 2024.

Companion diagnostics for CDK4 inhibitors, US9889135B2 **Granted**: February 13, 2018; EP2986740B1 **Granted**: January 15, 2020.

Methods for drug discovery, WO2016065044A1 Published: April 28, 2016.

ATRX as a companion diagnostic for CDK4 inhibitors, WO2016191740A1 Published: December 1, 2016.

SOCIETIES AND SCIENTIFIC COMMITTEES:

2025-present	Scientific Direction Advisory Committee (SDAC) Member, IFOM, Milan, Italy
2024-present	European Association for Cancer Research (EACR) Member
2024-present	International Cell Senescence Association (ICSA) Steering Committee Member
2022-present	Diversity, Equity & Inclusion (DEI) Working Group of the International Cell Senescence
	Association (ICSA); WG Chair since 2024
2018-2023	President, Young International Cell Senescence Association (yICSA)
2017-2023	Member, IRB Barcelona Postdoctoral Council
2012-2014	Co-Chair, Gerstner Sloan-Kettering Graduate School Student Council, New York, NY

MEETING ORGANIZATION:

2025	International Cell Senescence Association (ICSA) annual meeting, Rome, Italy
2025	IFOM Annual PI Retreat, Padenghe, Italy
2019-2023	International Cell Senescence Association (ICSA) annual meeting: yICSA invited speaker,
	poster prize judge, session moderator, & social events coordination, Athens, Greece;
	Osaka, Japan; and Groningen, The Netherlands
2020	International Cell Senescence Association (ICSA) Virtual Seminar Series, Global Online
2019	IRB Barcelona Postdoctoral Day, Barcelona, Spain
2018	Barcelona Institute of Science and Technology (BIST) Postdoctoral Day, CosmoCaixa,
	Barcelona, Spain

2012 HHMI Med Into Grad Northeast Peer Cluster Student Meeting, New York, NY

TECHNOLOGY TRANSFER, INDUSTRY RESEARCH CONTRACTS AND COLLABORATIONS:

2024-present Ad-hoc consultant. With Lever Bio (Italy). Role: Independently contracted consultant. Small-molecule inhibitors as selective senolytic drugs to treat senescence-associated 2019-2024 disorders, such as lung and kidney fibrosis. With CD3 (Belgium). Role: Lead Scientist & **Project Coordinator** Study in human dermal fibroblasts after the treatment with a proprietary compound. With 2023

Mesoestetic (Spain). Role: Principal Investigator & Consultant

2021-2022 Novel senolytic targets in inflammation and fibrosis. With Galapagos NV (Belgium).

Role: Lead Scientist & Project Coordinator

REVIEWER WORK:

2021-present **Review Editor**, Frontiers in Aging

2018-present Ad hoc reviewer for Nature Aging, Cancer Discovery, Aging Cell, iScience, FEBS Journal 2019-2020 Guest Editor, Mechanisms of Ageing & Development (Special Issue: The bright and dark

side of Cellular Senescence)

MAJOR INTERNATIONAL COLLABORATIONS (only ongoing):

2025-present: Yasuhiro Murakawa (Kyoto University, Japan): Control of intestinal cell plasticity by transposable elements as a driver of transformation

2025-present: Giorgia Ramponi (University of Zurich, Switzerland) & Rachel Zwick (New York University, USA): Using reinforcement learning to predict cell identity in intestinal development, aging, and cancer

2024-preset: Satomi Miwa (Newcastle University, UK): Sex-specific differences in aging and tissue degeneration

2022-present: Joan Montero (University of Barcelona, **Spain**): Dynamic BH3 profiling for personalized medicine and healthy aging

TEACHING, MENTORSHIP & SCIENTIFIC OUTREACH:

Current trainees (as PI):

2024-present PhD students: Devraj Parande, Saket Bhargava, Benedetta Mannelli; Postdoctoral Scientists: Valentina Ramponi, Christina Fissoun

PhD thesis committees:

2025-present	Maria Chiara Rainone (SEMM, Milan, Italy; Supervisor: Sara Sigismund; Role: Internal
	thesis advisor)
2025	Cristina Castellan (SEMM, Milan, Italy; Supervisor: Vincenzo Costanzo; Role: External
	examiner)
2024	Marta LaLinde (VHIO, Barcelona, Spain; Supervisor: Joaquin Arribas; Role: External
	examiner)
2024	Daniel Sampaio Goncalves (IBMBM, Strasbourg, France; Supervisor: William Keyes

Role: External examiner)

Marta Kovatcheva, PhD

Students trained (prior to PI):

2019-present	PhD students (co-supervisor): Valentina Ramponi, Elena Melendez (PhD 2022); IRB Barcelona, Spain
2018-2019	Master's student: Vanessa Lopez. Thesis: A CRISPR/Cas9-based screening approach to identify novel senolytic targets; UPF/BIST, Barcelona, Spain
2013-2017	Summer Undergraduate Students: Vivien Wong, Hannah Millimet, Jossie Yasshinskie, Mary Klein; Memorial Sloan Kettering Cancer Center, NY, USA

Programs & courses I have led:

2021-present	IRB Barcelona PhD-Postdoc Mentorship Program: Founder & Mentor
2019-present	IRB Barcelona-Catalan Foundation La Pedrera: Crazy about Biomedicine Course Leader,
	Barcelona, Spain
2014-2017	Scientist Educator, Science Outreach Program at Rockefeller University, New York, NY
2012-2016	Preparing and Presenting a Scientific Poster, Memorial Sloan-Kettering Cancer Center
	Medical Student Summer Fellowship, New York, NY
2012-2016	Journal Club Leader, Gerstner Sloan-Kettering Graduate School Summer
	Undergraduate Program, New York, NY
2012-2014	Teaching Assistant, Gerstner Sloan-Kettering Graduate School of Biomedical Sciences
	Core Course, Memorial Sloan-Kettering Cancer Center, New York, NY
2011	New York Academy of Sciences STEM Tutor, New York, NY

COURSE WORK & CERTFICATIONS:

2025	Newton Management & Leadership Academy, Milan, Italy
2022	Leadership Programme for Women in Science, IRB, Coach: Alicia Marin Muniesa
2019	Supervising Masters Students: Skills for Effective Supervision, Barcelona Institute of Science & Technology, Broad Associates Communication Collaboration Coach: Caroline Broad
2017	Pitch Coaching for Scientists, IRB, Coach: Beth Susanne

HONORS & AWARDS:

2025	International "Technovisionary®" Award, Women&Tech – Women and Technology Association
2024	European Association for Cancer Research (EACR) Rising Star Award
2017	Gerstner Sloan Kettering Graduate School of Biomedical Sciences Commencement Speaker
2016	American Society for Cell Biology (ASCB) New York Graduate School Symposium on Cell & Cancer Biology Oral Presentation Award
2015	Weill Cornell Graduate School & Gerstner Sloan Kettering Graduate School Three-Minute Thesis (3MT®) Competition Bronze Winner
2013	New York Academy of Sciences Science Education Fellowship Program Certificate