

ALESSIA ANGELIN

ASSISTANT PROFESSOR, PH.D.

University of Rome Tor Vergata, School of Medicine

Phone +39 06 7259 6927

Email alessia.angelin@uniroma2.it

<https://www.scopus.com/authid/detail.uri?authorId=6507126049>

<https://orcid.org/0000-0002-7115-5451>

<https://www.linkedin.com/in/alessia-angelin-ph-d-47308944/>

SUMMARY STATEMENT

Dr. Alessia Angelin obtained her Master's Degree in Biological Sciences from the University of Padua, Italy, with a thesis on the biochemical characterization of mitochondrial function under the supervision of Prof. Paolo Bernardi. At the University of Padua, she also obtained her PhD in Molecular and Cellular Biology and Pathology, working on the role of mitochondria in human muscular dystrophies, again under the supervision of Prof. Bernardi.

Dr. Angelin has 15 years of international research experience in biochemistry and biomedical sciences, and her research primarily focuses on the physiology of mitochondria in human diseases. During these years, she worked as a postdoctoral scholar at the University of Padova, the University of California-Irvine (USA), and the Children's Hospital of Philadelphia (USA) under the supervision of Prof. Douglas C. Wallace. Dr. Angelin consolidated her position as a Senior Research Scientist at the Center for Mitochondrial and Epigenomic Medicine, directed by Prof. Wallace, at the Children's Hospital of Philadelphia. She investigated the mitochondrial contribution to the pathophysiology of human cardiomyopathy and muscular dystrophy and recently developed an AAV-based gene therapy for treating human mitochondrial cardiac disorder. Over the years, her scientific interests have expanded to include the role of mitochondria in immune deficiencies and infection, metabolic diseases, and cancer. In 2024, she returned to Italy as an Assistant Professor in Biochemistry at the University of Rome Tor Vergata, School of Medicine. In Prof. Eleonora Candi's laboratory, she studies the metabolic changes that occur in the transformation of neoplastic cells and tumor proliferation using cellular and murine models with biochemical techniques and omics characterization, with particular attention to the role of mitochondria. Currently, Dr. Angelin is mainly involved in two projects: (i) characterizing the role of p63 in regulating muscular differentiation and (ii) investigating the contribution of the polyol pathway in tumor development. Dr. Angelin has published 32 peer-reviewed articles and attended about 40 scientific conferences, several as an invited speaker. Dr. Angelin teaches Biochemistry in the Pharmacy Degree Course.

EDUCATION – **(2002-2003)** Master's Degree in Biology (Molecular Biology) (110/110 with honors), University of Padua, Italy, with a biochemical thesis on mitochondrial function under the supervision of Prof. Paolo Bernardi **(2003)**; Internship at the Dept. of Experimental Biomedical Sciences and AIRC and Pfizer scholarships **(2002-2003)**

PhD – **(2004-2007)** PhD in Molecular and Cellular Biology and Pathology with a thesis on mitochondrial function in human muscular dystrophies (thesis title: "Mitochondrial pathogenesis in Ullrich congenital muscular dystrophy: new therapeutic perspectives") under the supervision of Prof. Paolo Bernardi, University of Padua, Italy **(2007)**; Doctoral student at the Dept. Experimental Biomedical Sciences and trainee at Pfizer Company (UK) as part of the Ph.D. program **(2004-2007)**

POST-DOCTORAL – **(2007-2023)** 3 years in Italy and 14 years in the USA: **2007-2010** Dept. Biomedical Sciences, University of Padua, Italy (DebioPharm fellowship; MIUR Research Grant); **2010** (03-07) Center for Molecular and Mitochondrial Medicine and Genetics, Univ. California Irvine (CA-USA); **2010-2014** Center for Mitochondrial and Epigenomic Medicine (CMEM), Children's Hospital Philadelphia (CHOP) (PA-USA); **2015-2024** Research Associate, Research Scientist, Senior Scientist at CMEM, Children's Hospital Philadelphia (USA); **2024-present** Assistant Professor in Biochemistry, University of Rome Tor Vergata, School of Medicine.

RESEARCH PROJECTS – Participated as a researcher in 19 international research projects (funded by NIH, DOD, UMDF, Frontier/CHOP, Gates Foundation, MIUR, Telethon, AIRC); she supervised and guided four scientific projects, and she was involved in two clinical studies (trials) and several collaborations with international institutes and companies. Scientific interests are focused on biochemical aspects of mitochondrial function and the

pathophysiology of mitochondrial genetic diseases, such as cardiomyopathy or muscular dystrophy. She has developed an AAV-based gene therapy to correct mitochondrial cardiomyopathy. Recently, she expanded her interest in the mitochondrial role in immune deficiencies, senescence, cancer, or during SARS-CoV-2 infection. Currently, at the Department of Experimental Medicine of the University of Rome Tor Vergata, she is mainly involved in two research projects: (i) characterization of the role of p63 in the regulation of muscle differentiation and (ii) study of the contribution of the polyol pathway in the tumor development.

TEACHING – **(2005-2006)** Integrative Teaching and Laboratory Trainer, Biochemistry Laboratory Course, Degree in Biology, University of Padua, Italy; **(2013-2014)** Guest Lecturer in the Molecular Biology Course (MolBiol305) and the Cellular Bioenergetics Course (Bio 599), Rutgers University, Candem (NJ – USA); **(2010-2024)** Speaker and organizer of the annual CMEM seminar series, at the Children's Hospital of Philadelphia (CHOP) (PA-USA), open to faculty, researchers, physicians and students of CHOP and the University of Pennsylvania (as of 2020 this includes the public international with the virtual option); **(2017-current)** Committee member of PhD thesis evaluation commission; **(2004-current)** Supervisor of undergraduate and graduate students; **(2024-2025)** Assistant Professor in Biochemistry, Biochemistry I Course, Degree Course in Pharmacy, University of Rome Tor Vergata.

CONFERENCES – Participation in over 40 international conferences as invited speaker (9), selected speaker (5), and poster presenter (19) in the following research areas: mitochondrial medicine; biochemistry and bioenergetics; human degenerative and genetic diseases; therapeutic targets; cellular metabolism and epigenomics; cell signaling and therapeutic strategy; translational research in aging and complex human diseases.

AWARDS and CERTIFICATES – **(2021)** Knight of Merit of the Italian Republic; **(2018)** National Scientific Qualification MIUR (Abilitazione Nazionale) Sector 05/E1 General Biochemistry (BIO10) Associate Professor; **(2009 and 2012)** Qualified for Faculty Member position by National Research Council CNR in Medical Sciences; **(2003)** National License (Esame di Stato) as Biologist; **(2005, 2009, 2013)** Awarded for best poster and research by international institutions and conferences; **(2011-2023)** Founder, President (for eight years), Board Member of the Association of Italian Professionals in Philadelphia (PIPhilly); **(2018)** Member AWIS Association for Women in Science; **(2014-2024)** ISSNAF Member Italian Scientist and Scholar of the North American Foundation.

MANAGEMENT ACTIVITY – **(2016-2024)** Staff member of the Administrative Director of CMEM (CHOP) with responsibility for coordination of multidisciplinary projects, department resources, and multi-million dollar budget for the laboratory and the department; **(2015-2024)** Laboratory and department manager at CMEM (CHOP) with responsibility for supervising supply orders, purchases/contracts of large equipment, biohazardous laboratory material (BSL2 and 3 levels) and safety regulations, as well as instrumentation of microscopy, sequencing and biochemistry suits.

EDITORIAL ACTIVITY – Reviewer for peer-reviewed journals "Cell Death Discovery" (nature.com/IF=6.1), "Biomolecules" (mdpi.com/IF=5.5), "Aging Clinical and Experimental Research" (springer.com/ IF=4.5), e "Human Molecular Genetics" (academic.oup.com/IF=5.1); Topic Editor Member for "Frontiers in Cellular Neuroscience - Cellular Neuropathology" (frontiersin.org/IF=5.3); Receiving Editor for peer-reviewed journal "Cell Death Disease" (nature.com/IF=8.1),

BIBLIOMETRICS – 32 Publications (Medline); 3368 Citations (Scopus ID); 23 H-index (Scopus ID)

FULL LIST OF PUBLICATIONS

Author of 32 papers published in leading peer-reviewed scientific journals

<https://pubmed.ncbi.nlm.nih.gov/?term=angelin+alessia&sort=date>

32 Publications (Medline); 3,368 Citations (Scopus ID); 23 H-index (Scopus ID)

<https://www.scopus.com/authid/detail.uri?authorId=6507126049>

<https://orcid.org/0000-0002-7115-5451>

1. Bianchi C., Fato R., **Angelin A.**, Trombetti F., Ventrella V., Borgatti AR., Fattorusso E., Ciminiello P., Bernardi P., Lenaz G., Parenti Castelli G. (2004)
Yessotoxin, a shellfish biotoxin, is a potent inducer of the permeability transition in isolated mitochondria and intact cells
Biochim. Biophys. Acta 1656:139-47 PMID: 15178475
2. Penzo D., Petronilli V., **Angelin A.**, Cusan C., Colonna R., Scorrano L., Pagano F., Prato M., Di Lisa F. and Bernardi P. (2004)
Arachidonic acid released by phospholipase A2 activation triggers Ca²⁺-dependent apoptosis through the mitochondrial pathway
J. Biol. Chem. 279(24):25219-25 PMID: 15070903
3. **Angelin A.**, Tiepolo T., Sabatelli P., Grumati P., Bergamin N., Golfieri C., Mattioli E., Gualandi F., Ferlini A., Merlini L., Maraldi NM., Bonaldo P. and Bernardi P. (2007)
Mitochondrial dysfunction in the pathogenesis of Ullrich congenital muscular dystrophy. Novel therapeutic perspectives with cyclosporins
Proc. Natl. Acad. Sci. USA 104(3):991-6 PMID: 17215366
4. **Angelin A.**, Bonaldo P., Bernardi P. (2008)
Altered threshold of the mitochondrial permeability transition pore in Ullrich congenital muscular dystrophy
Biochim. Biophys. Acta 1777:893-6 PMID: 18435905
5. Merlini L., **Angelin A.**, Tiepolo T., Braghett P., Sabatelli P., Zamparelli A., Ferlini A., Maraldi NM., Bonaldo P. and Bernardi P. (2008)
Cyclosporin A corrects mitochondrial dysfunction and muscle apoptosis in patients with collagen VI myopathies
Proc. Natl. Acad. Sci. USA 105(13):5225-9 PMID: 18362356
6. Tiepolo T.* , **Angelin A.***, Palma E., Sabatelli P., Merlini L., Nicolosi L., Finetti F., Braghett P., Vuagniaux G., Dumont JM., Baldari CT., Bonaldo P. and Bernardi P. (2009) ***Co-first Authorship**
The cyclophilin inhibitor Debio 025 normalizes mitochondrial function, muscle apoptosis and ultrastructural defects in Col6a1-/- myopathic mice
Br. J. Pharmacol. 157(6):1042-52 PMID: 19519726
7. Porcelli AM.* , **Angelin A.***, Ghelli A., Mariani E., Martinuzzi A., Carelli V., Petronilli V., Bernardi P. and Rugolo M. (2009) ***Co-first Authorship**
Respiratory complex I dysfunction due to mitochondrial DNA mutations shifts the voltage threshold for opening of the permeability transition pore toward resting levels
J. Biol. Chem. 284(4):2045-52 PMID: 19047048
8. Palma E., Tiepolo T., **Angelin A.**, Sabatelli P., Maraldi NM., Basso E., Forte MA., Bernardi P. and Bonaldo P. (2009)
Genetic ablation of cyclophilin D rescues mitochondrial defects and prevents muscle apoptosis in collagen VI myopathic mice
Hum. Mol. Genet. 18(11):2024-31 PMID: 19293339

9. Frka K., Facchinello N., Del Vecchio C., Carpi A., Curtarello M., Venerando R., **Angelin A.**, Parolin C., Bernardi P., Bonaldo P., Volpin D., Braghetta P., Bressan GM. (**2009**)
Lentiviral-mediated RNAi in vivo silencing of Col6a1, a gene with complex tissue specific expression pattern
J. Biotechnol. 141:8-17 PMID: 19428725
10. Grumati P., Coletto L., Sabatelli P., Cescon M., **Angelin A.**, Blaauw B., Tiepolo T., Urciuolo A., Maraldi NM., Bernardi P., Sandri M. and Bonaldo P. (**2010**)
Autophagy is defective in collagen VI myopathies and its induction protects against muscle degeneration
Nature Medicine 16(11):1313-20 PMID: 21037586
11. Sabatelli P., Palma E., **Angelin A.**, Squarzoni S., Urciuolo A., Pellegrini C., Tiepolo T., Bonaldo P., Gualandi F., Merlini L., Bernardi P., Maraldi NM. (**2011**)
Critical evaluation of the use of cell cultures for inclusion in clinical trials of patients affected by Collagen VI myopathies
J. Cell Physiol. 227(7):2927-35 PMID: 21953374
12. Merlini L., Sabatelli P., Armaroli A., Gnudi S., **Angelin A.**, Grumati P., Michelini ME., Franchella A., Gualandi F., Bertini E., Maraldi NM., Ferlini A., Bonaldo P., Bernardi P. (**2011**)
Cyclosporine a in Ullrich congenital muscular dystrophy: long-term result
Oxid. Med. Cell Longev. 2011:139194 PMID: 22028947
13. Ji F., Sharpley M., Derbeneva O., Alves L., Qian P., Wang Y., Chalkia D., Lvova M., Xu J., Yao W., Simon M., Platt J., Xu S., **Angelin A.**, Davila A., Huang T., Wang PH., Chuang LM., Moore L., Qiu G. and Wallace DC. (**2012**)
Mitochondrial DNA variant associated with Leber hereditary optic neuropathy and high-altitude Tibetans
Proc. Natl. Acad. Sci. USA 109(19):7391-6 PMID: 22517755
14. Chae YC., **Angelin A.**, Lisanti S., Kossenkov AV., Speicher KD., Wang H., Powers JF., Tischler AS., Pacak K., Fliedner S., Michalek RD., Karoly ED., Wallace DC., Languino LR., Speicher DW., Altieri DC. (**2013**)
Landscape of the mitochondrial Hsp90 metabolome in tumors
Nature Commun. 4:2139 PMID: 23842546 Corrigendum PMID: 26085380
15. Lin R., **Angelin A.**, Da Settimo F., Martini C., Taliani S., Zhu S. and Wallace DC. (**2014**)
Genetic analysis of dTSPO, an outer mitochondrial membrane protein, reveals its functions in apoptosis, longevity and A 42-induced neurodegeneration
Aging Cell 13(3):507-18 PMID: 24977274
16. Rivadeneira DB., Caino MC., Seo JH., **Angelin A.**, Wallace DC., Languino LR., Altieri DC. (**2015**)
Survivin promotes oxidative phosphorylation, subcellular mitochondrial repositioning, and tumor cell invasion
Sci. Signal. 8(389):ra80 PMID: 26268608
17. Beier UH., **Angelin A.**, Akimova T., Wang L., Liu Y., Xiao H., Koike MA., Hancock SA., Bhatti TR., Han R., Jiao J., Veasey SC., Sims CA., Baur JA., Wallace DC., Hancock WW. (**2015**)
Essential role of mitochondrial energy metabolism in Foxp3⁺ T-regulatory cell function and allograft survival
FASEB J. 29(6):2315-26 PMID: 25681462
18. Kokoszka JE., Waymire KG., Flierl A., Sweeney KM., **Angelin A.**, MacGregor GR., Wallace DC. (**2016**)
Deficiency in the mouse mitochondrial adenine nucleotide translocator isoform 2 gene is associated with cardiac noncompaction
Biochim. Biophys. Acta 1857(8):1203-12 PMID: 27048932

19. Kandel J., Angelin A., Wallace D.C., Eckmann D.M. (2016)
Mitochondrial respiration is sensitive to cytoarchitectural breakdown
Integrative Biology 8(11):1170-82 PMID: 27734042
20. Angelin A., Gil-de-Gomez L., Dahiya S., Jiao J., Guo L., Levine MH., Wang Z., Quinn WJ., Kopinski PK., Wang L., Akimova T., Liu Y., Bhatti TR., Han R., Laskin BL., Baur JA., Blair IA., Wallace DC., Hancock WW., Beier UH., (2017)
Foxp3 Reprograms T Cell Metabolism to Function in Low-Glucose, High-Lactate Environment
Cell Metabolism 25(6):1282-93 PMID: 28416194
21. Bertholet AM., Chouchani ET., Kazak L., Angelin A., Fedorenko A., Long JZ., Vidoni S., Garrity R., Cho J., Terada N., Wallace DC., Spiegelman BM., Kirichok Y. (2019)
H⁺ transport is an integral function of the mitochondrial ADP/ATP carrier
Nature 571(7766):515-520 PMID: 31341297
22. McManus MJ., Picard M., Chen HW., De Haas HJ., Potluri P., Leipzig J., Towheed A., Angelin A., Sengupta P., Morrow RM., Kauffman BA., Vermulst M., Narula J., Wallace DC. (2019)
Mitochondrial DNA Variation Dictates Expressivity and Progression of Nuclear DNA Mutations Causing Cardiomyopathy
Cell Metabolism 29(1):78-90 PMID: 30174309
23. Rossidis AC., Angelin A., Lawrence KM., Baumgarten HD., Kim AG., Mejaddam AY., Coons BE., Hartman HA., Hwang G., Monos S., Davey MG., Murdock D., Wallace DC., and Flake AW. (2019)
Premature Lambs Exhibit Normal Mitochondrial Respiration after Long-term Extrauterine Support
Fetal Diagnosis and Therapy 46(5):306-312 PMID: 30861524
24. Quinn WJ., Jiao J., TeSlaa T., Stadanlick J., Wang Z., Wang L., Akimova T., Angelin A., Schäfer PM., Cully MD., Perry C., Kopinski PK., Guo L., Blair IA., Ghanem LR., Leibowitz MS., Hancock WW., Moon EK., Levine MH., Eruslanov EB., Wallace DC., Baur JA., Beier UH. (2020)
Lactate Limits T Cell Proliferation via the NAD(H) Redox State
Cell Rep. PMID: 33326785, 33(11):108500. doi: 10.1016/j.celrep.2020.108500
25. Dahiya S., Beier UH., Wang L., Han R., Jiao J., Akimova T., Angelin A., Wallace DC., Hancock WW. (2020)
HDAC10 deletion promotes Foxp3+ T-regulatory cell function
Sci. Rep. 10(1):424 PMID: 31949209
26. Mereness JA., Bhattacharya S., Ren Y., Wang Q., Anderson CS., Donlon K., Dylag AM., Haak J., Angelin A., Bonaldo P., Mariani TJ. (2020)
Collagen VI Deficiency Results in Structural Abnormalities in the Mouse Lung
Am. J. Pathol. 190(2):426-441 PMID: 31837950
27. Angelin A. (2023)
Cyclin-dependent kinases regulate the adult nervous system via the one-carbon-metabolism
Cell Death & Disease. 14(7):429. doi: 10.1038/s41419-023-05950-6
(Commentary)
28. Guarnieri JW., Angelin A., Murdock DG., Schaefer P., Portluri P., Lie T., Huang J., Wallace DC. (2023)
SARS-CoV-2 viroporins activate the NLRP3-inflammasome by the mitochondrial permeability transition pore
Front. Immuno.I PMID: 36891303, doi: 10.3389/fimmu.2023.1064293
29. Guarnieri JW., Dybas JM., Fazelinia H., Kim MS., Frere J., Zhang Y., Albrecht YS., Murdock DG., Angelin A., Singh LN., Weiss SL., et al. (2023)
Core mitochondrial genes are down-regulated during SARS-CoV-2 infection of rodent and human

hosts **Sci Transl Med.** 2023 Aug 9;15(708). Epub 2023 Aug 9 PMID: 37556555

30. Jain S, Pei L, Spraggins JM, Angelo M, Carson JP, Gehlenborg N, Ginty F, Gonçalves JP, Hagood JS, Hickey JW, Kelleher NL, Laurent LC, Lin S, Lin Y, Liu H, Naba A, Nakayasu ES, Qian WJ, Radtke A, Robson P, Stockwell BR, Van de Plas R, Vlachos IS, Zhou M, **HuBMAP Consortium***, et al. (2023)
**Angelin Alessia Author as part of the HuBMAP Consortium*
Advances and prospects for the Human BioMolecular Atlas Program (HuBMAP)
Nat Cell Biol. Aug;25(8):1089-1100 PMID: 37468756 Author Correction PMID: 38429479
(Review)
31. Guarnieri JW., Haltom JA., Albrecht YES., Lie T., Olali AZ., Widjaja GA., Ranshing SS., **Angelin A.**, Murdock D., Wallace DC. (2024)
SARS-CoV-2 mitochondrial metabolic and epigenomic reprogramming in COVID-19.
Pharmacol Res. 2024 Apr 12;204:107170. doi: 10.1016/j.phrs.2024.107170.
PMID: 38614374
(Review)
32. Guarnieri JW., Lie T, Albrecht YES., Hewin P., Jurado KA., Widjaja GA., Zhu Y., McManus MJ., Kilbaugh TJ., Keith K., Potluri P., Taylor D., **Angelin A.**, Murdock DG., Wallace DC. (2024)
Mitochondrial antioxidants abate SARS-CoV-2 pathology in mice.
Proc Natl Acad Sci USA. 2024 Jul 23;121(30):e2321972121. doi: 10.1073/pnas.2321972121. Epub 2024 Jul 15. PMID: 39008677