

## **Chiara Mondello**

### **Curriculum vitae**

#### **Education**

1980: Graduation in Biological Sciences cum laude, University of Pavia, Italy.

1984: Advanced Studies in Genetics cum laude, University of Pavia, Italy.

1987: PhD in Genetics and Molecular Biology, University of Pavia, Italy.

#### **Professional positions**

1981: Fellowship of the University of Pavia, Istituto di Genetica Biochimica ed Evoluzionistica of the Consiglio Nazionale delle Ricerche (CNR).

1982-1983: CNR Fellowship, Istituto di Genetica Biochimica ed Evoluzionistica, CNR, Pavia (Italy).

1984-1985: Post-doctoral EMBO Fellowship, Imperial Cancer research Fund, London (UK).

1986-2001: CNR Researcher, Istituto di Genetica Biochimica ed Evoluzionistica, CNR, Pavia, Italy.

1992-1994: Professor a contratto, University of Pavia (Faculty of Science).

2001-present: Senior Researcher, Istituto di Genetica Molecolare of CNR, Pavia, Italy.

2002-2009; 2016-2020: Member of the Scientific Committee of the Istituto di Genetica Molecolare of CNR.

2005-present: Member of the CNR list of: "Esperti in gestione progettuale".

#### **Memberships**

Member of the New York Academy of Science.

Member of the Italian Genetic Association.

#### **Editorial activity**

Member of the Editorial Board of *Scientific Reports*; *Mutation Research – Reviews in Mutation Research*; *Oncology Letters*; *Experimental and Therapeutic Medicine*; *Stem Cell Discovery*.

#### **Leadership in projects**

1995-1997: AIRC project: "Genome Instability in XP fibroblasts used as a model to study cellular transformation".

1996-1998: Partner in the EU BIOMED 2 Concerted Action: "Mammalian chromosome stability and cancer".

2001-2003: Partner in the EU project "Telomere instability and the formation and transmission of radiation induced DNA damage".

2003-2005: Partner in the EU project "Telomeres and radiosensitivity of individuals".

2004-2008: FIRB Project from Italian Ministry of Research "Nuovi sistemi cellulari per studiare la riparazione delle rotture a doppio filamento del DNA in cellule di mammifero".

2007-2010: Project funded by Fondazione Cariplo "Genomics of tumor progression using a model cellular system".

2011-2014: Project funded by Fondazione Cariplò “Optofluidic chips for the study of cancer cell mechanical properties and invasive capacities.”

### **Reviewer activity**

Reviewer for the following scientific journals: Proc. Nat. Acad. Science USA; International Journal of Cancer; European Journal of Cancer; Molecular Cancer Research; Oncotarget; Carcinogenesis; Cancers; Scientific Reports; PLoS One; BMC Medical Genetics; BBA - Molecular Basis of Disease; Mechanisms of Ageing and Development; Biochimie; Bioorganic Chemistry; Biomolecules; Cytogenetics and Genome Research; International Journal of Oncology; Oncology Reports; Experimental and Therapeutic Medicine; Journal of Experimental Medicine; Molecular Medicine Reports; International Journal of Cell Biology; Environmental Pollution; Mutation Research Reviews; chapters in Springer books.

Evaluator of scientific projects for Swiss Cancer League, Swiss National Science Foundation and Belgian FNRS (Fonds de la Recherche Scientifique).

Member of the CNR panel of evaluators for the Projects launched by the Italian Ministero dello Sviluppo Economico, in the framework of the “Fondo per la crescita sostenibile”.

### **Research activity**

The main research interests have been: study of DNA repair in mammalian cells: isolation and characterization of UV sensitive mutants; cytological aspects of DNA damage. Molecular analysis of the terminal region of the human X chromosome short arm. Chromosomal instability in mammalian cells: mechanisms of origin of rearranged chromosomes and their role in cellular transformation and senescence. Telomere metabolism in mammalian cells. Interstitial telomeric sequences. Factors and genes controlling gene amplification in mammalian cells. Use of *in vitro* cellular model systems to study the process of neoplastic transformation, metabolic reprogramming and cancer stem cell generation during transformation.

### **Publications**

She is author of 93 full papers, 4 chapters in books and 112 Abstracts in Meetings. She has been the Editor of the book “Multiple Pathways in neoplastic transformation.” Transworld research network. Kerala (India).

H index: 28 ISI web of knowledge; 28 Scopus; 31 Google Scholar.

### **List of Publications**

- 1) Giulotto E. and Mondello C. Aphidicolin does not inhibit the repair synthesis of mitotic chromosomes. *Biochem. Biophys. Res. Comm.* 99: 1287-1294 (1981).
- 2) Stefanini M., Mondello C. and Nuzzo F. Studies on DNA repair in mammals: Isolation and characterization of mutagen sensitive cell lines. In: Proceeding of a Meeting on Cancer Risk and DNA Repair, p. 111-119. Ed. by A. Castellani (1982).

- 3) Mazza C., Mondello C. and Nuzzo F. Determinazione dell'attività mutagena e genotossica dell'antibiotico Distamicina A. Giornale Italiano di Chemioterapia. 30: 39-52 (1983).
- 4) Mondello C., Giorgi R. and Nuzzo F. Chromosomal effects of methotrexate on cultured human lymphocytes. Mutat. Res. 139: 67-70 (1984).
- 5) Mondello C., Zei G. and Nuzzo F. Correlation between unscheduled DNA synthesis and chromosome condensation in mitoses from human lymphocytes. Mutat. Res. 142: 45-48 (1985).
- 6) Mondello C. and Goodfellow P.N. Methylation and expression of a housekeeping gene. Trends Genet. 1: 124-125 (1985).
- 7) Buckle V., Mondello C., Darling S., Craig I.W. and Goodfellow P.N. Homologous expressed genes in the human sex chromosome pairing region. Nature 317: 739-741 (1985).
- 8) Stefanini M., Mondello C., Tessera L., Capuano V., Guerra B.R. and Nuzzo F. Sensitivity to DNA damaging agents and mutation induction in UV sensitive CHO cells. Mutat. Res. 174: 155-159 (1986).
- 9) Ballabio A., Parenti G., Tippet P., Mondello C., Di Maio S., Tenore A. and Andria G. X-linked ichthyosis due to steroid sulphatase deficiency, associated with Kallmann syndrome (hypogonadotropic hypogonadism and anosmia): linkage relationship with Xg and cloned DNA sequences from the distal short arm of the X chromosome. Hum. Genet. 72: 237-240 (1986).
- 10) Mondello C., Ropers H.H. , Craig I.W., Tolley E. and Goodfellow P.N. Physical mapping of genes and sequences at the end of the human X chromosome short arm. Ann. of Hum. Genet. 51: 137-143 (1987).
- 11) Goodfellow P.J., Darling S., Banting G., Pym B., Mondello C. and Goodfellow P.N. Pseudoautosomal genes in man. In: "The mammalian Y chromosome, Molecular search for the sex determining gene" Editors Goodfellow P.N., Wolf J. and Craig I.W., Development 101: 119-125(s) (1987).
- 12) Stefanini M., Mondello C., Tessera M.L., Botta E. and Nuzzo F. Cellular and genetic studies in three UV-sensitive Chinese hamster mutants. Cytotechnology 1: 91-94 (1987).
- 13) Talarico D., Peverali A.F., Ginelli E., Meneveri R., Mondello C. and Della Valle G. Satellite DNA induces unstable expression of the adjacent Herpes Simplex Virus TK gene cotransfected in mouse cells. Molecular and Cellular Biology 8: 1336-1344 (1988).
- 14) Goodfellow P.J., Mondello C., Darling S.M., Pym B., Little P. and Goodfellow P.N. Absence of methylation of a CpG rich region at the 5' end of the MIC2 gene is the same on the active X, inactive X and the Y chromosome. Proc. Natl. Acad. Sci. USA 85: 5605-5609 (1988).
- 15) Mondello C., Goodfellow P.J. and Goodfellow P.N. Analysis of methylation of a human X located gene which escapes X inactivation. Nucleic Acids Res. 16: 6813-6824 (1988).
- 16) Banting G., Mondello C., Hope R. and Goodfellow P.N. A monoclonal antibody, R1, and a polyclonal serum, S10, recognize the same molecules: a novel use of DNA transfectants. J. of Immunogenet. 15: 257-265 (1988).
- 17) Stefanini M., Mondello C., Botta E., Riboni R. and Nuzzo F. Cellular and genetic characterization of UV sensitive Chinese hamster mutants. Ann. Ist. Sup. Sanità 25: 123-130 (1989).

- 18) Mazzarello P., Verri A., Mondello C., Colombo S., Maga G., Spadari S. and Focher F. Enzymes of DNA metabolism in a patient with Wiedemann-Rautenstrauch progeroid syndrome. *Annales of the New York Academy of Sciences*. Vol. 663: 440-441 (1992).
- 19) Fontana M., Lestingi M., Mondello C., Braghetta A., Montecucco A., Ciarrocchi G. DNA binding properties of FCE 24517, an electrophilic distamycin analogue. *Anti-Cancer Drug Design*. 7: 131-141 (1992).
- 20) Casati A., Giorgi R., Lanza A., Raimondi E., Vagnarelli P., Mondello C., Ghetti P., Piazza G. and Nuzzo F. Trisomy mosaicism in two subjects from two generations. *Ann. Génét.* 35, 245-250(1992).
- 21) Braghetta A., Piazzesi G., Lanfranco L. and Mondello C. Multiple DNA-protein interactions at the CpG island of the human pseudoautosomal gene MIC2. *Somat. Cell. Molec. Genet.* 19: 51-63, (1993).
- 22) Montecucco A., Capolongo L., Melegara G., Mondello C. and Ciarrocchi G. Temperature influences both cytotoxicity and DNA nicking efficiency of the antitumor Distamycin analogue FCE24517. *Anticancer Res.* 14: 189-192 (1994).
- 23) Mondello C., Nardo T., Giliani S., Arrand J.E., Weber C.A., Lehmann A.R., Nuzzo F. and Stefanini M. Molecular analysis of the XP-D gene in Italian families with patients affected by trichothiodystrophy and xeroderma pigmentosum group D. *Mutat. Res. DNA repair* 314: 159-165 (1994).
- 24) Mondello C., Casati A., Riboni R. and Nuzzo F. Structural instability of a transmissible end-to-end dicentric chromosome in a xeroderma pigmentosum fibroblast clone. *Cancer Genet. Cytogenet.* 79: 41-48 (1995).
- 25) Mondello C., Riboni R., Rady M., Giulotto E. and Nuzzo F. Gene amplification in chinese hamster DNA repair deficient mutants. - *Mutation Res.* 346: 61-67 (1995)
- 26) Casati A., Riboni R., Caprioli J., Nuzzo F. and Mondello C. Condensation anomalies and exclusion in micronuclei of rearranged chromosomes in human fibroblasts cultured in vitro. *Chromosoma* 104: 137-142 (1995).
- 27) Riboni R., Casati A., Nardo T., Zaccaro E., Ferretti L., Nuzzo F. and Mondello C. Telomeric fusions in human cultured fibroblasts as a source of genomic instability. *Cancer Genetics and Cytogenetics* 95: 130-136 (1997).
- 28) Mondello C., Riboni R., Casati A., Nardo T. and Nuzzo F. Chromosomal instability and telomere length variations during the life span of human fibroblast clones. *Exp. Cell. Res.* 236: 385-396 (1997).
- 29) Giulotto E. and Mondello C. Telomeres. In "Advances in Genome Biology", Vol 5B pages 323-361. JAI Press Inc. (1998).
- 30) Mondello C. I telomeri. *BioTec* 3: 41-52 (1998).
- 31) Mondello C., Petropoulou C., Monti D., Gonos E.S., Franceschi C. and Nuzzo F. Telomere length in fibroblast and blood cells from healthy centenarians. *Exp. Cell Res.* 248: 234-242 (1999).
- 32) Mondello C., Moralli D., Franceschi C. and Nuzzo F. Occurrence and expansion of trisomy 7 in a centenarian fibroblast strain. *Exp. Gerontol.* 34: 717-719 (1999).
- 33) Franceschi C., Mondello C., Bonafè M., Valesin S., Sansoni P. and Sorbi S. Long telomeres and well preserved proliferative vigor in cells from centenarians: A contribution to longevity? *Aging Clin. Exp. Res.* 11: 69-72 (1999).

- 34) Mucciolo E., Bertoni L, Mondello C. and Giulotto E. Late onset of CAD gene amplification in unamplified PALA resistant Chinese hamster mutants. *Cancer Letters* 150: 119-127 (2000).
- 35) Mondello C., Pirzio L., Azzalin C. and Giulotto E. Instability of interstitial telomeric sequences in the human genome. *Genomics* 68: 111-117 (2000).
- 36) Mondello C., Faravelli M., Pipitone L., Rollier A., Di Leonardo A. and Giulotto E. Gene amplification in fibroblasts from ataxia telangiectasia patients and in X-ray hypersensitive Chinese hamster mutants. *Carcinogenesis* 22: 141-145 (2001).
- 37) Mondello C., Rebuzzini P., Dolzan M., Edmonson S., Taccioli G.E., and Giulotto E. Increased gene amplification in immortalized rodent cells deficient in the catalytic subunit of the DNA-dependent protein kinase. *Cancer Res.* 61: 4520-4525 (2001).
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- 39) Mondello C., Guasconi V., Giulotto E and Nuzzo F.  $\gamma$ -Ray and hydrogen peroxide induction of gene amplification in hamster cells deficient in DNA double strand break repair. *DNA repair* 1: 483-493 (2002).
- 40) Mondello C., Chiesa M., Rebuzzini P., Zongaro S., Verri A., Colombo T., Giulotto E., D'Incàlci M., Franceschi C. and Nuzzo F. Karyotype instability and anchorage independent growth in telomerase immortalized fibroblasts from two centenarian individuals. *Biochem. Biophys. Res. Commu.* 308: 914-921 (2003).
- 41) Rebuzzini P., Lisa A., Giulotto E. and Mondello C. Chromosomal end-to-end fusions in immortalized mouse embryonic fibroblasts deficient in the DNA-dependent protein kinase catalytic subunit. *Canc. Lett.* 203: 79-86 (2004).
- 42) Desmaze C., Pirzio L.M., Blaise R., Mondello C., Giulotto E., Murnane J.P. and Sabatier L. Interstitial telomeric repeats are not preferentially involved in radiation-induced chromosome aberrations in human cells. *Cytogenet Genome Res.* 104: 123-130 (2004).
- 43) Mondello C. and Scovassi A.I. Telomeres, telomerase and apoptosis. *Biochem.and Cell Biol.* 82: 498-507 (2004).
- 44) Nergadze S.G., Rocchi M., Azzalin C.M., Mondello C. and Giulotto E. Insertion of telomeric repeats at intrachromosomal break sites during primate evolution. *Genome Res.* 14: 1704-1710 (2004).
- 45) Rebuzzini P., Khoriauli L., Azzalin C. M., Magnani E., Mondello C.\* and Giulotto E. New mammalian cellular systems to study mutations introduced at the break site by non-homologous end-joining. *DNA repair* 4: 546-555 (2005). Corresponding author
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- 48) Mondello C., Zongaro S., and D'Incàlci M. Telomerase Expression in Somatic Cells: Fountain of Youth or Damocles' Sword? *Cell Cycle* 5: 465-466 (2006)

- 49) Mondello C., Bottone MG., Noriki S., Soldani C., Pellicciari C. and Scovassi A.I. Oxidative Stress Response in Telomerase-Immortalized Fibroblasts from a Centenarian. *Ann. N. Y. Acad. Sci.* 1091: 94-101 (2006).
- 50) Rebuzzini P., Martinelli P., Blasco M., Giulotto E. and Mondello C. Inhibition of gene amplification in telomerase deficient immortalized mouse embryonic fibroblasts. *Carcinogenesis* 283: 553-559 (2007). Epub 2006 Sep 14.
- 51) Paulis M., Bensi M., Orioli D., Mondello C., Mazzini G., D'Incalci M., Falcioni C., Radaelli E., Erba E., Raimondi E., and De Carli L. Transfer of a human chromosomal vector from a hamster cell line to a mouse embryonic stem cell line. *Stem Cells* 25: 2543-2550 (2007). Epub 2007 Jul 5.
- 52) Nergadze S.G., Santagostino M.A., Salzano A., Mondello C., and Giulotto E. Contribution of telomerase RNA retrotranscription to DNA double-strand break repair during mammalian genome evolution. *Genome Biol.* 8: R260 (2007)
- 53) Donà F., Mondello C. and Scovassi A.I. Poly (ADP-ribosylation) at telomeres. *Trends in Cell & Molecular Biology* 2: 77-88 (2007).
- 54) Zongaro S., Verri A., Giulotto E., and Mondello C. Telomere length and radiosensitivity in human fibroblast clones immortalized by ectopic telomerase expression. *Oncology Reports* 19: 1605-1609 (2008).
- 55) Donà F., Prosperi E., Savio M., Coppa T., Scovassi A. I., and Mondello C. Loss of Histone H2AX increases sensitivity of immortalized mouse fibroblasts to the topoisomerase inhibitor etoposide. *Int. J. Oncol.* 33: 613-621 (2008).
- 56) Belgiovine C., Chiodi I., and Mondello C. Telomerase: cellular immortalization and neoplastic transformation. Multiple functions of a multifaceted complex. *Cytogenetic and Genome Research*. 122: 255-262 (2008).
- 57) Maga G., Crespan E., Wimmer U., van Loon B., Amoroso A., Mondello C., Belgiovine C., Ferrari E., Villani G. and Hübscher U. Replication Protein A and Proliferating Cell Nuclear Antigen coordinate DNA polymerase selection in 8-oxo-guanine repair. *Proc. Natl. Acad. Sci. USA.* 105: 20689-20694 (2008).
- 58) Salzano A., Kochiashvili N., Nergadze S.G., Khoriauli L., Smirnova A., Herrera A., Mondello C., and Giulotto E. Enhanced gene amplification in human cells knocked down for DNA-PKcs. *DNA Repair* 8: 19-28 (2009).
- 59) Mondello C., Smirnova A., and Giulotto E. Gene amplification, radiation sensitivity and DNA double strand breaks. *Mutat. Res.* 704: 29-37 (2010). Epub ahead of print January 2010.
- 60) Belgiovine C., Frapolli R., Bonezzi K., Chiodi I., Mello-Grand M., Favero F., Dei Tos A. P., Giulotto E., Tarabocchetti G., D'Incalci M., Mondello C. ROCK inhibitor RhoE is associated with increased invasiveness and metastatic potential in mesenchymal tumor cells. *PLoS ONE* 5: e141542010 (2010).
- 61) Mondello C. and Scovassi I. Apoptosis: a way to maintain healthy individuals. In "Genome Stability and Human Diseases" Nasheuer, H.P. (Ed.). *Subcell Biochem.* 50: 307-323 (2010). ISBN 978-90-481-3470-0
- 62) Ruiz-Herrera A., Smirnova A., Khoriauli L., Nergadze S.G., Mondello C. and Giulotto E. Gene amplification in human cells knocked down for RAD54. *Genome Integrity.* 2: 5 (2011).
- 63) Chiodi I., Belgiovine C., Donà I., Scovassi A.I. and Mondello C. Drug treatment of cancer cell line: a way to select for cancer stem cells? *Cancers* 3: 1111-1128 (2011).

- 64) Belgiovine C., Chiodi I. and Mondello C. Relocalization of cell adhesion molecules during neoplastic transformation of human fibroblasts. *Int. J. Oncol.* 39: 1199-1204 (2011).
- 65) Ostano P., Bione S., Belgiovine C., Chiodi I., Ghimenti C., Scovassi A.I., Chiorino G. and Mondello C. Cross-analysis of gene and miRNA genome-wide expression profiles in human fibroblasts at different stages of transformation. *OMICS* 16: 24-36 (2012).
- 66) Donà F., Chiodi I., Belgiovine C., Raineri T., Ricotti R., Mondello C. and Scovassi A.I. Poly(ADP-ribosylation) and neoplastic transformation: effect of PARP inhibitors. *Current Pharm. Biotechnol.* 2012 Mar 20. [Epub ahead of print].
- 67) Vidale P., Magnani E., Nergadze S.G., Santagostino M., Cristofari G., Smirnova A., Mondello C. and Giulotto E. The catalytic and the RNA subunits of human telomerase are required to immortalize equid primary fibroblasts. *Chromosoma* 121: 475-488, 2012. Epub 2012 Jul 14.
- 68) Bragheri F., Minzioni P., Martinez Vazquez R., Bellini N., Paiè P., Mondello C., Ramponi R., Cristiani I. and Osellame R. Optofluidic integrated cell sorter fabricated by femtosecond lasers. *Lab Chip* 12: 3779-384, 2012.
- 69) Chiodi I. and Mondello C. Telomere-independent functions of telomerase in nuclei, cytoplasm, and mitochondria. *Frontiers in Oncology “Telomeres: structure, functions and therapeutic opportunities”*. *Front Oncol.* 2:133, 2012. doi: 10.3389/fonc.2012.00133. Epub 2012 Sep 28.
- 70) Chiodi I., Belgiovine C., Zongaro S., Ricotti R., Horard B., Lossani A., Focher F., Gilson E., Giulotto E. and Mondello C. Super-telomeres in transformed human fibroblasts. *Biochim Biophys Acta*. 1833:1885-1893, 2013. Epub 2013 Apr 6. doi: 10.1016/j.bbamcr.2013.03.030.
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- \*Corresponding author
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- Mondello C, Raju J, Salem HK, Woodrick J, Scovassi I, Singh N, Vaccari M, Roy R, Forte S, Memeo L, Kim SY, Bisson WH, Lowe L and Park HH. Disruptive environmental chemicals and cellular mechanisms that confer resistance to cell death. *Carcinogenesis*. 2015; 36 Suppl 1:S89-S110.
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