

Michela Clerici



Personal data

Name Michela Clerici
Birth February 27th, 1975 in Bollate, Italy
Citizenship Italian
Work address Università di Milano-Bicocca, Piazza della Scienza 2, 20126 Milano, Italy
Phone +39 0264483547
e-mail michela.clerici@unimib.it

Present position

Associate Professor in Genetics (SSD BIO18) at the University of Milano-Bicocca

Education

Gen. 2004 PhD in Industrial Biotechnology at the University of Milano-Bicocca
July 2000 Degree in Biology, Summa cum Laude, at the University of Milano

Research experience

March 2015-present Associate Professor in Genetics at the Department of Biotechnology and Biosciences, University of Milano-Bicocca
2008-2015 Assistant Professor in Genetics at the Department of Biotechnology and Biosciences, University of Milano-Bicocca
2004-2008 Post-doctoral fellow in the laboratory of Prof. Maria Pia Longhese at the Department of Biotechnology and Biosciences, University of Milano-Bicocca (fellowships from University of Milano-Bicocca and Fondazione Telethon)
2000-2003 PhD student in the laboratory of Prof. Maria Pia Longhese at the University of Milano-Bicocca
2000 Graduate fellow in the laboratory of Dr. Maria Pia Longhese (fellowship from “Biopolo”)
1999-2000 Undergraduate student in the laboratory of Prof. Giovanna Lucchini and Dr. Maria Pia Longhese at the Department of Genetics, University of Milano.

Awards

2007 Giovanni Magni award from Adriano-Buzzati Traverso Foundation for the best paper published by a young researcher in microorganism genetics (The *S. cerevisiae* Sae2 protein negatively regulates DNA damage checkpoint signaling. 2006. *EMBO Reports*, 7:212-218).

Research grants

2010-2013 Exploring the link between gene transcription and telomere regulation in *Saccharomyces cerevisiae*. MIUR-PRIN2009

Technical skills and competences

Expert in yeast genetics, Michela Clerici is experienced with genetic, biochemistry, molecular biology and cell biology techniques, as well as with fluorescence microscopy and image processing. She is particularly skilled in generating mutants by both site-specific and random mutagenesis and in performing genome-wide genetic screenings. Finally, she contributed to generate a wide collection of yeast strains and mutants.

Scientific activity

Michela Clerici has always been using the budding yeast *Saccharomyces cerevisiae* as a eukaryotic model system to explore the mechanisms preserving genome stability in eukaryotes, focusing in particular on the cellular response to DNA damage and replication stress, as well as on the control of the homeostasis of telomeres, the nucleoprotein specialized structures that protect the natural ends of chromosomes. Genetic instability can induce tumorigenesis in multicellular organisms, and replication stress is emerging as a hallmark of cancer cells. Therefore, unveil the molecular mechanisms underlying the response to DNA perturbations is of considerable biological relevance.

The cellular response to DNA damage and replication stress is orchestrated by evolutionarily conserved checkpoint pathways, which preserve genome stability by delaying cell cycle progression in the presence of DNA alterations, concomitantly activating DNA repair systems and regulating DNA replication. Thanks to the use of multidisciplinary approaches, Michela Clerici's research has provided significant insights into these mechanisms. In particular she contributed to elucidate i) the functions and regulation of different checkpoint proteins, among which the central conserved protein kinases Mec1/ATR and Tel1/ATM, ii) the complex relationships among checkpoint and DNA repair systems, iii) the role of the master regulator of cell cycle progression, the cyclin-dependent kinase Cdk1, in promoting specific homologous recombination-dependent repair pathways, and iv) the molecular mechanisms regulating the nucleolytic processing of DNA ends, an event particularly important to repair DNA double-strand breaks and to maintain telomere stability.

Furthermore, her more recent research activity has been unveiling a novel and specific function of the checkpoint kinase Tel1/ATM in supporting DNA replication upon the generation of a topological stress imposed by the poisoning of DNA topoisomerase.

Bibliometry

The results of Michela Clerici's research activity are documented by 32 papers, which have been published in international peer-reviewed journals. She is corresponding author of 8 of these papers.

Total peer-reviewed publications	32
H-index (Scopus source)	21
Total number of citations (Scopus source)	1713
Publications in the last 5 years	9
Total citations in the last 5 years (Scopus source)	693

Publications of the last 10 years

1. Menin L, Colombo CV, Maestrini G, Longhese MP*, Clerici M*. 2019. Tel1/ATM Signaling to the Checkpoint Contributes to Replicative Senescence in the Absence of Telomerase. *Genetics* 213:411-429. doi: 10.1534/genetics.119.302391.
2. Colombo CV, Menin L, Ranieri R, Bonetti D, Clerici M*, Longhese MP*. 2019 Uncoupling Sae2 Functions in Downregulation of Tel1 and Rad53 Signaling Activities. *Genetics* 211:515-530. doi: 10.1534/genetics.118.301830.
3. Bonetti D, Colombo CV, Clerici M, Longhese MP. 2018. Processing of DNA Ends in the Maintenance of Genome Stability. *Front Genet.* 9:390. doi: 10.3389/fgene.2018.00390.
4. Menin L, Ursich S, Trovesi C, Zellweger R, Lopes M, Longhese MP*, Clerici M*. 2018. Tel1/ATM prevents degradation of replication forks that reverse after topoisomerase poisoning. *EMBO Rep.* 19 pii: e45535. doi: 10.15252/embr.201745535.
5. Colombo CV, Menin L, Clerici M*. 2018. Alkaline Denaturing Southern Blot Analysis to Monitor Double-Strand Break Processing. *Methods Mol Biol.* 1672:131-145. doi: 10.1007/978-1-4939-7306-4_11.
6. Colombo CV, Trovesi C, Menin L, Longhese MP*, Clerici M*. 2017. The RNA binding protein Npl3 promotes resection of DNA double-strand breaks by regulating the levels of Exo1. *Nucleic Acids Res.* 45:6530-6545. doi: 10.1093/nar/gkx347.
7. Cassani C, Gobbin E, Wang W, Niu H, Clerici M, Sung P, Longhese MP. 2016. Tel1 and Rif2 Regulate MRX Functions in End-Tethering and Repair of DNA Double-Strand Breaks. *PLoS Biol.* 14:e1002387. doi: 10.1371/journal.pbio.1002387.
8. Gobbin E, Villa M, Gnugnoli M, Menin L, Clerici M, Longhese MP. 2015. Sae2 Function at DNA Double-Strand Breaks Is Bypassed by Dampening Tel1 or Rad53 Activity. *PLoS Genet.* 11:e1005685. doi: 10.1371/journal.pgen.1005685.
9. Manfrini N, Clerici M, Wery M, Colombo CV, Descrimes M, Morillon A, d'Adda di Fagagna F, Longhese MP. 2015. Resection is responsible for loss of transcription around a double-strand break in *Saccharomyces cerevisiae*. *Elife.* 4. doi: 10.7554/eLife.08942.
10. Clerici M, Trovesi C, Galbiati A, Lucchini G, Longhese MP. 2014. Mec1/ATR regulates the generation of single-stranded DNA that attenuates Tel1/ATM signaling at DNA ends. *EMBO J.* 33:198-216. doi: 10.1002/emboj.201386041.
11. Bonetti D, Anbalagan S, Lucchini G, Clerici M*, Longhese MP*. 2013. Tbf1 and Vid22 promote resection and non-homologous end joining of DNA double-strand break ends. *EMBO J.* 32:275-89. doi: 10.1038/emboj.2012.327.
12. Fumagalli M, Rossiello F, Clerici M, Barozzi S, Cittaro D, Kaplunov JM, Bucci G, Dobрева M, Matti V, Beausejour CM, Herbig U, Longhese MP, d'Adda di Fagagna F. 2012. Telomeric DNA damage is irreparable and causes persistent DNA-damage-response activation. *Nat Cell Biol.* 14:355-65. doi: 10.1038/ncb2466.
13. Martina M, Clerici M, Baldo V, Bonetti D, Lucchini G, Longhese MP. 2012. A balance between Tel1 and Rif2 activities regulates nucleolytic processing and elongation at telomeres. *Mol Cell Biol.* 32:1604-17. doi: 10.1128/MCB.06547-11.
14. Trovesi C, Falcettoni M, Lucchini G, Clerici M*, Longhese MP*. 2011. Distinct Cdk1 requirements during single-strand annealing, noncrossover, and crossover recombination. *PLoS Genet.* 7:e1002263. doi: 10.1371/journal.pgen.1002263.

15. Bonetti D, Clerici M, Manfrini N, Lucchini G, Longhese MP. 2010. The MRX complex plays multiple functions in resection of Yku- and Rif2-protected DNA ends. *PLoS One* 5:e14142. doi: 10.1371/journal.pone.0014142.
16. Longhese MP, Bonetti D, Manfrini N, Clerici M. 2010. Mechanisms and regulation of DNA end resection. *EMBO J.* 29:2864-74. doi: 10.1038/emboj.2010.165.
17. Bonetti D, Clerici M, Anbalagan S, Martina M, Lucchini G, Longhese MP. 2010. Shelterin-like proteins and Yku inhibit nucleolytic processing of *Saccharomyces cerevisiae* telomeres. *PLoS Genet.* 6:e1000966. doi: 10.1371/journal.pgen.1000966.
18. Donnianni RA1, Ferrari M, Lazzaro F, Clerici M, Tamilselvan Nachimuthu B, Plevani P, Muzi-Falconi M, Pelliccioli A. 2010. Elevated levels of the polo kinase Cdc5 override the Mec1/ATR checkpoint in budding yeast by acting at different steps of the signaling pathway. *PLoS Genet.* 6:e1000763. doi: 10.1371/journal.pgen.1000763.

Selected publications before 2010

1. Bonetti D, Martina M, Clerici M, Lucchini G, Longhese MP. 2009. Multiple pathways regulate 3' overhang generation at *S. cerevisiae* telomeres. *Mol Cell.* 35:70-81. doi: 10.1016/j.molcel.2009.05.015.
2. Clerici M*, Mantiero D, Guerini I, Lucchini G, Longhese MP*. 2008. The Yku70-Yku80 complex contributes to regulate double-strand break processing and checkpoint activation during the cell cycle. *EMBO Rep.* 9:810-8. doi: 10.1038/embor.2008.121.
3. Mantiero D**, Clerici M**, Lucchini G, Longhese MP. 2007. Dual role for *Saccharomyces cerevisiae* Tel1 in the checkpoint response to double-strand breaks. *EMBO Rep.* 8(4):380-7.
4. Clerici M, Mantiero D, Lucchini G, Longhese MP. 2006. The *Saccharomyces cerevisiae* Sae2 protein negatively regulates DNA damage checkpoint signalling. *EMBO Rep.* 7:212-8.
5. Clerici M, Mantiero D, Lucchini G, Longhese MP. 2005. The *Saccharomyces cerevisiae* Sae2 protein promotes resection and bridging of double strand break ends. *J Biol Chem.* 280:38631-8.
6. Clerici M, Baldo V, Mantiero D, Lottersberger F, Lucchini G, Longhese MP. 2004. A Tel1/MRX-dependent checkpoint inhibits the metaphase-to-anaphase transition after UV irradiation in the absence of Mec1. *Mol Cell Biol.* 24:10126-44.
7. Clerici M, Paciotti V, Baldo V, Romano M, Lucchini G, Longhese MP. 2001. Hyperactivation of the yeast DNA damage checkpoint by TEL1 and DDC2 overexpression. *EMBO J.* 20:6485-98.
8. Paciotti V**, Clerici M**, Scotti M, Lucchini G, Longhese MP. 2001. Characterization of mec1 kinase-deficient mutants and of new hypomorphic mec1 alleles impairing subsets of the DNA damage response pathway. *Mol Cell Biol.* 21:3913-25.
9. Paciotti V, Clerici M, Lucchini G, Longhese MP. 2000. The checkpoint protein Ddc2, functionally related to *S. pombe* Rad26, interacts with Mec1 and is regulated by Mec1-dependent phosphorylation in budding yeast. *Genes Dev.* 14:2046-59.

* Corresponding authors

** The authors contribute equally to these works