

## CURRICULUM VITAE

Titia de Lange  
 Born: November 11, 1955 in Rotterdam, The Netherlands  
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### Address

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### Education

1981 Master's equivalent: Doctoraal examen, highest honors. Univ. Amsterdam and NIMR, Mill Hill, UK. Thesis: Chromatin structure of the human  $\beta$ -globin gene locus. Advisor: Richard A. Flavell  
 1981 - 1985 Ph.D. Biochemistry, cum laude. Univ. of Amsterdam and The Netherlands Cancer Institute, Amsterdam. Thesis: Activation and transcription of surface antigen genes in trypanosomes. Advisor: Piet Borst

### Appointments

1981 - 1985 Graduate teaching assistant, Department of Biochemistry, Univ. of Amsterdam  
 1985 - 1990 Postdoctoral fellow with Harold E. Varmus at UCSF  
 1990 - 1991 University Fellow, Rockefeller University  
 1991 - 1994 Assistant Professor, Rockefeller University  
 1994 - 1997 Associate Professor, Rockefeller University  
 1997 - Professor, Rockefeller University  
 1999 - Leon Hess Professor, Rockefeller University  
 2006- 2011 Associate Director, Anderson Center for Cancer Research, Rockefeller University  
 2010- American Cancer Society Research Professor  
 2011- Director, Anderson Center for Cancer Research, Rockefeller University

### Honors and Awards

1980 Dr. Catharine van Tussenbroek Award  
 1985 - 1987 Christiaan and Constantijn Huygens Award  
 1987 - 1995 Lucille P. Markey Trust Scholar Award  
 1993 - 1998 The Irma T. Hirschl - Monique Weill-Caulier Trust Award  
 1995 - 2000 Rita Allen Award  
 1997 - 2002 Burroughs Wellcome Toxicology Scholar Award  
 1997 - 1999 New York Community Trust Cancer Research Award  
 2000 - 2004 Ellison Medical Foundation Senior Scholar Award  
 2000 - 2001 President of the Harvey Society  
 2000 Elected Member and Correspondent of the Royal Dutch Academy of Sciences  
 2001 Elected Foreign Member of the European Molecular Biology Organization  
 2001 Paul Marks Prize for Cancer Research (with S. Elledge, W. Kaelin, and X. Wang)  
 2003 Honorary Degree, University of Utrecht, The Netherlands  
 2004 AACR Charlotte Friend Memorial Award

2005	NIH MERIT award (GM)
2005	Elected Fellow of the New York Academy of Sciences
2005	NIH Director Pioneer Award
2006	Elected Foreign Associate of the National Academy of Sciences, USA
2006	Elected Fellow of the American Society for Microbiology
2007	Elected Fellow of the American Academy for Arts and Sciences
2007	Elected Fellow of the American Association for the Advancement of Science
2007	Rockefeller University Award for Excellence in Teaching
2008	Third Annual Massachusetts General Hospital Cancer Center Prize
2010	AACR G.H.A. Clowes Memorial Award
2010	Elected into Institute of Medicine of the National Academy of Sciences, USA
2011	Vilcek Prize in Biomedical Sciences
2012	Rosalind E. Franklin Award, National Cancer Institute
2012	Vanderbilt Prize in Biomedical Science
2012	Dr. H.P. Heineken Prize for Biochemistry and Biophysics, Royal Netherlands Academy for Arts and Sciences
2013	Breakthrough Prize in Life Science
2013	Elected Foreign Member of the Koninklijke Hollandsche Maatschappij der Wetenschappen (the second Royal Dutch Academy)
2013	Katharine Berkan Judd Award, Memorial Sloan-Kettering Cancer Center
2013	Jill Rose Award, Breast Cancer Research Foundation
2014	Elected Fellow of the AACR Academy
2014	Canada Gairdner International Award
2015	Honorary Degree, University of Chicago

#### **Keynote and named lectures**

2000	Keynote, "Telomeres and Telomerase" meeting, Montreal, Canada
2001	Keynote, "ADPR 2001" meeting, NYC
2001	Keynote, "Genes and Cancer" meeting Warwick, UK
2002	Keynote, Fifth Annual Cell and Dev. Biol. Symp., U. Penn.
2004	Keynote, Dutch Tumor Biology Meeting
2004	Keynote, IMP meeting, Vienna
2004	Keynote, AACR meeting on Role of Telomeres and Telomerase in Cancer
2005	Reginal Harris Lecturer. CSH Symp. Quant. Biol.
2006	EMBO Lecture; Telomeres and the DNA damage response, Villars, Switzerland
2006	Tolmach Lecture; Telomeres and DNA repair Symposium Wash. U, St. Louis
2007	Keynote, Gordon Conference on Mammalian DNA Repair
2007	EMBO Lecture; 32 <sup>nd</sup> Ann. FEBS Congress. Vienna
2007	Frederick Ruysch Lecture, Amsterdam Medical Center
2007	Mendel Lecture, Brno, Czech Republic
2008	Keynote, DNA Repair Symposium, Univ. Pittsburgh
2009	Keynote, Keystone Symposium on Genome Instability and DNA repair
2009	Rupert E. Billingham Lecture, UTSW
2009	Alexander Cruickshank Lecture, Gordon Conference on Chromosome Dynamics
2010	Harvey Lecture, Harvey Society, New York
2010	Osborne Lecture, Univ. Connecticut
2010	Marguerite Vogt Lecture, Salk Institute
2010	"Speaker of the year" lectures, Netherlands Soc. Biochem. Mol. Biol.
2011	Rose Winer Levin Lecture, Dana Farber Cancer Institute
2011	Keynote, Gordon Conference on Cell Growth and Proliferation
2012	Cancer Biology Distinguished Lecture, Rockefeller University
2012	Keynote, EMBO meeting on Telomeres and the DNA damage response
2012	Keynote, Gordon Conference on Mutagenesis
2013	Distinguished Cancer Researcher Lecture, USCF
2013	Cynthia and Alexander Tseng, Jr., MD Memorial Lecture, Stanford University

2014 Don W. Fawcett Lectures, Department Cell Biology, Harvard Medical School  
 2014 Nicholson Lecture, Nobel Forum, Karolinska Institute  
 2014 Jean Weigle Lecture, Univ. Geneva  
 2014 Keynote, EMBO meeting on Telomeres  
 2016 Cyma Rubin Lecture, Weill Cornell Medical School  
 2016 George Palade Lecture, Yale  
 2016 Keynote, Abcam meeting on Recombination  
 2016 Keynote, Boston Mitosis meeting, Whitehead Institute, MIT

### Scientific advisory boards and selection committees

2001- Board of Scientific Consultants of Memorial Sloan-Kettering Cancer Center  
 2001- External Advisory Committee for the Endowed Scholars Program in Biomedical Sciences at UT Southwestern Medical School at Dallas  
 2002-2009 Trustee, Cold Spring Harbor Laboratory, LI  
 2002-2008 Scientific Advisory Board, Institute for Molecular Pathology, Vienna  
 2003-2011 Scientific Advisory Board, MIT Cancer Center  
 2003-2009 Paul Marks Award Selection Committee (Chair 2009)  
 2003- Pearl Meister Greengard Prize Selection Committee  
 2004 Overseers' Committee for Dept. MCB at Harvard  
 2004 Search committee for Director of IMP, Vienna  
 2005-2012 Scientific Advisory Board, CRUK, Clare Hall and Lincoln's Inn Field (LRI)  
 2005- Lasker Award Jury  
 2005- Scientific Advisory Board, Dutch Cancer Institute  
 2006- Executive Committee of the Starr Cancer Consortium  
 2008- Advisory Council, Dep. Mol. Biol., Princeton University  
 2009 W.M. Keck Foundation, Subcommittee Evaluation Cell Biology  
 2010-2013 AACR Special Conference Committee  
 2011-2014 Scientific Advisory Committee, Ludwig Institute for Cancer Research  
 2011- Selection Committee Vilcek Prize in Biomedical Sciences  
 2012- Biomedical Scientific Advisory Board, Vanderbilt University  
 2012- Executive Committee American Italian Cancer Foundation  
 2013 Search committee for Director of IMP, Vienna  
 2013 External advisor, search committee for Director of SKI, MSKCC  
 2013- Wiley Prize Jury  
 2014- Selection Committee NAS Award in Molecular Biology  
 2015- Board member, Vilcek Foundation

### Editorial Boards

1997- Molecular and Cellular Biology  
 2000- Trends in Biological Science  
 2004 -2007 PLoS Biology  
 2008 - Genes and Development  
 2009- Nucleus  
 2010- Journal of Cell Biology  
 2011- Current Opinion in Genetics and Development

**BIBLIOGRAPHY (Reviews, book chapters, and commentaries in italics.)**

1. S. Hueting, **T. de Lange**, D. Tempest (1978) Properties and regulation of synthesis of the glycerol dehydrogenase present in *Klebsiella aerogenes* NCTC 418, growing in chemostat culture. FEMS 4: 195-198.
2. S. Hueting, **T. de Lange**, D.W. Tempest (1979) Energy requirement for maintenance of the transmembrane potassium gradient in *Klebsiella aerogenes* NCTC 418: a continuous culture study. Arch. Microbiol. 123: 183-188.
3. F. van der Mark, **T. de Lange**, H.F. Bienfait (1981) The role of ferritin in developing primary bean leaves under various light conditions. Planta 153: 338-342.
4. **T. de Lange** and P. Borst (1982) Genomic environment of the expression-linked extra copies of genes for surface antigens of *Trypanosoma brucei* resembles the end of the chromosome. Nature 299: 451-453.
5. L.H.T. van der Ploeg, D. Valerio, **T. de Lange**, A. Bernards, P. Borst, F.G. Grosveld (1982) An analysis of cosmid clones of nuclear DNA from *Trypanosoma brucei* shows that the genes for Variant Surface Glycoprotein are clustered in the genome. Nucl. Acids Res. 10: 5905-5923.
6. L.H.T. van der Ploeg, A.Y.C. Liu, P.A.M. Michels, **T. de Lange**, P. Borst, H.K. Majumder, H. Weber, G.H. Veeneman, J.C. van Boom (1982) RNA splicing is required to make the messenger RNA for a Variant Surface Glycoprotein in trypanosomes. Nucl. Acids Res. 10: 3591-3604.
7. **T. de Lange**, A.Y.C. Liu, L.H.T. van der Ploeg, P. Borst, M.C. Tromp, J.C. van Boom (1983) Tandem repetition of the 5' mini-exon of Variant Surface Glycoprotein genes. Cell 34: 891-900.
8. **T. de Lange**, J.M. Kooter, P.A.M. Michels, P. Borst (1983) Telomere conversion in trypanosomes. Nucl. Acids Res. 11: 8149-8165.
9. P. Borst, A. Bernards, L.H.T. van der Ploeg, P.A. Michels, A.Y. Liu, **T. de Lange**, J.M. Kooter (1983) The control of variant surface antigen synthesis in trypanosomes. Eur. J. Biochem. 137: 383-389.
10. D. Kioussis, D. Vanin, **T. de Lange**, R.A.F. Flavell, F.G. Grosveld (1983)  $\beta$ -globin gene inactivation by DNA translocation in  $\gamma/\beta$ -thalassemia. Nature 306: 662-666.
11. A. Bernards, **T. de Lange**, P.A.M. Michels, M.J. Huisman, P. Borst (1984) Two modes of activation of a single surface antigen gene of *Trypanosoma brucei*. Cell 36: 163-170.
12. **T. de Lange**, P.A.M. Michels, H.J.G. Veerman, A.W.C.A. Cornelissen, P. Borst (1984) Many trypanosome messenger RNAs share a common 5' terminal sequence. Nucl. Acids Res. 12: 3777-3790.
13. J.M. Kooter, **T. de Lange**, P. Borst (1984) Discontinuous transcription in trypanosomes. EMBO J. 3: 2387-2392.
14. T. de Lange, T.M. Berkvens, H.G. Veerman, A.C.C. Frasch, J.D. Barry, P. Borst (1984) Comparison of the genes coding for the common 5' terminal sequence of messenger RNAs in three trypanosome species. Nucl. Acids Res. 12: 4431-4443.
15. **T. de Lange**, J.M. Kooter, J. Luirink, P. Borst (1985) Transcription of a transposed trypanosome surface antigen gene starts upstream of the transposed segment. EMBO J. 4: 3299-3306.
16. A. Bernards, L.H.T. van der Ploeg, W.C. Gibson, P. Leegwater, F. Eijgenraam, **T. de Lange**, P. Weyers, J. Calafat, P. Borst (1986). Rapid change of the repertoire of Variant Surface Glycoprotein genes in trypanosomes by gene duplication and deletion. J. Mol. Biol. 190: 1-10.
17. M. Imboden, B. Blum, **T. de Lange**, R. Braun, T. Seebeck (1986) The tubulin mRNAs of *Trypanosoma brucei*. J. Mol. Biol. 188: 393-402.
18. **T. de Lange** (1986) *The molecular biology of antigenic variation in trypanosomes: Gene rearrangements and discontinuous transcription.* Intern. Rev. Cytol. 99: 85-113.
19. M. Timmers, **T. de Lange**, J.M. Kooter, P. Borst (1987) Coincident multiple activations of the same surface antigen gene in *Trypanosoma brucei*. J. Mol. Biol. 194: 81-90.
20. J. Stone, **T. de Lange**, G. Ramsay, E. Jacobovits, J.M. Bishop, H.E. Varmus, W.M.F. Lee (1987) Definition of regions in human c-myc that are involved in transformation and nuclear localization. Mol. Cell. Biol. 7: 1697-1709.

21. **T. de Lange**, L. Shiue, R.M. Myers, D.R. Cox, S.L. Naylor, A.M. Killery, H.E. Varmus (1990) Structure and variability of human chromosome ends. *Mol. Cell. Biol.* 10: 518-527.
22. M. Burmeister, S.W. Kim, E.R. Price, **T. de Lange**, U. Tantravahi, R.M. Myers, D.R. Cox (1991) A map of the distal long arm of human chromosome 21, constructed by using radiation hybrids and pulsed field gel electrophoresis. *Genomics* 9: 19-30.
23. **T. de Lange** (1992) Human telomeres are attached to nuclear matrix. *EMBO J.* 11: 717-724.
24. Z. Zhong, L. Shiue, S. Kaplan, T. de Lange (1992) A mammalian factor that binds telomeric TTAGGG repeats in vitro. *Mol. Cell. Biol.* 12: 4834-4843.
25. D. Saltman, R. Morgan, M. Cleary, **T. de Lange** (1993) Telomeric structure in cells with chromosome end associations. *Chromosoma* 102: 121-128.
26. M.E. Cardenas, A. Bianchi, **T. de Lange** (1993) A *Xenopus* egg factor with DNA-binding properties characteristic of terminus-specific telomeric proteins. *Genes and Development*. 7: 883-894.
27. H. Tommerup, A. Dousmanis, **T. de Lange** (1994) Unusual chromatin in human telomeres. *Mol. Cell. Biol.* 14: 5777-5785.
28. **T. de Lange** (1994) Activation of telomerase in a human tumor. *Proc. Natl. Acad. Sci. USA* 91: 2882-2885.
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30. D. Broccoli, J.W. Young, **T. de Lange** (1995) Telomerase activity in normal and malignant hematopoietic cells. *Proc. Natl. Acad. Sci. USA* 92: 9082-9086.
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32. **T. de Lange** (1995) *Telomere dynamics and genome instability in human cancer. In: Telomeres. E.H. Blackburn and C.W. Greider, eds. CSH press, pp. 265-293.*
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39. B. van Steensel and **T. de Lange** (1997) Control of telomere length by the human telomeric protein TRF1. *Nature* 385: 740-744.
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42. J. Griffith, A. Bianchi, **T. de Lange** (1998) TRF1 promotes parallel pairing of telomeric DNA in vitro. *J. Mol. Biol.* 278: 79-88.
43. B. van Steensel, A. Smogorzewska, **T. de Lange** (1998) TRF2 protects human telomeres from end-to-end fusions. *Cell* 92: 401-413.
44. **T. de Lange** (1998) *Telomeres and senescence: ending the debate. Science* 279: 334-335.
45. X. Dong, M.A. Michelis, J. Wang, R. Bose, **T. de Lange**, W.H. Reeves (1998) Autoantibodies to DEK oncoprotein in a patient with systemic lupus erythematosus and sarcoidosis. *Arthritis Rheum.* 41:1505-1510.
46. S. Smith, I. Gariat, A. Schmitt, **T. de Lange** (1998) Tankyrase, a poly(ADP-ribose)polymerase at human telomeres. *Science* 282: 1484-1488.

47. L. Zhu, S. Smith, **T. de Lange**, M. F. Seldin. (1999) Chromosomal mapping of the Tankyrase gene in human and mouse. Genomics 57: 320-321.
48. **T. de Lange** and R. DePinho (1999) Unlimited mileage from telomerase? Science 283: 947-949.
49. J. Karlseder, D. Broccoli, Y. Dai, S. Hardy, **T. de Lange** (1999) p53- and ATM-dependent apoptosis induced by telomeres lacking TRF2. Science 283: 1321-1325.
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51. A. Bianchi and **T. de Lange** (1999) Ku binds telomeric DNA in vitro. J. Biol. Chem. 274: 21223-21227.
52. S. Smith and **T. de Lange** (1999) Cell cycle dependent localization of a telomeric PARP, tankyrase, to centrosomes and nuclear pore complexes. J. Cell Sci. 112: 3649-3656.
53. **T. de Lange** and T. Jacks (1999) For better or worse? *Telomerase inhibition and cancer*. Cell 98: 273-275.
54. A. Bianchi, R. M. Stansel, L. Fairall, J. D. Griffith, D. Rhodes, **T. de Lange** (1999) TRF1 binds a bipartite telomeric site with extreme spatial flexibility. EMBO J. 18: 5735-5744.
55. A. Smogorzewska, B. van Steensel, A. Bianchi, G. Schnapp, M. R. Schaefer, S. Oelmann, **T. de Lange** (2000) Control of human telomere length by TRF1 and TRF2. Mol. Cell Biol. 20: 1659-1668.
56. B. Li, S. Oestreich, and **T. de Lange** (2000) Identification of Human Rap1: Implications for Telomere Evolution. Cell 101: 471-483.
57. X.-D. Zhu, B. Küster, M. Mann, J.H.J. Petrini, **T. de Lange** (2000) Cell cycle regulated association of the Rad50/Mre11/Nbs1 complex with TRF2 and human telomeres. Nature Genet. 25: 347-352.
58. **T. de Lange** and J.H.J. Petrini. (2000) A new connection at human telomeres: association of the Mre11 complex with TRF2. CSH Symp. Quant. Biol. Vol LXV: 265-273.
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73. **T. de Lange** (2003) *Protection and maintenance of human telomeres. Encyclopedia of the Human Genome. D. Cooper, editor. Nature Publishing group. McMillan press.*
74. B. Li and **T. de Lange** (2003) Rap1 affects the length and heterogeneity of human telomeres. Mol. Biol. Cell 14: 5060-5068. Epub 2003 Oct 17.
75. X.-D. Zhu, L. Niedernhofer, B. Kuster, M. Mann, J. H. J. Hoeijmakers, **T. de Lange** (2003) ERCC1/XPF removes the 3' overhang from uncapped telomeres and represses formation of telomeric DNA containing double minute chromosomes. Mol. Cell 6: 1489-1498.
76. D. Loayza\*, H. Parsons\*, J. Donigian, L. Hoke, **T. de Lange** (2004) DNA binding features of human POT1: A nonamer 5'-TAGGGTTAG-3' minimal binding site, sequence specificity, and internal binding to multimeric sites. J. Biol. Chem. 279: 13241-13248. Epub 2004 Jan 7.
77. J. Z.-S. Ye and **T. de Lange** (2004) TIN2 is a tankyrase 1 PARP-modulator in the TRF1 telomere length control complex. Nature Genet. 36: 618-623. Epub 2004 May 9.
78. A. Smogorzewska and **T. de Lange** (2004) *Regulation of telomerase by telomeric proteins. Ann. Rev. Biochem.* 73: 177-208.
79. **T. de Lange** (2004) *T-loops and the origin of telomeres. Nature Rev. Mol. Cell Biol.* 5: 323-329.
80. D. Loayza and **T. de Lange** (2004) *Telomerase regulation at the telomere: a binary switch. Cell* 117: 279-280.
81. J. Z.-S. Ye, D. Hockemeyer, A. Krutchinsky, D. Loayza, S. Hooper, B. Chait, **T. de Lange** (2004) POT1 Interacting Protein PIP1: a telomere length regulator that recruits POT1 to the IN2/TRF1 complex. Genes and Development 18: 1649-1654. Epub 2004 Jul 1.
82. J. Karlseder, K. Hoke, O. Mirzoeva, C. Bakkenist, M. Kastan, J.H.J. Petrini, **T. de Lange** (2004) The telomeric protein TRF2 binds the ATM kinase and can inhibit the ATM-dependent DNA damage response. PLoS Biology 2: 1150-1157. Epub 2004 Aug 17.
83. J. Silverman, H. Takai, S.B.C. Buonomo, F. Eisenhaber, **T. de Lange** (2004) Human Rif1, ortholog of a yeast telomeric protein, is regulated by ATM and 53BP1 and functions in the S-phase checkpoint. Genes and Development 18: 2108-2119.
84. J.Z.-S. Ye\*, J. Donigian\*, M. van Overbeek\*, D. Loayza, B. Chait, A. Krutchinsky, **T. de Lange** (2004) TIN2 binds TRF1 and TRF2 simultaneously and stabilizes the TRF2 complex on telomeres. J. Biol. Chem., 279: 47264-47271. Epub 2004 Aug 17.
85. R. Wang, A. Smogorzewska, **T. de Lange** (2004) Homologous recombination generates t-loop sized deletions at human telomeres. Cell 119: 355-368. (cover)
86. J.J.L. Jacobs and **T. de Lange** (2004) Significant role for p16<sup>INK4a</sup> in p53-independent telomere-directed senescence. Curr. Biol. 14: 1-20.
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88. G. Celli and **T. de Lange** (2005) DNA processing not required for ATM activation or the telomere damage response after conditional deletion of mouse TRF2. Nature Cell Biol 7: 712-718. (cover) Epub 2005 Jun 19.
89. **T. de Lange** (2005). *Shelterin: the protein complex that shapes and safe-guards human telomeres. Genes and Development* 19: 2100-2110.
90. **T. de Lange** (2005) *Mammalian telomeres. In "Telomeres" T. de Lange, V. Lundblad, and E.H. Blackburn eds., CSH press.*
91. J.J.L. Jacobs and **T. de Lange** (2005) *p16INK4as a second effector of the telomere damage response. Cell Cycle* 4: 1364-1368. Epub 2005 Oct 17.
92. **T. de Lange** (2005) *Telomere-related genome instability in cancer. CSH Symp. Quant. Biol.* 70: 197-204.
93. G.B. Celli, E. Lazzarini Denchi, **T. de Lange** (2006) Ku70 stimulates fusion of dysfunctional telomeres yet protects chromosome ends from homologous recombination. Nature Cell Biol. 8: 855-862. Epub 2006 July 16.
94. M. van Overbeek and **T. de Lange** (2006) Apollo, an Artemis related nuclease, interacts with TRF2 and protects human telomeres in S phase. Curr. Biol. 16: 1295-1302. Epub 2006 May 25.
95. D. Hockemeyer, J.-P. Daniels, H. Takai, **T. de Lange** (2006) Recent expansion of the telomeric complex in rodents: two distinct POT1 proteins protect mouse telomeres. Cell 126: 63-77.

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97. N. Dimitrova and **T. de Lange** (2006) MDC1 accelerates nonhomologous end-joining of dysfunctional telomeres. Genes and Development 20: 3238-3243.
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