

CURRICULUM VITAE

Michael W. Young  
Nobel Laureate in Physiology or Medicine (2017)

Birth: March 28, 1949, Miami, Florida

Married: Laurel A. Eckhardt; two children

Education: The University of Texas, Austin B.A. Biology 1971  
The University of Texas, Austin Ph.D. Genetics 1975

Address: Laboratory of Genetics,  
The Rockefeller University, 1230, York Avenue,  
New York, New York 10021. Tel. (212) 327-8645.

Positions:

Postdoctoral Fellow, Department of Biochemistry, Stanford  
University School of Medicine, Palo Alto, CA 1975-1977

Assistant Professor, Meyer Foundation Fellow, The Rockefeller  
University, New York, NY 1978-1984

Associate Professor, The Rockefeller University 1984-1988

Investigator, Howard Hughes Medical Institute 1987-1996

Professor, Rockefeller University 1988-

Head, Rockefeller Unit, National Science Foundation Science  
and Technology Center for Biological Timing 1991-2001

Vice President for Academic Affairs, Rockefeller University 2004-

Richard and Jeanne Fisher Professor, The Rockefeller  
University 2004-

Selected Committees:

Member, Advisory Panel for Genetic Biology, National  
Science Foundation 1984-1988

Member, Editorial Board, Molecular and Cellular Biology 1986-1991

Member, Research Advisory Committee, National Science  
Foundation Science and Technology Center for  
Biological Timing 1991-

Member, Cell Biology Study Section, National Institutes  
of Health 1993-1997

<u>NYSTAR</u> , Evaluation Committee for Capital Facility and Faculty Development Programs in New York State	2001-2002
Member, Harvey Society Council	1999-2004
Treasurer, Harvey Society	2001-2004
Associate Editor/Advisory Board, <u>Journal of Biological Rhythms</u>	2000-
Associate Editor, <u>Neuron</u>	2000-2016
Co-organizer, Fondation des Treilles Conference, <u>Cellular Mechanisms of Circadian Clocks</u> , Nice, Fr.	2000
Organizer, Keystone Symposium, <u>Molecular Clocks</u> , Tahoe, CA	2001
Chairman, External Review Panel, Life Sciences Task Force Program of Excellence, Texas A&M University	2003
Member, Neurodifferentiation, Plasticity, Regeneration and Rhythmicity Study Section, National Institutes of Health	2012-2016

University Service:

Member, Executive Committee of the Faculty, The Rockefeller University	1990-1993, 2000-2003
Chairman, Faculty Senate, The Rockefeller University	1992-1993 2002-2003
Chairman, Faculty Nominating Committee, The Rockefeller University	1996-2003

Selected Honors and Awards:

Hayaishi Prize, University of Tokyo	2001
Ashoff/Pittendrigh Award for Pioneering Contributions to the Field of Circadian Biology, Society for Research on Biological Rhythms	2006
Member, National Academy of Sciences, USA	2007
Fellow, American Academy of Microbiology	2007
Neuroscience Prize, Peter and Patricia Gruber Foundation	2009
Louisa Gross Horwitz Prize, Columbia University	2011
Canada Gairdner International Award	2012
Massry Prize, USC Keck School of Medicine	2012
Wiley Prize in Biomedical Sciences	2013
Shaw Prize in Life Science and Medicine	2013
Nobel Prize in Physiology or Medicine	2017

Memberships:

National Academy of Sciences, USA  
 American Academy of Microbiology  
 Genetics Society of America  
 Society for Research on Biological Rhythms  
 New York Academy of Sciences  
 American Society for Microbiology  
 Harvey Society

Selected Publications

Judd, B. H. and Young, M. W. (1974). An examination of the one cis-tron: one chromomere concept. Cold Spring Harbor Symp. Quant. Biol. 38: 573-579.

Finnegan, D. J., Rubin, G. M., Young, M. W., and Hogness, D. S. (1978). Repeated gene families in *Drosophila melanogaster*. Cold Spring Harbor Symp. Quant. Biol. 42: 1053-1063.

Young M. W., Judd B. H. (1978). Nonessential Sequences, Genes, and the Polytene Chromosome Bands of *DROSOPHILA MELANOGASTER*. Genetics 88: 723-742.

Young, M. W. (1979). Middle repetitive DNA: A fluid component of the *Drosophila* genome. Proc. Natl. Acad. Sci. U. S. A. 76: 6274-6278.

Dowsett, A. P. and Young, M. W. (1982). Differing levels of dispersed, repetitive DNA among closely related species of *Drosophila*. Proc. Natl. Acad. Sci. U. S. A. 79: 4570-4574.

Kidd, S., Lockett, T. J., and Young, M. W. (1983). The *Notch* locus of *Drosophila melanogaster*. Cell 34: 421-433.

Bargiello, T. A. and Young, M. W. (1984). Molecular genetics of a biological clock in *Drosophila*. Proc. Natl. Acad. Sci. U. S. A. 81: 2142-2146.

Bargiello, T.A., Jackson, F.R., and Young, M.W. (1984). Restoration of circadian behavioural rhythms by gene transfer in *Drosophila*. Nature 312: 752-754.

Shin, H. -S., Bargiello, T. A., Clark, B. T., Jackson, F. R., and Young, M. W. (1985). An unusual coding sequence from a *Drosophila* clock gene is conserved in vertebrates. Nature 317: 445-448.

Young, M. W., Jackson, F. R., Shin, H. -S., and Bargiello, T. A. (1985). A biological clock in *Drosophila*. Cold Spring Harbor Symp. Quant Biol. 50: 865-875.

Jackson, F. R., Bargiello, T. A., Yun, S. -H., and Young, M. W. (1986). Product of the *per* locus of *Drosophila* shares homology with proteoglycans. Nature 320: 185-188.

- Kidd, S. J., Kelley, M. R., and Young, M. W. (1986). Sequence of the *Notch* locus of *Drosophila melanogaster*; relationship of the encoded protein to mammalian clotting and growth factors. *Mol. Cell. Biol.* **6**: 3094-3108.
- Kidd, S. J., and Young, M. W. (1986). Transposon dependent mutant phenotypes at the *Notch* locus of *Drosophila*. *Nature* **323**: 89-91.
- Baylies, M. K., Bargiello, T. A., Jackson, F. R., and Young, M. W. (1987). Changes in the abundance or structure of the *per* gene product can affect the periodicity of the *Drosophila* clock. *Nature* **326**: 390-392.
- Kelley, M. R., Kidd, S. J., Deutsch, W. A., and Young, M. W. (1987). Mutations altering the structure of Epidermal Growth Factor-like coding sequences at the *Drosophila Notch* locus. *Cell* **51**: 539-548.
- Kidd, S. J., Baylies, M. K., Gasic, G. P., and Young, M. W. (1989). Structure and distribution of the *Notch* protein in developing *Drosophila*. *Genes and Development* **3**:1113-1129.
- Corbin, V., Michelson, A. M., Abmayr, S. M., Neel, V., Alcamo, E., Maniatis, T., and Young, M. W. (1991). A role for the *Drosophila* neurogenic genes in mesoderm differentiation. *Cell*, **67**: 311-323.
- Sehgal, A., Price, J., and Young, M. W. (1992). Ontogeny of a biological clock in *Drosophila melanogaster*. *Proc. Natl. Acad. Sci., USA*, **89**: 1423-1427.
- Baylies, M.K., Sehgal, A., and Young, M.W. (1992). New short period mutations of the *Drosophila* clock gene *per*. *Neuron*, **9**: 575-581.
- Lieber, T., Wesley, C. S., Alcamo, E., Hassel, B., Krane, J.F., Campos-Ortega, J.A., and Young, M.W. (1992). Single amino acid substitutions in EGF-like elements of *Notch* and *Delta* modify *Drosophila* development and affect cell adhesion *in vitro*. *Neuron*, **9**: 847-859.
- Molecular Genetics of Biological Rhythms. (1992). Young, M. W., ed. Marcel Dekker, New York, 319 pp.
- Lieber, T., Kidd, S., Alcamo, E., Corbin, V., and Young, M.W. (1993). Anti-neurogenic phenotypes induced by truncated Notch proteins indicate a role in signal transduction, and may point to a novel function for Notch in nuclei. *Genes & Dev.*, **7**: 1949-1965.
- Lyman, D. and Young, M.W. (1993). Further evidence for function of the *Drosophila* Notch protein as a transmembrane receptor. *Proc. Natl. Acad. Sci. U.S.A.*, **90**: 10395-10399.
- Sehgal, A., Price, J.L., Man, B., and Young, M.W. (1994). Loss of circadian behavioral rhythms and *per* RNA oscillations in the *Drosophila* mutant *timeless*. *Science*, **263**: 1603-1606.
- Vosshall, L.B., Price, J.L., Sehgal, A., Saez, L., and Young, M.W. (1994). Block in nuclear localization of *period* protein by a second clock mutation, *timeless*. *Science*, **263**: 1606-1609.
- Neel, V. A. and Young, M.W. (1994). *igloo*, a GAP-43-related gene expressed in the developing nervous system of *Drosophila*. *Development*, **120**: 2235-2243.

Vosshall, L.B. and Young, M.W. (1995). Circadian Rhythms in *Drosophila* can be driven by *period* expression in a restricted group of central brain cells. *Neuron*, 15: 345-360.

Price, J.L., Dembinska, M.E., Young, M.W. and Rosbash, M. (1995). Suppression of PERIOD protein abundance and circadian cycling by the *Drosophila* clock mutation *timeless*. *EMBO J.*, 14: 4044-4049.

Myers, M.P., Wager-Smith, K., Wesley, C.S., Young, M.W. and Sehgal, A. (1995). Positional cloning and sequence analysis of the *Drosophila* clock gene, *timeless*. *Science*, 270: 805-808.

Sehgal, A., Rothenfluh-hilfiker, A., Hunter-Ensor, M., Chen, Y., Myers, M.P., and Young, M.W. (1995). Rhythmic expression of *timeless*: A basis for promoting circadian cycles in *period* gene autoregulation. *Science*, 270: 808-810.

Gekakis, N., Saez, L., Delahaye-Brown, A.-M., Myers, M.P., Sehgal, A., Young, M.W., and Weitz, C.J. (1995). Isolation of *timeless* by PER protein interaction: Defective interaction between *timeless* protein and long-period mutant PER<sup>L</sup>. *Science*, 270: 811-815.

Myers, M.P., Wager-Smith, K., Rothenfluh-Hilfiker, A., and Young, M.W. (1996). Light-induced degradation of TIMELESS and entrainment of the *Drosophila* circadian clock. *Science*, 271: 1736-1740.

Saez, L. and Young, M.W. (1996). Regulation of nuclear entry of the *Drosophila* clock proteins PERIOD and TIMELESS. *Neuron*, 17: 911-920.

Young, M.W., Wager-Smith, K., Vosshall, L.B., Saez, L., and Myers, M.P. (1996). Molecular anatomy of a light-sensitive circadian pacemaker in *Drosophila*. *Cold Spring Harbor Symp. Quant. Biol.* 61: 279-284.

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Price, J.L., Blau, J., Rothenfluh, A., Abodeely, M., Kloss, B, and Young, M.W. (1998). *double-time* is a novel *Drosophila* clock gene that regulates PERIOD protein accumulation. *Cell*, 94: 83-95.

Kloss, B., Price, J.L., Saez, L., Blau, J., Rothenfluh, A., Wesley, C.S., and Young, M.W. (1998). The *Drosophila* clock gene *double-time* encodes a protein closely related to human casein kinase I $\epsilon$ . *Cell*, 94: 97-107.

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Kidd, S., Lieber, T., and Young, M.W. (1998). Ligand-induced cleavage and regulation of nuclear entry of Notch in *Drosophila melanogaster* embryos. *Genes & Dev.*, 12: 3728-3740.

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Young, M.W. (2000). The tick-tock of the biological clock. *Scientific American*, 282: 64-71.

Young, M.W. (2000). Circadian rhythms: Marking time for a kingdom. *Science*, 288: 451-453.

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Rothenfluh, A., Abodeely, M., Price, J.L., and Young, M.W. (2000). Isolation and analysis of six *timeless* alleles which cause short- or long-period circadian rhythms in *Drosophila*. *Genetics*, 156:665-675.

Martinek, S. and Young, M.W. (2000). Specific genetic interference with behavioral rhythms in *Drosophila* by expression of inverted-repeats. *Genetics*, 156: 1717-1725.

Rothenfluh, A., Abodeely, and Young, M.W. (2000). Short-period mutations of *per* affect a *double-time*-dependent step in the *Drosophila* circadian clock. *Current Biology*, 10: 1399-1402.

Young, M.W. (2000). Life's 24 hour clock: Molecular control of circadian rhythms in animal cells. *Trends in Biochem. Sci.* 25<sup>th</sup> Anniversary, 25: 601-606.

Martinek, S., Inonog, S., Manoukian, A.S. and Young, M.W. (2001). A Role for the Segment Polarity Gene *shaggy*/GSK-3 in the *Drosophila* Circadian Clock. *Cell*, 105: 769-779.

Kloss, B., Rothenfluh, A., Young, M.W., and Saez, L. (2001). Phosphorylation of PERIOD is influenced by cycling physical associations of DOUBLE-TIME, PERIOD AND TIMELESS in the *Drosophila* clock. *Neuron* 30: 699-706.

Young, M.W. and Kay, S.A. Time zones: a comparative genetics of circadian clocks. (2001). *Nat Rev Genet.* 2: 702-715.

Claridge-Chang, A., Wijnen, H., Naef, F., Boothroyd, C., Rajewsky, N., and Young, M.W. (2001).. Circadian regulation of gene expression systems in the *Drosophila* head. *Neuron*, 32: 657-671.

Lieber, T, Kidd S, Young MW. (2002). *kuzbanian*-mediated cleavage of *Drosophila* Notch. *Genes Dev.* 16: 209-221.

Wijnen, H, Boothroyd C, Young MW, Claridge-Chang A. (2002). Molecular genetics of timing in intrinsic circadian rhythm sleep disorders. *Ann Med.* 34:386-393.

Young, M.W. (2002). Big Ben rings in a lesson on biological clocks. *Neuron* 36:1001-1005.

Cyran, S.A., Buchsbaum, A.M., Reddy, K.L., Lin, M.C., Glossop, N.R., Hardin, P.E., Young, M.W., Storti, R.V., Blau, J. (2003). *vriille*, *Pdp1* and *dClock* form a second feedback loop in the *Drosophila* circadian clock. *Cell* 112:329-341.

Harms, E., Young M.W., Saez, L. (2003). CK1 and GSK3 in the *Drosophila* and mammalian circadian clock. *Novartis Found Symp.* 253:267-277.

Young. MW. (2004). An ultradian clock shapes genome expression in yeast. *Proc Natl Acad Sci U S A.* 101:1118-1119.

Wijnen H, Naef F, Young MW. (2005). Molecular and statistical tools for circadian transcript profiling. *Methods Enzymol.* 393:341-365.

Cyran SA, Yiannoulos G, Buchsbaum AM, Saez L, Young MW, Blau J. (2005). The double-time protein kinase regulates the subcellular localization of the *Drosophila* clock protein period. *J Neurosci.* 25:5430-5437.

Meyer P, Saez L, Young MW. (2006). PER-TIM interactions in living *Drosophila* cells: an interval timer for the circadian clock. *Science* 311:226-229.

Wijnen H, Naef F, Boothroyd C, Claridge-Chang A, Young MW. (2006). Control of Daily Transcript Oscillations in *Drosophila* by Light and the Circadian Clock. *PLoS Genet* 2(3): e39.

Wijnen H. and Young MW. (2006). Interplay of Circadian Clocks and Metabolic Rhythms. *Annu. Rev. Genet.* 40:409-448.

Schwartz EC, Saez L, Young MW, Muir TW. (2007). Post-translational enzyme activation in an animal via optimized conditional protein splicing. *Nat Chem Biol.* 3: 50-54.

Boothroyd, C.E., Wijnen, H., Naef, F., Saez, L., Young, M.W. (2007). Integration of Light and Temperature in the Regulation of Circadian Gene Expression in *Drosophila*. *PLoS Genet* 3(4):e54, pp. 492-507.

Meyer P, Young MW. (2007). The 2006 Pittendrigh/Aschoff lecture: New roles for old proteins in the *Drosophila* circadian clock. *J Biol Rhythms* 22:283-290.

Sekine T, Yamaguchi T, Hamano K, Young MW, Shimoda M, Saez L. (2008). Casein kinase I epsilon does not rescue double-time function in *Drosophila* despite evolutionarily conserved roles in the circadian clock. *J Biol Rhythms.* 23:3-15.

Saez L, Meyer P, Young MW. (2008). A PER/TIM/DBT interval timer for *Drosophila*'s circadian clock. *Cold Spring Harb Symp Quant Biol.* 72:69-74.

Kivimäe S, Saez L, Young MW. (2008). Activating PER repressor through a DBT-directed phosphorylation switch. *PLoS Biol.* 6(7):e183, pp 1570-1583.

King HA, Hoelz A, Crane BR, Young MW. (2011). Structure of an enclosed dimer formed by the *Drosophila* Period protein. *J Mol Biol.* 413:561-572. Epub 2011 Sep 3.

Saez L, Derasmo M, Meyer P, Stieglitz J, Young MW. (2011). A key temporal delay in the circadian cycle of *Drosophila* is mediated by a nuclear localization signal in the Timeless protein. *Genetics* 188:591-600.

Syed S, Saez L, Young MW. (2011). Kinetics of Doubletime Kinase-dependent degradation of the *Drosophila* Period protein. *J Biol Chem.* 286:27654-27662.

Zoltowski BD, Vaidya AT, Top D, Widom J, Young MW, Crane BR. (2011). Structure of full-length *Drosophila* cryptochrome. *Nature* 480:396-399.

Stavropoulos N, Young MW. (2011). *insomniac* and Cullin-3 regulate sleep and wakefulness in *Drosophila*. *Neuron* 72:964-976.

Rogulja D, Young MW. (2012). Control of Sleep by Cyclin A and its regulator. *Science* 335: 1617-1621.

Levy C, Zoltowski BD, Jones AR, Vaidya AT, Top D, Widom J, Young MW, Scrutton NS, Crane BR, Leys D. (2013). Updated structure of *Drosophila* cryptochrome. *Nature* 495: E3-4.

Bamne MN, Ponder CA, Wood JA, Mansour H, Frank E, Kupfer DJ, Young MW, Nimgaonkar VL. (2013). Application of an ex vivo cellular model of circadian variation for bipolar disorder research: a proof of concept study. *Bipolar Disord.* 15: 694-700.

Vaidya AT, Top D, Manahan CC, Tokuda JM, Zhang S, Pollack L, Young MW, Crane BR. (2013). Flavin reduction activates *Drosophila* cryptochrome. *Proc. Natl. Acad. Sci. U. S. A.*, 110:20455-20460.

Crane BR, Young MW. (2014). Interactive features of proteins composing eukaryotic circadian clocks. *Annu. Rev. Biochem.*, 83:191-219.

Axelrod S, Saez L, Young MW. (2015). Studying circadian rhythm and sleep using genetic screens in *Drosophila*. *Methods Enzymol.* 551:3-27.

Jang AR, Moravcevic K, Saez L, Young MW, Sehgal A. (2015). *Drosophila* TIM binds Importin  $\alpha 1$ , and acts as an adapter to transport PER to the nucleus. *PLoS Genet.*, 11(2): e1004974. doi:10.1371/journal.pgen.1004974.

Kidd PB, Young MW, Siggia ED. (2015) Temperature compensation and temperature sensation in the circadian clock. *Proc. Natl. Acad. Sci. U. S. A.*, 112: E6284-E6292.

Garaulet DL, Sun K, Li W, Wen J, Panzarino AM, O'Neil JL, Hiesinger PR, Young MW, Lai EC. (2016). miR-124 Regulates Diverse Aspects of Rhythmic Behavior in *Drosophila*. *J Neurosci.* 36:3414-3421.

Patke A, Murphy PJ, Onat OE, Krieger AC, Ozcelik T, Campbell SS, Young MW. (2017). Mutation of the Human Circadian Clock Gene CRY1 in Familial Delayed Sleep Phase Disorder. *Cell* 169:203-215.

#### Biosketch:

Vice President for Academic Affairs, Richard and Jeanne Fisher Professor, Head of the Laboratory of Genetics, The Rockefeller University. B.A. in Biology (1971), Ph.D. in Genetics (1975), University of Texas, Austin. Doctoral work with Burke Judd -cytogenetic studies of *Drosophila* chromosomes. Postdoctoral work, Biochemistry, Stanford University Medical School with David Hogness -identification and cloning of *Drosophila* transposable elements. Joined the faculty of The Rockefeller University in 1978 (Fellow of the Andre and Bella Meyer Foundation). Work at Rockefeller has focused on two areas of research: Neuromuscular development, stemming from the laboratory's isolation and studies of the *Notch* locus of *Drosophila*, and the genetics of behavior, primarily circadian rhythms including initial cloning of the *period* gene of *Drosophila*, discovery and functional characterizations of the circadian clock genes *timeless*, *double-time*, *shaggy*, *vriille*, and *pdp1*, and modeling of principal molecular features of the *Drosophila* circadian system.