| TANIA A. BAKER  |                       |
|---|-----------------------|
| Massachusetts Institute of Technology/Howard Hughes Medical Institute | Phone: (617) 253-3594 |
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### **EDUCATION AND TRAINING:**

University of Wisconsin, Madison B.S. Biochemistry, with distinction, 1983

Stanford University Medical School Ph.D. Biochemistry, 1988 Advisor: Professor Arthur Kornberg

Stanford University Medical School, Department of Biochemistry Postdoctoral Fellow, 1988-89 Advisor: Professor Arthur Kornberg

National Institute of Diabetes and Digestive and Kidney Disease, National Institutes of Health Postdoctoral Fellow, 1989-1992 Advisor: Kiyoshi Mizuuchi

### **POSITIONS HELD**:

#### Massachusetts Institute of Technology - Department of Biology

| Assistant Professor                   | 1992-1997    |
|---------------------------------------|--------------|
| Associate Professor                   | 1997-2002    |
| Associate Department Head             | 1999-2004    |
| E. C. Whitehead Professor             | 2002-present |
| Co-Director, Biology Graduate Program | 2008-present |
| Department Head                       | 2012-present |
|                                       |              |

| Howard Hughes Medical Institute |              |
|---------------------------------|--------------|
| Assistant Investigator          | 1994-1997    |
| Associate Investigator          | 1997-2002    |
| Investigator                    | 2002-present |

### HONORS:

- 2008 MacVicar Faculty Fellow for Innovation & Dedication to Undergraduate Education
- 2007 Elected Member of the National Academy of Sciences
- 2005 Elected Fellow of the American Association for the Advancement of Science
- 2004 Elected Fellow of the American Academy of Arts and Sciences
- 2002 Elected Fellow of the American Society for Microbiology
- 2002 E. C. Whitehead Professor of Biology
- 2001 Eli Lilly and Co. Research Award from the American Society for Microbiology
- 2000 MIT School of Science Teaching Prize for Excellence in Undergraduate Education
- 1999 Harold E. Edgerton Award for Distinction in Teaching, Research, and Service to MIT
- 1998 ASBMB Schering-Plough Research Institute Award
- 1993 NSF Young Investigator Award
- 1992-94 Robert A. Swanson Career Development Professorship in the Life Sciences

1992-93 Surdna Foundation Research Award for Support of Junior Faculty in Life Sciences

1989-92 Helen Hay Whitney Foundation Fellowship for Postdoctoral Research

1982-83 Mary Shine Peterson Fellowship for Undergraduate Research

1982 Undergraduate Summer Research Fellowship to Cold Spring Harbor Laboratory

# Named Lectureships

| NIH Directors Lecture, NIH Bethesda, MD                             | Fall, 2005   |
|---|--------------|
| Joan Wood Lecture, Dept. Biology, Indiana University                | Spring, 2008 |
| Gavin Borden Visiting Fellow, Cold Spring Harbor Laboratory         | Spring, 2008 |
| Storer Endowment "Major Issues in Modern Biology Lecture", UC Davis | Fall, 2010   |
| Gerhard Schmidt Memorial Lecture, Tufts University                  | Spring 2011  |

# SERVICE:

# Massachusetts Institute of Technology

| Department Head, Biology  | 2012-2014    |
|---|--------------|
| Associate Department Head, Biology  | 1999-2004    |
| Co-Director, Biology Graduate Program   | 2008-present |
| Graduate Program Steering Committee   | 1994-present |
| Chair, "Target of Opportunity" Faculty Search Committee   | 2006-present |
| Biochemistry/Molecular Mechanism Faculty Search Committee<br>(Chaired 3 years)                          | 7 years      |
| Dean of Science Search Committee  | 2007         |
| School of Science Committee Overseeing Institute Communications<br>Requirement (Co-Chair/Chair 3 years) | 2004-2008    |
| Gender Equity Committee   | 2007-present |
| Gender Equity Chairs Committee  | 2008-present |
| Steering Committee for the Chemistry/Biology Interface Training Program                                 | 2008-present |
| Institute-wide Planning Task Force  | 2009         |
| Provost's Committee for Design of the Faculty Housing Benefit   | 2003         |
| Organized/Taught "Responsible Research Conduct" for Biology PhD Students                                | 2005-2008    |
| Dean's Committee of to recommend Biology Department Head  | 1998         |
| Co-organizer of the MIT Scientific Retreat for Junior Faculty in the<br>School of Science               | 1998         |
| Subcommittee to Design Biology Graduate Lab Rotations   | 1997         |
| Other Service   |              |
| CSHL Academic Affairs Committee   |              |
| CSHL Board of Trustees  |              |
| NIGMS Council: Ad Hoc Member  | May 2010     |
| NIH Study Section Member (PCMB)   | 2006-2009    |
| Ad Hoc Reviewer for: NIH Microbial Physiology and Genetics  |              |
| Ad Hoc Reviewer for NCI   |              |
| National Academy of Science Section 21 Canvassing Panel   | 2007-present |
| ASM Promega Biotechnology Research Award Panel (current Chair)  | 2007-present |
| Protein Society Nominating Committee  | 2006, 2009   |

| Editorial Boards (current and former): <i>Molecular Microbiology, Molecular</i><br>Cell, Genes & Development, Annual Review of Biochemistry |                  |
|---|------------------|
| Member, Nat L. Sternberg Thesis Prize Committee   | 2002-2004        |
| Co-organizer, Keystone Symposium on Transposition and Other<br>Genome Rearrangements, Santa Fe, NM  | 2003             |
| Co-organizer, Molecular Genetics of Bacteria and Phages Annual<br>Meeting (U. Wisconsin, Madison and Cold Spring Harbor Labs)               | 1999, 2000, 2001 |
| Co-organizer, Keystone Symposium on Transposition and DNA<br>Rearrangements, Santa Fe, NM   | 2000             |
| Keystone Symposia of Molecular and Cellular Biology<br>Scientific Advisory Board for Molecular Biology                                      | 2002-2006        |
| Chair, Review Committee for the Department of Biochemistry and<br>Molecular Biology, USC Keck School of Medicine                            | 2007             |
| TEACHING:   |                  |

| Experimental Biology (undergraduate lab)                      | 1993-1995    |
|---|--------------|
| Method and Logic in Molecular Biology (for graduate students) | 1995-1997    |
| Molecular Biology (joint undergraduate/graduate subject)      | 1997-present |
| Principles of Biochemical Analysis (for graduate students)    | 1999-2007    |

## **PUBLICATIONS:**

- Olivares AO, Nager AR, Yosefson O, Sauer RT, Baker TA (2014) Mechanochemical basis of protein degradation by a double-ring AAA+ machine. *Nat. Struct. Mol. Biol.* (in press).
- Cordova JC, Olivares AO, Shin Y, Stinson BM, Calmat S, Schmitz KR, Aubin-Tam ME, Baker TA, Lang MJ, Sauer RT (2014) Stochastic but Highly Coordinated Protein Unfolding and Translocation by the ClpXP Proteolytic Machine. *Cell.* 158(3):647-58.
- Barthelme D, Chen JZ, Grabenstatter J, Baker TA, Sauer RT (2014) Architecture and assembly of the archaeal Cdc48\*20S proteasome. *Proc Natl Acad Sci.* [Epub ahead of print].
- Vieux EF, Wohlever ML, Chen JZ, Sauer RT, Baker TA (2013) Distinct quaternary structures of the AAA+ Lon protease control substrate degredation. *Proc Natl Acad Sci U S A.* 110(22):E2002-8.
- Wohlever ML, Baker TA, Sauer, RT (2013) Roles of the N domain of the AAA+ Lon protease in substrate recognition, allosteric regulation and chaperone activity. *Mol Microbiol*. 91(1):66-78.
- Wohlever ML, Baker TA, Sauer RT (2013) A mutation in the N domain of the *Escherichia coli* lon stabilizes dodecamers and selectively alters degradation of model substrates. *J Bacteriol*. 195(24):5622-8.
- Wohlever ML, Nager AR, Baker TA, Sauer RT (2013) Engineering fluorescent protein substrates for the AAA+ Lon protease. *Protein Eng Des Sel.* [Epub ahead of print]
- Stinson BM, Nager AR, Glynn SE, Schmitz KR, Baker TA, Sauer RT (2013) Nucleotide binding and conformational switching in the hexameric ring of AAA+ machine. Cell. 153(3):628-39.
- Glynn SE, Nager AR, Baker TA, Sauer RT(2012) Dynamic and static components power unfolding in topologically closed rings of a AAA+ proteolytic machine. *Nat Struct Mol Biol.* 19(6):616-22.
- Peterson CN, Levchenko I, Rabinowitz JD, Baker TA, Silhavy TJ (2012) RpoS proteolysis is controlled directly by ATP levels in *Escherichia Coli. Genes Dev.* 26(6):548-53.
- Landgraf D, Okumus B, Chien P, Baker TA, Paulsson J (2012) Segregation of molecules at bacterial cell division reveals native protein localization. *Nature Methods* 9(5):480-2.
- Sundar S, Baker TA, Sauer RT (2012) The I domain of the AAA+ HsIUV protease coordinates substrate binding, ATP hydrolysis, and protein degradation. *Protein Sci.* 21(2):188-98.
- Davis JH, Baker TA, Sauer RT (2011) Small-molecule control of protein degradation using split adaptors. ACS Chem Biol. 6(11):1205-13.

- Abel S, Chien P, Wassmann P, Schirmer T, Kaever V, Laub MT, Baker TA, Jenal U (2011) Regulatory cohesion of cell cycle and cell differentiation through interlinked phosphorylation and second messenger networks. *Mol Cell.* 43(4):550-60.
- Nager AR, Baker TA, Sauer RT (2011) Stepwise Unfolding of a β Barrel Protein by the AAA+ ClpXP Protease. *J Mol Biol.* 413(1):4-16.
- Román-Hernández G, Hou JY, Grant RA, Sauer RT, Baker TA (2011) The ClpS Adaptor Mediates Staged Delivery of N-End Rule Substrates to the AAA+ ClpAP Protease. *Mol Cell* 43(2):217-28.
- Baker TA, Sauer RT (2011) ClpXP, an ATP-powered unfolding and protein-degradation machine. *Biochim Biophys Acta* 1823(1):15-28
- Aubin-Tam ME, Olivares AO, Sauer RT, Baker TA, Lang MJ (2011) Single-molecule protein unfolding and translocation by an ATP-fueled proteolytic machine. *Cell* 145(2):257-67.
- Sauer RT, Baker TA (2011) AAA+ proteases: ATP-fueled machines of protein destruction. *Annu Rev Biochem* 7;80:587-612.
- Sundar S, McGinness KE, Baker TA, Sauer RT (2010) Multiple sequence signals direct recognition and degradation of protein substrates by the AAA+ protease HsIUV. 403(3):420-9.
- Lee Me, Baker TA, Sauer RT (2010) Control of Substrate Gating and Translocation into ClpP by Channel Residues and ClpX Binding. *J Mol Biol.* 399(5):707-18.
- Bissonnette SA, Rivera-Rivera I, Sauer RT, Baker TA (2010) The IbpA and IbpB small heat-shock proteins are substrates of the AAA+ Lon protease. *Mol Microbiol.* 5(6):1539-49.
- Abdelhakim AH, Sauer RT, and Baker TA (2010) The AAA+ ClpX machine unfolds a keystone subunit to remodel the Mu transpososome. *Proc Natl Acad Sci U S A* (6):2437-42.
- Chowdhury T, Ebrahim S, Chien P, Sauer RT, and Baker TA (2010) Versatile modes of peptide recognition by the ClpX N domain mediate alternative adaptor-binding specificities in different bacterial species. *Protein Sci.* 219(2):242-54.
- Glynn SE, Baker TA, and Sauer RT (2009) Crystal structures of asymmetric ClpX hexamers reveal nucleotide-dependent motions in a AAA+ protein-unfolding machine. *Cell* 139(4):744-56.
- Shin Y, Davis JH, Brau RR, Martin A, Kenniston JA, Baker TA, Sauer RT, Lang MJ. (2010) Singlemolecule denaturation and degradation of proteins by the AAA+ ClpXP protease. *Proc Natl Acad Sci U S A* 106(46):19340-5.
- Pruteanu M, Baker TA (2009). Proteolysis in the SOS response and metal homeostasis in *Escherichia coli*. *Res Microbiol*. 160(9):677-83.
- Davis JH, Baker TA, and Sauer RT (2009). Engineering synthetic adaptors and substrates for controlled ClpXP degradation. *J Biol Chem*. 284(33):21848-55
- Barkow SR, Levchenko I, Baker TA, and Sauer RT (2009). Polypeptide translocation by the AAA+ ClpXP protease machine. *Chem Biol.* 16(6):605-12
- Román-Hernández G, Grant RA, Sauer RT, and Baker TA (2009) Molecular basis of substrate selection by the N-end rule adaptor protein ClpS. *Proc Natl Acad Sci U S A*. 106(22):8888-93.
- Pruteanu M, and Baker TA (2009) Controlled degradation by ClpXP protease tunes the levels of the excision repair protein UvrA to the extent of DNA damage. *Mol Microbiol*. 71(4):912-24.
- Wang KH, Roman-Hernandez G, Grant RA, Sauer RT, and Baker TA (2008) The molecular basis of Nend rule recognition. *Mol Cell* 32: 406-14
- Martin A, Baker TA, and Sauer RT (2008) Pore loops of the AAA+ ClpX machine grip substrates to drive translocation and unfolding. *Nat Struct Mol Biol.* 15: 1147-51
- Schweidenback CT & Baker TA (2008) Dissecting the roles of MuB in Mu transposition: ATP regulation of DNA binding is not essential for target delivery. *Proc Natl Acad Sci USA* 105: 12101-7.
- Moore SD, Baker TA and Sauer RT (2008) Forced extraction of targeted components from complex macromolecular assemblies. *Proc Natl Acad Sci USA* 105: 11685-90.
- Yakamavich JA, Baker TA, and Sauer RT (2008) Asymmetric Nucleotide Transactions of the HsIUV Protease. *J Mol Biol.* 380: 946-57.
- Wang KH, Oakes ES, Sauer RT, and Baker TA (2008) Tuning the strength of a bacterial N-end rule degradation signal. *J Biol Chem.* 283: 24600-7.
- Martin A, Baker TA, and Sauer RT (2008) Diverse pore loops of the AAA+ ClpX machine mediate unassisted and adaptor-dependent recognition of ssrA-tagged substrates. *Mol. Cell* 29: 441-50.

- Abdelhakim A, Oakes EC, Sauer RT, and Baker TA (2008) Unique contacts direct high-priority recognition of the tetrameric Mu transposase-DNA complex by the AAA+ unfoldase ClpX. *Mol. Cell* 30: 39-50.
- Hou JY, Sauer RT, and Baker TA (2008) Distinct structural elements of the adaptor proteins ClpS are required for activation and inhibition of degradation by AAA+ protease ClpAP. *Nat Struct Mol. Biol* 15: 288-94.
- Martin A, Baker TA, and Sauer RT (2008) Protein unfolding by AAA+ protease: critical dependence on ATP-hydrolysis rates, energy landscapes, and substrate engagement. *Nat Struct Mol Biol.* 15: 139-145.
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- Chien P, Grant RA, Sauer RT, and Baker TA (2007) Structure and substrate specificity of a SspB ortholog: design implications for AAA+ adaptors. *Structure* 15: 1296-1305
- Martin A, Baker TA and Sauer RT (2007) Distinct static and dynamic interactions control ATPasepeptidase communication in a AAA+ protease. *Mol Cell* 27: 41-52.
- Chien P, Perchuk BS, Laub MT, Sauer RT and Baker TA (2007) Direct and adaptor-mediated substrate recognition by an essential AAA+ protease. *Proc. Natl. Acad. Sci. USA* 104: 6590-5.
- McGinness KE, Bolon DN, Kaganovich M, Baker TA and Sauer RT (2007) Altered tethering of the SspB adaptor to the ClpXP protease causes changes in substrate delivery.
- J Biol Chem. 282: 11465-73.
- Pruteanu M, Neher SB and Baker TA (2007) Ligand-controlled proteolysis of the *Escherichia coli* transcriptional regulator ZntR. *J Bacteriol.* 189: 3017-25.
- Wang KH, Sauer RT and Baker TA (2007) ClpS modulates but is not essential for bacterial N-end rule degradation. *Genes Dev.* 21: 403-8.
- Farrell CM, Baker TA and Sauer RT (2007) Altered specificity of a AAA+ protease. Mol Cell. 25: 161-6.
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- Baker TA and Sauer RT (2006) ATP-dependent proteases of bacteria: recognition logic and operating principles. *Trends Biochem Sci.* 31: 647-653.
- McGinness KE, Baker TA and Sauer RT (2006) Engineering Controllable Protein Degradation. *Mol Cell* 22: 701–707.
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- Burton BM and Baker TA (2005) Remodeling protein complexes: insights from the AAA+ unfoldase ClpX and Mu transposase. *Protein Sci.* 14: 1945-54.
- Hersch GL, Burton RE, Bolon DN, Baker TA and Sauer RT (2005) Asymmetric interactions of ATP with the AAA+ ClpX6 unfoldase: allosteric control of a protein machine. *Cell* 121: 1017-1027.
- Levchenko I, Grant RA, Flynn JM, Sauer RT and Baker TA (2005) Versatile modes of peptide recognition by the AAA+ adaptor protein SspB. *Nat Struct Mol Biol.* 12: 520-525.
- Kenniston JA, Baker TA, and Sauer RT (2005) Partitioning between unfolding and release of native domains during ClpXP degradation determines substrate selectivity and partial processing. *Proc. Natl. Acad. Sci. USA* 102: 1390-5.
- Burton RE, Baker TA and Sauer RT (2005) Nucleotide-dependent substrate recognition by the AAA+ HsIUV protease. *Nat Struct Mol Biol.* 12: 245-51.
- Bolon DN, Grant RA, Baker TA and Sauer RT (2004) Nucleotide-dependent substrate handoff from the SspB adaptor to the AAA+ ClpXP protease. *Mol Cell* 16: 343-50.

- Sauer RT, Bolon DN, Burton BM, Burton RE, Flynn JM, Grant RA, Hersch GL, Joshi SA, Kenniston JA, Levchenko I, Neher SB, Oakes ES, Siddiqui SM, Wah DA, and Baker TA (2004) Sculpting the proteome with AAA(+) proteases and disassembly machines. *Cell* 119: 9-18.
- Flynn JM, Levchenko I, Sauer RT and Baker TA (2004) Modulating substrate choice: the SspB adaptor delivers a regulator of the extracytoplasmic-stress response to the AAA+ protease ClpXP for degradation. *Genes Dev.* 18: 2292-2301.
- Hersch GL, Baker TA and Sauer RT (2004) SspB delivery of substrates for ClpXP proteolysis probed by the design of improved degradation tags. *Proc. Natl. Acad. Sci. USA* 101: 12136-12141.
- Siddiqui SM, Sauer RT and Baker TA (2004) Role of the protein-processing pore of ClpX, an AAA+ ATPase, in recognition and engagement of specific protein substrates. *Genes Dev.* 18: 369-74.
- Joshi SA, Hersch GL, Baker TA and Sauer RT (2004) Communication between ClpX and ClpP during substrate processing and degradation. *Nat Struct Mol Biol.* 11: 404-11.
- Kenniston JA, Burton RE, Siddiqui SM, Baker TA and Sauer RT (2004) Effects of local protein stability and the geometric position of the substrate degradation tag on the efficiency of CIpXP denaturation and degradation. *J Struct Biol.* 146: 130-40.

Bolon DN, Wah DA, Hersch GL, Baker TA and Sauer RT (2004) Bivalent tethering of SspB

- to ClpXP is required for efficient substrate delivery: a protein-design study. *Mol Cell.* 13: 443-9.
- Williams, TL and Baker TA (2004) Reorganization of the Mu transpososome active sites during a cooperative transition between DNA cleavage and joining. *J Biol Chem.* 279: 5135-45.
- Spector S, Flynn JM, Tidor B, Baker TA and Sauer RT (2003) Expression of N-formylated proteins in *Escherichia coli. Protein Expr. Purif.* 32: 317-322.
- Goldhaber-Gordon I, Early MH and Baker TA. (2003) MuA transposase separates DNA sequence recognition from catalysis. *Biochem.* 42: 14633-42.
- Neher SB, Sauer RT and Baker TA (2003) Distinct peptide signals in the UmuD and UmuD' subunits of the UmuD/D' heterodimer mediate tethering and substrate-processing by the ClpXP protease. *Proc. Natl. Acad. Sci. USA* 100: 13219-24.
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- Coros CJ, Sekino Y, Baker TA and Chaconas G (2003) Effect of mutations in the C-terminal domain of Mu B on DNA binding and interactions with Mu A transposase. *J Biol Chem.* 278: 31210-7.
- Kenniston JA, Baker TA, Fernandez JM and Sauer RT (2003) Linkage between ATP consumption and mechanical unfolding during the protein processing reactions of an AAA+ degradation machine. *Cell* 114: 511-20.
- Goldhaber-Gordon I, Early MH and Baker TA (2003) The terminal nucleotide of the Mu genome controls catalysis of DNA strand transfer. *Proc. Natl. Acad. Sci. USA* 100:7509-14.
- Burton RE, Baker TA and Sauer RT (2003) Energy-dependent degradation: linkage between ClpXcatalyzed nucleotide hydrolysis and protein-substrate processing. *Protein Science* 12: 893 - 902.
- Burton BM and Baker TA (2003) Mu transpososome architecture ensures that unfolding by ClpX or proteolysis by ClpXP remodels but does not destroy the complex. *Chem Biol.* 10:463-72.
- Joshi SA, Baker TA and Sauer RT. (2003) C-terminal domain mutations in ClpX uncouple substrate binding from an engagement step required for unfolding. *Mol Microbiol.* 48: 67-76.
- Neher SB, Flynn JM, Sauer RT and Baker TA (2003) Latent ClpX-recognition signals ensure LexA destruction after DNA damage. *Genes and Development* 17: 1084-1089.
- Sokolsky TD and Baker TA (2003) DNA gyrase requirements distinguish the alternate pathways of Mu transposition. *Mol Microbiol.* 47:397-409
- Flynn JM, Neher SB, Kim YI, Sauer RT and Baker TA (2003) Proteomic discovery of cellular substrates of the ClpXP protease reveals five classes of ClpX-recognition signals. *Mol Cell*. 11: 671-83.

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- Lo JH, Baker TA, and Sauer RT (2001) Characterization of the N-terminal repeat domain of *Escherichia coli* ClpA-A class I Clp/HSP100 ATPase. *Protein Science* 10: 551-559.
- Roldan LAS, and Baker TA (2001) Differential role of the Mu B protein in phage Mu integration versus replication: mechanistic insights into two transposition pathways. *Mol Microbiol* 40: 141-55.
- Burton BM, Williams TL, and Baker TA (2001) ClpX-mediated remodeling of Mu transpososomes: selective unfolding of subunits destabilizes the entire complex. *Mol Cell.* 8: 449-54.
- Flynn JM, Levchenko I, Seidel M, Wickner SH, Sauer RT, and Baker TA (2001) Overlapping recognition determinants within the ssrA degradation tag allow modulation of proteolysis. *Proc Natl Acad Sci USA* 11: 10584-9.
- Burton RE, Siddiqui SM, Kim Y-I, Baker TA, and Sauer RT (2001) Effects of protein stability and structure on substrate processing by the ClpXP unfolding and degradation machine. *EMBO J.* 20: 3092-100.
- Rice PA, and Baker TA (2001) Comparative architecture of transposase and integrase complexes. *Nat Struct Biol.* 8: 302-307.
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- Levchenko I, Seidel M, Sauer RT, and Baker TA (2000) A specificity-enhancing factor for the ClpXP degradation machine. *Science* 289: 2354-6.
- Williams TL, and Baker TA (2000) Transposase team puts a headlock on DNA. Science 289: 73-4.
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- Smith CK, Baker TA, and Sauer RT (1999) Lon and Clp family proteases and chaperones share homologous substrate-recognition domains. *Proc. Natl. Acad. Sci. USA* 96: 6678-6682.
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- Baker TA (1994) Replication initiation: a new controller in Escherichia coli. Current Biology 4: 945-946.
- Baker TA, Mizuuchi M, Savilahti H, and Mizuuchi K (1993) Division of labor among monomers within the Mu transposase tetramer. *Cell* 74: 723-733.
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