# CURRICULUM VITAE JAMES M. MANNING

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# A. EDUCATION

1956-1960	Boston College, Chestnut Hill, Massachusetts, B.S. Chemistry, Honors
1960-1966	Tufts University, Medford, Massachusetts, Ph.D. Biochemistry

# **B. APPOINTMENTS**

1996-Present	Professor of Biochemistry, Northeastern University.
1996-Present	Adjunct Professor, The Rockefeller University.
1977-1995	The Rockefeller University, Associate Professor of Biochemistry (tenured); 1988-1995, Head of Laboratory (Biochemistry).
1975-1995	Cornell University Medical College, Adjunct Associate Professor of Biochemistry.
1972-1977	The Rockefeller University, Associate Professor of Biochemistry (untenured).
1969-1972	The Rockefeller University, Assistant Professor of Biochemistry.
1967-1969	The Rockefeller University, Research Associate.
1966-1967	University of Rome, Italy, National Science Foundation Postdoctoral Fellow.
1960-1966	Tufts University, USPHS Pre-doctoral Fellow.

# C. PROFESSIONAL SOCIETIES AND ACTIVITIES

American Society of Biological Chemists

American Chemical Society

The Harvey Society

American Society of Hematology

**Protein Society** 

# D. PROFESSIONAL ACTIVITIES AND AWARDS

1976-1978	American Heart Association, Grants Review Committee
1977-1979	American Society of Biological Chemists, Membership Committee, Chairman (1979)
1977-1979	National Science Foundation, Postdoctoral Fellowship Evaluation Panel, Chairman (1979)
1975-1986	The New England Enzyme Center, Scientific Advisory Committee
1973-Present	National Institutes of Health ad hoc Study Sections

1979-1984	Journal of Biological Chemistry, Editorial Board
1990-Present	Journal of Biological Chemistry, Editorial Board
1987-1997	NIH MERIT Award
1990-Present	Biochemistry and Molecular Biology/Chemtracts, Editorial Board
1992-Present	National Institutes of Health Parent Committee- Sickle Cell Centers, Chairman, Site Visits-
1996	FDA Committee on Evaluation of Blood Substitutes
1999	International Advisory Board, Vitamin 85 and Carbonyl Catalysis

# E. INTERNATIONAL AND NATIONAL MEETINGS - INVITED PRESENTATIONS

1994	Japan Society for Promotion of Science (Lectures throughout Japan)
1994	International Conference on Vitamin 86 and Carbonyl Catalysis, Capri, Italy
1994	International Conference on Strategies for Sickle Cell Anemia, Paris, France
1994	European Peptide Symposium, Braga, Portugal
1995	French Red Cross, Lecture on Blood Substitute Research, Paris, France
1996	National Conference on Blood Substitutes, University of California at San Diego
1996	International Biocatalysis Workshop, Kyoto, Japan
1999	International Conference on Vitamin 86 and Carbonyl Catalysis, Santa Fe NM
2000	Pacifichem International Congress, Honolulu, Hawaii

# F. PUBLICATIONS

- 1. J.S. Nishimura. J.M. Manning, and A. Meister. Studies on the Mechanism of Activation of Aspartic Acid  $\beta$ -Decarboxylase by  $\alpha$ -Keto Acids and Pyridoxal-5'-Phosphate Biochemistry, 1, 442 (1962).
- 2. A. Meister, N. Stone, and J.M. Manning. Conversion of Proline to Hydroxyproline in Collagen Synthesis *Adv. Chem.*, **44**, 67 (1964).
- J.M. Manning, and A. Meister.
   Conversion of Proline to Collagen Hydroxyproline.
   Biochemistry, 5, 1154 (1966).
- J.M. Manning, and A. Meister.
   Conversion of Proline to Collagen Hydroxyproline.
   In "Biological and Chemical Aspects of Oxygenases", K. Bloch, and 0. Hayaishi, Editors, Tokyo, pp. 27 (1966).
- P. Fasella, and J.M. Manning.
   Substrate Structure and Reaction Specificity in Pyridoxal Dependent Enzymes.
   Seventh International Congress of Biochemistry, Supplement I, 1074 (1967).
- 6. J.M. Manning, R.M. Khomutov, and P. Fasella. The Reaction of B-Chloroglutamic Acid with Glutamate-Aspartate Transaminase. Eur. J. Biochem., 5, 109 (1968).

7. J.M. Manning, and S. Moore.

Determination or D- and L-Amino Acids by Ion Exchange Chromatography as L-D and L-L Dipeptides.

J. Biol. Chem., 243, 5591 (1968).

8. J.M. Manning, S. Moore, W.B. Rowe, and A. Meister.
Identification of L-Methionine S-Sulfoximine as the Diastereoisomer of L-MethionineSR-Sufoximine That Inhibits Glutamine Synthetase.
Biochemistry, 8, 2681 (1969).

9. E. Antonini, M. Brunori, P. Fasella, R. Khomutov, J.M. Manning, and E.S. Severin. Kinetic Study of the Interaction Between Aspartate Aminotransferase and Threo-,  $\beta$  - Chloroglutamate. Biochemistry, 9, 1211 (1970).

10. J.M. Manning.

Determination of D- and L-Amino Acid Residues in Peptides. Use of Tritiated Hydrochloric Acid to Correct for Racemization During Acid Hydrolysis. J. Am. Chem. Soc., 92, 7449 (1970).

11. J.M. Manning, A. Marglin, and S. Moore.

Determination of the Stereochemical Purity of Amino Acid Residues in Peptides. Proceedings of the Second American Peptide Symposium, Cleveland, Gordon and Breach, New York (1970).

12. J.M. Manning.

Chromatographic Determination of the D- and L-Amino Acid Residues m Pneumococcal C-Polysaccharide.

J. Biol. Chem., 2-46, 2926 (1971).

13. A. Cerami, and J.M. Manning.

Potassium Cyanate as an Inhibitor of the Sickling of Erythrocytes *In Vitro. Proc. Natl. Acad . Sci., USA,* **68,** 1180 (1971).

14. P.N. Gillette, J.M. Manning, and A. Cerami.

Increased Survival of Sickle Cell Erythrocytes after Treatment *In Vitro* with NaNCO. *Proc. Natl. Acad . Sci., USA,* **68,** 2791 (1971).

15. J.M. Manning,

Determination of D- and L-Amino Acid Residues by Ion Exchange Chromatography. *Method s in Enzymol.*, 25, 9 (1972).

16. F.G. de Furia, D.R. Miller, A. Cerami, and J.M. Manning.

The Effects of Cyanate *In Vitro* on Red Blood Cell Metabolism and Function in Sickle Cell Anemia.

J. Clin. Invest., 51, 566 (1972).

17. J.M. Manning, A. Cerami, P.N. Gillette, F.G. de Furia, and D.R. Miller. Chemical and Biological Aspects of the Inhibition of Red Cell Sickling by Cyanate. *Adv. Exp. Med . Biol.*. **28**, 253 (1972).

18. P.N. Gillette, C.M. Peterson, J.M. Manning, and A. Cerami. Preliminary Clinical Trials with Cyanate.

Adv. Exp. Med . Biol., 28, 261 (1972).

19. J.M. Manning, A. Cerami, P.N. Gillette. F.G. de Furia, and D.R. Miller. Cyanate Inhibition of Red Blood Cell Sickling.

In "Sickle Cell Disease", A. Abramson, J.F. Bertles, and D.L. Wethers, Editors, The C.V. Mosby Company, Saint Louis, pp. 177 (1973).

- 20. A. Cerami, T.A. Allen, J.H. Graziano, F.G. de Furia, J.M. Manning, and P.N. Gillette. Pharmacology of Cyanate I. General Effects on Experimental Animals. *J. Pharmacol. Exp. Therap.*, **185.** 653 (1973).
- J.M. Manning, C.K. Lee, A. Cerami, and P.N. Gillette.
   Gas Chromatographic Determination of the Carbamylation of Hemoglobin by Cyanate.
   J. Lab. Clin. Med., 81, 941 (1973).
- C.K. Lee, J.M. Manning, Kinetics of the Carbamylation of the Amino Groups of Sickle-Cell Hemoglobin by Cyanate. J. Biol. Chem. 2 48, 5861 (1973).
- A. Cerami, J.M. Manning, P.N. Gillette, F.G. de Furia, D.R. Miller, J.H. Graziano, and C.M. .Peterson.
   Effect of Cyanate on Red Blood Cell Sickling.
   Fed . Proc., 32, 1168 (1973).
- 24. N. Njikam. W.M. Jones, A.M. Nigen, P.N. Gillette, R.C. Williams, Jr., and J.M. Manning. Carbamylation of the Chains of Hemoglobin S By Cyanate *In Vitro* and *In Vivo. J. Biol. Chem.*, **248**, 8052 (1973).
- 25. J.M. Manning, A. Cerami, P.N. Gillette, F.G. de Furia, and D.R. Miller. Biochemical and Physiological Properties of Carbamylated Hemoglobin S. *Ad v. Enzymol.*, **4 0.** 1 (1974).
- A.M. Nigen, C.M. Peterson, P.N. Gillette, and J.M. Manning.
   Determination of the Blood Concentrations of Cyanate after Intravenous Administration to Patients with Sickle-Cell Disease.

   J. Lab. Clin. Med., 83, 139 (1974).
- 27. J.M. Manning, A. Cerami, P.N. Gillette, F.G. de Furia, D.R. Miller, and C.M. Peterson. Inhibition of Red Blood Cell Sickling by Cyanate. *Ann. Okla. Acad . Sci.*, **4**, 47 (1974).
- J. M. Manning, N.E. Merrifield, W.M. Jones, and E.C. Gotschlich. Inhibition of Bacterial Growth by , β-Chloro- D-Alanine. Proc. Natl. Acad. Sci., USA, 7 1, 417 (1974).
- 29. A.M. Nigen, N. Njikam, C.K. Lee, and J.M. Manning. Studies on the Mechanism of Action of Cyanate in Sickle Cell Disease. *J. Biol. Chem.*, **249**, 6611 (1974).
- 30. N. Njikam, A.M. Nigen, C.K. Lee, and J.M. Manning.
  Preparation and Properties of Hemoglobin S Hybrids Carbamylated on Specific Chains.
  First National Symposium on Sickle Cell Disease, Washington, D.C., pp. 157 (1974).
- 31. A.M. Nigen, J.M. Manning, C.M. Peterson, and J.M. White.
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- A.M. Nigen, and J.M. Manning.
   The Interaction of Anions with Hemoglobin Carbamylated on Specific NH2-Terminal Residues.
   J. Biol. Chem., 250, 8248 (1975).

33. T.S. Soper, and J.M. Manning.

Synergy in the Antimicrobial Action of Penicillin and ,  $\beta$  -Chloro-D-Alanine *In Vitro*. *Antimicrob*. *Agents Chemotherap.*, **9**, 347 (1976).

34. J.M. Manning, and A.M. Nigen.

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35. A.M. Nigen, B.D. Bass, and J.M. Manning. Reactivity of Cyanate with Valine-I(a) of Hemoglobin as a Probe of Conformational Change. *J. Biol. Chem.*, **2 51**, 7638 (1976).

36. R.S. Lane, J.M. Manning, and E.E. Snell.
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37. T.S. Soper, W.M. Jones, B. Lerner, M. Trop, and J.M. Manning. Inactivation of Bacterial D-Amino Acid Transaminase by. *β*-Chloro-D-Alanine. *J. Biol. Chem.*, **2 52**, 3170 (1977).

38. T.S. Soper, J.M. Manning, P.A. Marcotte, and C.T. Walsh. Inactivation of Bacterial D-Amino Acid Transaminase by the Olefinic Amino Acid D-Vinylglycine. *J. Biol. Chem.*, **252**, 1571 (1977).

39. A.M. Nigen, and J.M. Manning.
Inhibition of Erythrocyte Sickling *In Vitro* by DL-Glyceraldehyde.

Proc. Natl. Acad . Sci., USA, 7 4, 367 (1977).
40. A.M. Nigen, and J.M. Manning.
Effects of Glyceraldehyde on the Structural and Functional Properties of Sickle

Erythrocytes. *J. Clin. Invest.*, **61**, 11 (1978).

J.M. Manning, A.M. Nigen, S. Charache, and J.O. Alben.
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42. J.M. Manning, and A.M. Nigen.
Inhibition of Erythrocyte Sickling *In Vitro* by Glyceraldehyde.
In "Clinical and Biochemical Aspects of Hemoglobin Abnormalities", W.S. Caughey, Editor, Academic Press, New York, pp. 665 (1978).

43. J.M. Manning, and T.S. Soper.  $\beta$  -Elimination of  $\beta$  -Halo Substrates by D-Amino Acid Transaminase Associated with Inactivation of the Enzyme. Trapping of a Key Intermediate in the Reaction. *Biochemistry.* **17.** 3377 (1978).

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44. J.M. Manning, and T.S. Soper.
Inactivation of D-Amino Acid Transaminase and the Antibacterial Action of P-Halo

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45. J. M. Manning, and D.A. Driscoll.

Effects of Glyceraldehyde on Erythrocyte Sickling.

In "NIH/INSERM International Meeting on Sickle Cell Disease", Paris, Elsevier, New York, pp. 187 (1979).

46. T.S. Soper, and J.M. Manning.

Effects of Substrates on the Selective Modification of the Cysteinyl Residues of D- Amino Acid Transaminase.

J. Biol. Chem., 254, 10901 (1979).

47. A.S. Acharya, and J.M. Manning.

Reactivity of the Amino Groups of Carbonmonoxy HemoglobinS with Glyceraldehyde. *J. Biol. Chem.*, 255, 1406 (1980).

48. A.M. Nigen. J.M. Manning, and J.O. Alben.

Oxygen-Linked Binding Sites for Inorganic Anions to Hemoglobin.

J. Biol. Chem., 255, 5525 (1980).

49. D.A. Uvelli, M.Y. Lee, J.M. Manning, M.P. Hlastala, and A.L. Babb. Measurement of the Carbamylation Kinetics and Anti-Sickling Mechanism in HbS Blood. J. *Lab. Clin. Med.*, 95, 748 (1980).

50. A.S. Acharya, and J.M. Manning.

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Mouse Peritoneal Macrophages Release Leukotriene C in Response to a Phagocytic Stimulus.

Proc. Natl. Acad. Sci., USA, 77, 4928 (1980).

52. J.M. Manning.

Preparation of Hemoglobin Carbamylated at Specific NH2-Terminal Residues. *Methods Enzymol.*, 7 6, 159 (1981).

53. T.S. Soper, and J.M. Manning.

Different Modes of Action of Inhibitors of Bacterial D-Amino Acid Transaminase. A Target Enzyme for the Design of New Antibacterial Agents.

J. Biol. Chem., 256, 4263 (1981).

54. R. Benesch, R.E. Benesch, S. Kwong, A.S. Acharya, and J.M. Manning. Labeling of Hemoglobin with Pyridoxal Phosphate.

J. Biol. Chem., 257, 1320 (1982).

55. D.E. Metzler, J.N. Jansonius, A. Arnone, M. Martinez-Carrion, and J.M. Manning. Transaminases (ASBC Minisymposium).

Fed. Prod., 41, 2432 (1982).

56. T.S. Soper, and J.M. Manning.

Inactivation of Pyridoxal Phosphate Enzymes by Gabaculine. Correlation with Enzymic Exchange of  $\beta$ -Protons.

J. Biol. Chem., 257, 13930 (1982).

57. A.S. Acharya, A. Di Donato, B.N. Manjula, V.A. Fischetti, and J.M. Manning. Influence of Trifluoroacetic Acid on Retention Times of Histidine-Containing Peptides in Reverse Phase HPLC.

Int. Pep. Prot. Res., 22, 78 (1983).

58. A.S. Acharya, L.G. Sussman, and J.M. Manning.
Schiff Base Adducts of Glyceraldehyde with Hemoglobin. Differences in the Amadori Rearrangement at the a-Amino Groups.

J. Biol. Chem., 258, 2296 (1983).

59. A.S. Acharya, and J.M. Manning.

The Reaction of Glycolaldehyde with Proteins. Latent Cross-Linking Potential of a-Hydroxyaldehydes.

Proc. Natl. Acad. Sci., USA, 80, 3590 (1983).

60. Di Donato, W.J. Fanti, A.S. Acharya, and J.M. Manning.

Selective Carboxymethylation of the a-Amino Groups of Hemoglobin. Effect on Functional Properties.

J. Biol. Chem., 258, 11890 (1983).

61. R. Seetharam, J.M. Manning, and A.S. Acharya.

Specific Modification of the Carboxyl Groups of Hemoglobin S.

J. Biol. Chem., 258, 14810 (1983).

62. A.S. Acharya, L.G. Sussman, W.M. Jones, and J.M. Manning. Inhibition of Deoxyhemoglobin S Polymerization by Glyceraldehyde. *Anal. Biochem.*, (Stanford Moore Dedicatory Issue), 136, 101 (1984).

63. J.M. Manning, and A.S. Acharya.

The Mechanism of Action of Two Anti-Sickling Agents: Cyanate and Glyceraldehyde. Am. J. *Ped. Hematol/Oncol.*, 6, 51 (1984).

64. Ueno, T.S. Soper, and J.M. Manning.

Enzyme-Activated Inhibition of Bacterial D-Amino Acid Transaminase by .B-Cyano-D-Alanine.

Biochem. Biophys. Res. Commun., 122, 485 (1984).

65. W.M. Jones, T.S. Soper, H. Ueno, and J.M. Manning.

Bacterial D-Amino Acid Transaminase.

Methods Enzymol., 113, 108 (1985).

66. A.S. Acharya, L.G. Sussman, and J.M. Manning.

Selective Modification of the a-Amino Groups of Hemoglobin S by Reductive Alkylation. *J. Biol. Chem.*, 260, 6039 (1985).

67. T.S. Soper, H. Ueno, and J.M. Manning.

Substrate-Induced Changes in Sulfhydryl Reactivity of Bacterial D-Amino Acid Transaminase.

Arch. Biochem. Biophys., 240, I (1985).

68. W.M. Jones, and J.M. Manning.

Acylpeptide Hydrolase Activity from Erythrocytes.

Biochem. Biophys. Res. Commun., 126, 933 (1985)

69. T.S. Soper, and J.M. Manning.

Enzyme-Activated Inhibition of Pyridoxal-Phosphate Enzymes.

In "Transaminases", P. Christen, and D.E. Metzler, Editors, John Wiley and Sons, Inc., New York, pp. 266 (1985).

70. L.J. Benjamin, and J.M. Manning.

Enhanced Survival of Sickle Erythrocytes Upon Treatment . with Glyceraldehyde. Blood, 67, 544 (1986).

- 71. N. Mori, and J.M. Manning.
  - Studies on the Amadori Rearrangement in a Model system: Chromatographic Isolation of Intermediates and Products.
  - Anal. Biochem., 152, 396 (1986).
- 72. Ueno, M.A. Pospischil, R. Kluger, and J.M. Manning. Site-Specific Modification of Hemoglobin by Methyl Acetyl Phosphate. *Arch. Biochem. Biophys.*, 244, 795 (1986).
- 73. Ueno, M.A. Pospischil, R. Kluger, and J.M. Manning. Methyl Acetyl Phosphate: A Novel Acetylating Agent. *J. Chromatogr.*, 359, 193 (1986).
- 74. J.M. Manning, A.S. Acharya, A. Di Donato, W.J. Fantl, W.M. Jones, N. Mori, M. Pospischil, and H. Ueno.
  - Factors that Influence the Reaction of the Amino Groups of Hemoglobin S with Different Compounds.
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- L.J. Benjamin, A. Arthur, E. Sneed, and J.M. Manning. Enhanced Survival and Reduced Density of Sickle Erythrocytes Upon Treatment with Glyceraldehyde. Inserm Symposium, #141, "Approaches to the Therapy of Sickle Cell Anemia", pp. 217 (1986).
- W.M. Jones, L.R. Manning, and J.M. Manning.
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- 78. W.J. Fantl, L.R. Manning, H. Ueno, A. Di Donato, and J.M. Manning. Properties of Carboxymethylated, Cross-Linked Hemoglobin A. *Biochemistry*, 26, 5755 (1987).
- 79. W.J. Fantl, A. Di Donato, J.M. Manning, P.H. Rogers, and A. Arnone. Specifically Carboxymethylated Hemoglobin as an Analogue of Carbamino Hemoglobin: Solution and X-Ray Studies of Carboxymethylated Hemoglobin and X-Ray Studies of Carbamino Hemoglobin.

  J. Biol. Chem., 262, 12700 (1987).
- 80. J.M. Manning, T.S. Soper, P. Recsei, A. Di Donato, M. Merola, and H. Ueno. . . Enzyme-Activated Inhibitors of Bacterial D-Amino Acid Transaminase as Antimicrobial Agents. In "Biochemistry of Vitamin B<sub>6</sub>", I.U.B. Symposium, T. Korpela and P. Christen, Editors, pp. 305 (1987).
- 81. H. Ueno, L.J. Benjamin, and J.M. Manning.
  Effects of Methyl Acetyl Phosphate on Hemoglobin S: A Novel Acetylating Agent Directed Towards the DPG Binding Site.
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82. W.M. Jones, and J.M. Manning.

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Properties of the Hydrolase that Catalyzes Removal of the Blocked NH2-Terminal Residues from Polypeptides.

In "Proteins: Structure and Function", Plenum, pp. 675 (1988).

84. W.J. Fanti, A. Di Donato, A. Arnone, and J.M. Manning.
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In "Proteins: Structure and Function", Plenum, pp. 141 (1988).

85. Ueno, S.S. Koide, J.M. Manning, and S.J. Segal. Gossypol: Interaction with Ribonuclease A. In "Proteins: Structure and Function", Plenum, pp. 149 (1988).

86. N. Mori, and J.M. Manning.

Studies on Nonenzymic Glycosylation of Peptides in a Simple Model System.

In "Proteins: Structure and Function", Plenum. pp. 161 (1987).

87. L.R. Manning, and J.M. Manning.

Influence of the Ligation State and the Concentration of Hemoglobin A on Its Cross-Linking by Glycolaldehyde: Functional Properties of Cross-Linked Carboxymethylated Hemoglobin. Biochemistry, 27, 6640 (1988).

88. N. Mori, Y. Bai, H. Ueno, and J.M. Manning.
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- 89. M. Merola, A. Martinez del Pozo, H. Ueno, P. Recsei, A. Di Donato, J.M. Manning, K. Tanizawa, Y. Masu, S. Asano, H. Tanaka, K. Soda, D. Ringe, and G. Petsko. Site-Directed Mutagenesis of the Cysteinyl Residues of D-Amino Acid Transaminase. Biochemistry. 28, 505 (1989).
- A. Martinez del Pozo, M. Merola, H. Ueno, J.M. Manning, K. Tanizawa, Y. Masu, S. Asano, H. Tanaka, K. Soda, D. Ringe, and G. Petsko.
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- 93. Ueno, and J.M. Manning.

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