Curriculum vitae

ALEXANDER M. KLIBANOV

Date and Place of Birth: July 15, 1949, in Moscow (Russia)

Naturalized U.S. Citizen (1983)

Education:

1974 Ph.D. in Chemical Enzymology, Moscow University

1971 M.S. in Chemistry, Moscow University

Honors:

| U | 13. | |
|---|-------------------|---|
| | 2015 | Tau Beta Pi's Leonardo Da Vinci Lecturer, MIT |
| | 2007-11 and 2014- | Novartis Chair Endowed Professorship, MIT |
| | 2012-13 | Roger and Georges Firmenich Endowed Professorship, MIT |
| | 2011 | MIT Biological Engineering Senior Class Faculty Award |
| | 2006 | Distinguished GRUM Lecturer in Drug Discovery & Development, University of Montreal (Canada) |
| | 2004 | UNAM Distinguished Lecturer, National University of Mexico (Mexico City) |
| | 2001 | Walter Enz Lecturer in Pharmaceutical Chemistry, University of Kansas |
| | 2001 | Elected a Corresponding Fellow of the Royal Society of Edinburgh (Scotland's National Academy of Science and Letters) |
| | 2000 | Merck Distinguished Lecturer, Rutgers University |
| | 2000 | Top 20 Biotechnology & Bioengineering Papers of the Last Forty Years |
| | 1998 | Robert Lutz Lecturer, University of Virginia |
| | 1996 | Perkin-Elmer Distinguished Lecturer, University of Pittsburgh |
| | 1995 | Elected to the National Academy of Sciences of the U.S.A. |
| | 1995 | Nathan O. Kaplan Memorial Lecturer in Biological Chemistry, University of California at San Diego |
| | 1994 | R.W. Johnson PRI Lecturer, Pharmaceutical Research Institute |
| | 1994 | Warren McCabe Lecturer, North Carolina State University |
| | 1993 | Elected to the National Academy of Engineering of the U.S.A. |
| | 1993 | Arthur C. Cope Scholar Award of the American Chemical Society |
| | 1993 | Biotechnology Divisional Lectureship Award of the Institute of Food Technologists |
| | 1992 | Charles Sabat Lecturer, Rutgers University |
| | 1992 | Elected a Founding Fellow of the American Institute for Medical and Biological Engineering |
| | 1992 | Louis C. Jordy Research Scholar Lecturer, Drew University |
| | 1991 | International Enzyme Engineering Award |
| | 1991 | Marvin J. Johnson Award of the American Chemical Society |
| | 1990 | Monsanto Lecturer, Ohio State University |
| | 1990 | NRC Distinguished Lecturer, Academia Sinica (Taiwan) |
| | 1989 | Ipatieff Prize of the American Chemical Society |
| | 1989 | Backer Lecturer, Groningen University (Holland) |
| | 1988 | Dow Lecturer, University of Ottawa (Canada) |
| | 1987 | Distinguished Scholar Lecturer, Hope College |
| | 1986 | Leo Friend Award of the American Chemical Society |
| | 1984 | Who's Who in Frontier Science and Technology |



Mylan v. Regeneron IPR2021-00881 U.S. Pat. 9,254,338 Exhibit 2082

| 1981-1983 1975 | U.S.S.R. Ministry of Higher Education Prize |
|---------------------------------------|--|
| Professional Experience: 2019-present | Novartis Chair Professor Emeritus of Chemistry and Bioengineering, MIT |
| 2014-2019 | Novartis Chair Endowed Professor of Chemistry and Bioengineering, MIT |
| 2012-2013 | Roger and Georges Firmenich Professor of Natural Products Chemistry Department of Chemistry, MIT |
| 2007-2011 | Novartis Chair Endowed Professor of Chemistry and Bioengineering, MIT |
| 2000-2019 | Professor of Bioengineering Department of Biological Engineering, MIT |
| 1988-2019 | Professor of Chemistry Department of Chemistry, MIT |
| 1987-1988 | Professor of Applied Biochemistry Department of Applied Biological Sciences, MIT |
| 1983-1987 | Associate Professor of Applied Biochemistry Department of Applied Biological Sciences (formerly Department of Nutrition and Food Science), MIT |

Assistant Professor of Applied Biochemistry
Department of Nutrition and Food Science, MIT

University of California at San Diego

Research Chemist

Postdoctoral Associate, Department of Chemistry

Department of Chemistry, Moscow University

Sohio Lecturer, Case Western Reserve University

Henry L. Doharty Career Development Professorship, MIT

American Men and Women of Science

Current Journal Editorial/Advisory Boards:

"Biocatalysis and Biotransformation", "Applied Biochemistry and Biotechnology", "Open Chemistry Journal", "Biotechnology Progress", "Biotechnology & Bioengineering", "Microbial Biotechnology", "Open Journal of Pharmacology", "Nanocarriers", "Journal of Antivirals and Antiretrovirals", "Open Access Academic Books in Chemistry", "Journal of Biological Chemistry and Molecular Pharmacology", "Archives of Natural Products and Medicinal Chemistry", "Recent Patents in Biotechnology", "Current Pharmaceutical Biotechnology", and "Archives of Medical Biotechnology", "International Journal of Drug Design, Delivery and Safety"

Professional Societies:

1979-1983

1977-1979

1974-1977

1984

1982

U.S. National Academy of Sciences, U.S. National Academy of Engineering, American Chemical Society, American Institute for Medical and Biological Engineering

Current Research Interests:

Enzyme chemistry and biotechnology Medicinal and formulation chemistry



Protein drug delivery

Antimicrobial polymers
Enzymes as stereoselective catalysts in organic syntheses
Stabilization and formulation of macromolecular pharmaceuticals

Publications:

- 1. Varfolomeyev, S.D., Klibanov, A.M., Berezin, I.V. 1971. Light-initiated enzymic activity caused by photostereoisomerization of *cis*-4-nitrocinnamoyl-α-chymotrypsin. <u>FEBS Lett</u>. <u>15</u>: 118 -120.
- Varfolomeyev, S.D., Klibanov, A.M., Martinek, K., Berezin, I.V. 1972. Light-sensitive catalysts.
 4-Nitrocinnamoyl residue as a chromophoric reporter group in the α-chymotrypsin active center.
 Dokl. Acad. Nauk SSSR 203: 616-619.
- 3. Klibanov, A.M., Samokhin, G.P., Martinek, K., Berezin, I.V. 1974. Mechanochemistry of catalytic systems. Regulation by a mechanical action of the enzymatic properties of α-chymotrypsin covalently attached to a nylon fiber. <u>Dokl. Acad. Nauk SSSR 218</u>: 715-718.
- 4. Berezin, I.V., Varfolomeyev, S.D., Klibanov, A.M., Martinek, K. 1974. Light and ultrasonic regulation of α-chymotrypsin catalytic activity. Proflavin as a light- and sound- sensitive competitive inhibitor. <u>FEBS Lett.</u> 39: 329-331.
- 5. Berezin, I.V., Klibanov, A.M., Martinek, K. 1974. The mechanochemistry of immobilized enzymes. How to steer a chemical process at the molecular level by a mechanical device. Biochim. Biophys. Acta 364: 193-199.
- Berezin, I.V., Klibanov, A.M., Goldmacher, V.S., Martinek, K. 1974. Mechanochemistry of catalytic systems. Regulation by a mechanical action of the enzymatic activity of trypsin entrapped in polyacrylamide gel. <u>Dokl. Acad. Nauk SSSR</u> 218: 367-370.
- 7. Klibanov, A.M., Martinek, K., Berezin, I.V. 1974. The effect of ultrasound on α-chymotrypsin. A novel approach to studying conformational transitions in active centers of enzymes. <u>Biochemistry SSSR</u> 39: 878-887.
- 8. Berezin, I.V., Klibanov, A.M., Klyosov, A.A., Martinek, K., Svedas, V.K. 1975. The effect of ultrasound as a new method of studying conformational transitions in enzyme active centers. pH- and temperature-induced conformational transitions in the active center of penicillin amidase. <u>FEBS Lett. 49</u>: 325-328.
- 9. Berezin, I.V., Klibanov, A.M., Martinek, K. 1975. Kinetic and thermodynamic aspects of catalysis by immobilized enzymes. Russ. Chem. Revs. 44: 17-47.
- Martinek, K., Goldmacher, V.S., Klibanov, A.M., Berezin, I.V. 1975. Denaturing agents (urea, acrylamide) protect enzymes against irreversible thermoinactivation: a study with native and immobilized α-chymotrypsin and trypsin. <u>FEBS Lett.</u> 51: 152-155.
- 11. Martinek, K., Klibanov, A.M., Tchernysheva, A.V., Berezin, I.V. 1975. The stabilization of α-chymotrypsin by entrapment in polymethacrylate gels. Dokl. Acad. Nauk SSSR 223: 233-236.
- 12. Tchernysheva, A.V., Goldmacher, V.S., Klibanov, A.M., Martinek, K., Berezin, I.V. 1975. The catalytic activity and thermostability of α-chymotrypsin oligomers entrapped in cross-linked polymeric gels. Bull. Moscow Univ. 19: 428-431.



- 13. Tchernysheva, A.V., Martinek, K., Klibanov, A.M., Mevkh, A.T., Berezin, I.V. 1975. The catalytic properties and thermostability of α-chymotrypsin in different polymethacrylate gels. <u>Izvestia Acad. Nauk SSSR, Ser. Khim.</u> <u>No.8</u>: 1764-1768.
- 14. Klibanov, A.M., Samokhin, G.P., Martinek, K., Berezin, I.V. 1976. Enzymatic mechanochemistry: a new approach to studying the mechanism of enzyme action. <u>Biochim. Biophys.</u> Acta 438: 1-12.
- 15. Berezin, I.V., Klibanov, A.M., Samokhin, G.P., Martinek, K. 1976. Mechanochemistry of immobilized enzymes: a new approach to studies in fundamental enzymology. Regulation by a mechanical device of the catalytic properties of enzymes covalently attached to elastic polymeric supports. Meth. Enzymol. 44: 558-571.
- 16. Berezin, I.V., Klibanov, A.M., Goldmacher, V.S., Martinek, K. 1976. Mechanochemistry of immobilized enzymes: a new approach to studies in fundamental enzymology. Regulation by a mechanical device of the catalytic activity of enzymes trapped in polyacrylamide gel. Meth. Enzymol. 44: 571-576.
- 17. Klibanov, A.M., Kazanskaya, N.F., Larionova, N.I., Martinek, K., Berezin, I.V. 1976. A comparative study of the dynamic structures of the active centers of proteolytic enzymes by the ultrasonic method. The effect of ultrasound on α-trypsin, β-trypsin, and trypsinogen. J. Bioorg. Chem. (Russ.) 2: 828-836.
- 18. Martinek, K., Goldmacher, V.S., Klibanov, A.M., Torchilin, V.P., Smirnov, V.N., Chazov, E.I., Berezin, I.V. 1976. Thermal stabilization of α-chymotrypsin by covalent attachment to a complementary surface of a polymeric matrix. <u>Dokl. Acad. Nauk SSSR</u> 228: 1468-1471.
- 19. Poglazov, B.F., Samokhin, G.P., Klibanov, A.M., Levitsky, D.I., Martinek, K., Berezin, I.V. 1977. Mechanochemistry of the myosin molecule. <u>Dokl. Acad. Nauk SSSR</u> 234: 482-485.
- 20. Berezin, I.V., Klibanov, A.M., Samokhin, G.P., Goldmacher, V.S., Martinek, K. 1977. Mechanosensitive and sound-sensitive systems as chemical amplifiers of weak signals. In: Biomedical Applications of Immobilized Enzymes and Proteins (T.M.S. Chang, ed.), vol. 2, pp. 237-251, Plenum Press, New York.
- 21. Klibanov, A.M., Samokhin, G.P., Martinek, K., Berezin, I.V. 1977. A new mechanochemical method of enzyme immobilization. <u>Biotechnol. Bioeng.</u> 19: 211-218.
- 22. Martinek, K., Klibanov, A.M., Samokhin, G.P., Semenov, A.M., Berezin, I.V. 1977. Preparative enzymatic synthesis in biphasic water-organic systems. J. Bioorg. Chem. (Russ.) 3: 696-702.
- 23. Klibanov, A.M., Samokhin, G.P., Martinek, K., Berezin, I.V. 1977. A new approach to preparative enzymatic synthesis. <u>Biotechnol. Bioeng.</u> 19: 1351-1361.
- 24. Martinek, K., Klibanov, A.M., Goldmacher, V.S., Berezin, I.V. 1977. The principles of enzyme stabilization. I. Increase in thermostability of enzymes covalently bound to a complementary surface of a polymeric support in a multipoint fashion. <u>Biochim. Biophys. Acta</u> 485: 1-12.



- 25. Martinek, K., Klibanov, A.M., Goldmacher, V.S., Tchernysheva, A.V., Mozhaev, V.V., Berezin, I.V., Glotov, B.O. 1977. The principles of enzyme stabilization. II. Increase in thermostability of enzymes as a result of multipoint non-covalent interactions with a polymeric support. <u>Biochim. Biophys. Acta</u> 485: 13-28.
- 26. Torchilin, V.P., Maksimenko, A.V., Smirnov, V.N., Martinek, K., Klibanov, A.M., Berezin, I.V. 1978. The principles of enzymes stabilization. III. The effect of the length of intramolecular cross-linking agents on the thermostability of enzymes. <u>Biochim. Biophys. Acta</u> 522: 277-283.
- 27. Samokhin, G.P., Klibanov, A.M., Martinek, K. 1978. Photochemical immobilization of enzymes. Bull. Moscow Univ. 19: 433-436.
- 28. Poglazov, B.F., Samokhin, G.P., Klibanov, A.M., Levitsky, D.I., Martinek, K., Berezin, I.V. 1978. The effect of mechanical stretching of myosin rod part (fragment LMM + HMM S-2) on the ATPase activity of myosin. <u>Biochim. Biophys. Acta</u> 524: 245-253.
- 29. Klibanov, A.M., Semenov, A.N., Samokhin, G.P., Martinek, K. 1978. Enzymatic reactions in water-organic solutions. The criterion for selecting optimal organic co-solvents. <u>J. Bioorg. Chem. (Russ.)</u> 4: 236-242.
- 30. Sinitsyn, A.P., Klibanov, A.M., Klesov, A.A., Martinek, K. 1978. The dependence of stability of immobilized glycoamylase on the method of immobilization. <u>Appl. Biochem. Microbiol. (Russ.)</u> 14: 236-242.
- 31. Klibanov, A.M., Mozhaev, V.V. 1978. On the mechanism of irreversible thermoinactivation of enzymes and possibilities for reactivation of "irreversibly" inactivated enzymes. <u>Biochem. Biophys. Res. Commun.</u> <u>83</u>: 1012-1017.
- 32. Klibanov, A.M., Kaplan, N.O., Kamen, M.D. 1978. A rationale for stabilization of oxygen-labile enzymes: application to a clostridial hydrogenase. Proc. Natl. Acad. Sci. USA 75: 3640-3643.
- 33. Torchilin, V.P., Maksimenko, A.V., Smirnov, V.N., Berezin, I.V., Klibanov, A.M., Martinek, K. 1979. The principles of enzyme stabilization. IV. Modification of "key" functional groups in the tertiary structure of proteins. Biochim. Biophys. Acta 567: 1-11.
- 34. Klibanov, A.M., Kaplan, N.O., Kamen, M.D. 1979. Chelating agents protect hydrogenase against oxygen inactivation. <u>Biochim. Biophys. Acta</u> <u>547</u>: 411-416.
- 35. Klibanov, A.M. 1979. Stabilization of enzymes by immobilization. Anal. Biochem. 93: 1-25.
- 36. Klibanov, A.M., Kaplan, N.O., Kamen, M.D. 1980. Thermal stabilities of membrane-bound, solubilized and artificially immobilized hydrogenase from *Chromatium vinosum*. <u>Arch. Biochem. Biophys.</u> 199: 545-549.
- 37. Klibanov, A.M., Puglisi, A.V. 1980. The regeneration of coenzymes using immobilized hydrogenase. <u>Biotechnol. Lett. 2</u>: 445-450.
- 38. Klibanov, A.M., Kaplan, N.O., Kamen, M.D. 1980. Approaches to stabilization of hydrogenase and nitrogenase against oxygen inactivation. In: Enzyme Engineering V (H.H. Weetall and G.P. Royer, eds.), pp.135-142, Plenum Press, New York.



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