

Lipase

Molecular Weight: 45,000 - 50,000

CAS # : 9001-62-1

Physical Description: Off-white to tan powder

Source: *Porcine pancreas*

Activity: See Desnuell (1972) on "Catalytic Properties" (page 586). Momsen and Brockman (1976a and b) report the effects of taurodeoxycholate and co-lipase. At low concentrations, up to 0.3 mM, the bile salt increases the stability of the lipase to 5 fold. At higher levels (0.3 - 0.8 mM), but below the critical micelle concentration, it interferes with enzyme adsorption on the substrate interface, thus inhibiting lipolysis. Co-lipase counters this inhibitory effect by providing high affinity binding sites at the surface of the lipase-bile salt complex. See also Borgstrom and Elanson (1973), Borgstrom et. al. (1974), and Kaimal and Saroja (1989). Co-lipase without bile-salts only mildly stimulates activity. Brockman et. al. (1973) report on lipase activity toward soluble triglycerides such as tripropionin. It is stimulated in the presence of hydrophobic surfaces. Santhanam and Wagle (1971) indicate that protein kinase, Mg^{2+} , ATP and cAMP stimulate lipase activity.

Unit Definition: One USP unit of lipase activity is contained in the amount of pancreatin that liberates 1.0 uEq of acid per minute at a pH of 9.0 at 37°C under the conditions of the assay for lipase activity.

Activators: Ca^{2+} is required for activity [Sr^{2+} and Mg^{2+} are less effective activators (Sarda, et. al. 1957)].

Inhibitors: Versene, Zn^{2+} , Cu^{2+} , Hg^{2+} , iodine, PCMB (Wills, 1960). DFP does not inhibit.

Stabilizers: DFP may be used to stabilize impure preparations containing proteinases in solutions.

Description: Lipase is effective in splitting dietary fats regardless of source. The relative digestive power on various edible fats is illustrated as follows:

Type of Fat	Grams of fat digested in 1 hour by 1 gram of lipase at pH 9.0
Olive Oil (Triolein)	531.4 grams
Tallow (Tristearin)	534.8 grams
Butter (Tributyrin)	181.4 grams
Coconut Oil (Tripalmitin)	484.4 grams

Lipase catalyzes the hydrolysis of emulsified esters of glycerol and long chain fatty acids. The substrate is not a single molecule but a nonaqueous phase of aggregated lipid (Brockhoff and Jensen 1974). The operative substrate characteristic is aggregates of ester molecules, micelles or monomolecular film, interfacing an aqueous medium. Enzyme activity is directly related to the concentration of substrate molecules on the interface (Esposito et. al. 1973; Lagocki et. al. 1973). Lipase attacks the primary ester groups most readily. Monoglycerides are poor substrates (it is the 2-monoglycerides that are absorbed through the intestinal wall and reformed into lymph chylomicrons). Pancreatic lipases have been thoroughly reviewed by Brockhoff and Jensen (1974), and Desnuelle (1972). Lieberman and Ollis (1975) have reported on lipase immobilized on stainless steel and polyacrylamide beads. Using a fluidized bed recycle reactor it is indicated that enzyme-substrate affinity is not altered.

Recommended Storage: 0°C

Composition of Lipase: Two lipases are present. Lipase A is more acidic than Lipase B; otherwise, the two isoenzymes are nearly the same (Verger et. al. 1969). Normally, a cofactor is bound to the enzymes (Maylie et. al. 1971). Two co-lipases were purified by Erlanson et. al. (1973). They were quite similar polypeptide chains with a molecular weight of 11,000. Borstrom and Erlanson (1973) indicated that co-lipase might be classified as a co-enzyme for lipase in that they interact in a stoichiometrical relationship.

The amino acid composition, which is almost identical except for isoleucine, is shown in Brockhoff and Jensen (1974). Both contain a carbohydrate moiety (Garner and Smith 1972). Histidine is involved in the active site (Semeriva et. al. 1971). See Hultin (1992). Modification of the free carboxyl group by amide formation inactivates the enzyme (Semeriva et. al. 1972). According to Desnuelle (1972) the carboxyl in lipase stabilizes the active enzyme, i.e., the enzyme conformation resulting from adsorption at a hydrophobic interface. Although lipase contains two disulfide groups, they are not involved in enzymatic activity (Verger et. al. 1971). Diisopropylphosphorofluoridate (DFP) binds to a tyrosine residue but is not inhibitory (Maylie et. al. 1969).

Extinction coefficient: $E_{280}^{1\%} = 13.3$ (Desnuelle 1972)

Isoelectric point: Lipase A = 4.9 (Brockhoff and Jensen 1974) and Lipase B = 5.0

Solubility: Partly soluble in water and insoluble in alcohol.

References:

1. Albro, P., and Latimer, A.: Pancreatic Nonspecific Lipase, an Enzyme Highly Specific for Micelles, *Biochem.*, **13**, 1431 (1974).
2. Antonian, E.: Recent Advances in the Purification, Characterization and Structure Determination of Lipases, *Lipids*, **23**, 1101 (1988).
3. Arzoglou, P., Tavidou, A., and Balaska, C.: Rapid Turbidimetric Determination of Lipase Activity in Biological Fluids, *Anal. Letters*, **22**, 1459 (1989).
4. Borgström, B., and Erlanson, C.: Pancreatic Lipase and Co-Lipase. Interactions and Effects of Bile Salts and Other Detergents, *Eur. J. Biochem.*, **37**, 60 (1973).
5. Borgström, B., Erlanson, C., and Sterby, B.: Further Characterization of Two Co-Lipases from Pancreas, *Biochem. Biophys. Res. Comm.*, **59**, 902 (1974).
6. Bradshaw, W., and Rutter, W.: Multiple Pancreatic Lipases. Tissue Distribution and Pattern of Accumulation During Embryological Development, *Biochem.*, **11**, 1517 (1972).
7. Bockerhoff, H.: Action of Pancreatic Lipase on Emulsions of Water-Insoluble Esters, *Arch. Biochem. Biophys.*, **134**, 366 (1969a).
8. Bockerhoff, H.: Esters of Phenols as Substrate for Pancreatic Lipase, *Biochim. Biophys. Acta*, **191**, 181 (1969b).
9. Bockerhoff, H., and Jensen, R.: in *Lipolytic Enzymes*, Academic Press, NY (1974).
10. Brockman, H., Law, J., and Kézdy, F.: Catalysis by Adsorbed Enzymes. The Hydrolysis of Tripalmitin by Pancreatic Lipase Adsorbed to Silicized Glass Beads, *J. Biol. Chem.*, **248**, 4965 (1973).
11. Chailan, C., Kerfelec, B., Foglizzo, E., and Chapus, C.: Direct Involvement of the C-Terminal Extremity of Pancreatic Lipase (403-449) in Colipase Binding, *Biochem. Biophys. Res. Comm.*, **184**, 206 (1992).
12. Chapus, C., Rovey, M., Sarda, L., and Verger, R.: Minireview on Pancreatic Lipase and Colipase, *Biochimie*, **70**, 1223 (1988).
13. Clement, G., Clement, J., and Bezard, J.: Action of Human Pancreatic Lipase on Synthetic Mixed Symmetrical Triglycerides of Long-Chain Acids and Butyric Acid, *Biochem. Biophys. Res. Comm.*, **8**, 238 (1962).
14. Coleman, M.: A Rapid Lipase Purification, *Biochim. Biophys. Acta*, **67**, 146 (1963).
15. Deckelbaum, R., Hamilton, J., Moser, A., Bengtsson-Olivecrona, G., Butbul, E., Carpentier, Y., Gutman, A., and Olivecrona, T.: Medium Chain versus Long-chain Triacylglycerol Emulsion Hydrolysis by Lipoprotein Lipase and Hepatic Lipase: Implications for the Mechanisms of Lipase Action, *Biochem. USA*, **29**, 1136 (1990).
16. Desnuelle, P.: *Pancreatic Lipase*, in *Advances in Enzymology*, 23 (Nord, F., ed.), Interscience Publ., Inc., NY, 129 (1961).
17. Desnuelle, P.: in *The Enzymes*, Vol. VII, 3rd Ed., (Boyer, P., ed.) Academic Press, NY, 575 (1972).
18. Desnuelle, P., Naudet, M., and Constantin, M.: Action des Sels de Calcium Sur L'Hydrolyse des Triglycerides Par La Pancreatin et le Suc Pancreatique, *Biochim. Biophys. Acta*, **5**, 561 (1950).
19. Desnuelle, P., Sarda, L., and Ailhaud, G.: Inhibition de la Lipase Pancreatique par la Diethyl-p-nitrophenyl Phosphate en Emulsion, *Biochim. Biophys. Acta*, **37**, 570 (1960).
20. Dufour, C., Séméria, M., and Desnuelle, P.: The Role of Carboxyl Groups in the Activity of Pancreatic Lipase, *Biochim. Biophys. Acta*, **327**, 101 (1973).
21. Ergon, F., and André, G.: Simple High Performance Liquid Chromatography Methods for Monitoring Lipase Reactions, *Lipids*, **24**, 76 (1989).
22. Erlanson, C., and Borgström, B.: Purification and Further Characterization of Co-Lipase from Porcine Pancreas, *Biochim. Biophys. Acta*, **271**, 400 (1972).
23. Erlanson, C., Fernlund, P., and Borgström, B.: Purification and Characterization of Two Proteins with Co-Lipase Activity from Porcine Pancreas, *Biochim. Biophys. Acta*, **310**, 437 (1973).
24. Esposito, S., Séméria, M., and Desnuelle, P.: Effect of Surface Pressure on the Hydrolysis of Ester Monolayers by Pancreatic Lipase, *Biochim. Biophys. Acta*, **302**, 293 (1973).
25. Estessangles, B., Sari, H., and Desnuelle, P.: On the Positional Specificity of Pancreatic Lipase, *Biochim. Biophys. Acta*, **125**, 597 (1966).
26. Fischer, B., and Kleber, H.: Isolation and Characterization of the Extracellular Lipase of *Acinetobacter calcoaceticus* 69 V, *J. Basic Microbiol.*, **27**, 427 (1987).
27. Fischer, C., and Höll, W.: Lipoprotein Lipase as a New Tool in Steryl Ester Analysis, *Lipids*, **25**, 292 (1990).
28. Garner, C., and Smith, L.: Porcine Pancreatic Lipase. A Glycoprotein, *J. Biol. Chem.*, **247**, 561 (1972).
29. Hahn, P.: Abolishment of Alimentary Lipemia Following Injection of Heparin, *Science*, **98**, 19 (1943).
30. Huang, J., Roheim, P., Sloop, C., and Wong, L.: A Sensitive, Inexpensive Method for Determining Minute Quantities of Lipase Activity, *Anal. Biochem.*, **179**, 413 (1989).
31. Hultin, P., and Jones, J.: Dilemma Regarding an Active Site Model for Porcine Pancreatic Lipase, *Tetrahedron Lett.*, **33**, 1399 (1992).
32. Julien, R., Canioni, P., Rathelot, J., Sarda, L., and Plummer, T.: Studies on Bovine Pancreatic Lipase and Colipase, *Biochim. Biophys. Acta*, **280**, 215 (1972).
33. Kaimal, T., and Saroja, M.: Enhancement of Catalytic Activity of Porcine Pancreatic Lipase by Reductive Alkylation, *Biotech. Letters*, **11**, 31 (1989).
34. Kaimal, T., and Saroja, M.: The Active Site Composition of Porcine Pancreatic Lipase: Possible Involvement of Lysine, *Biochim. Biophys. Acta*, **999**, 331 (1989).
35. Kimura, H., Kitamura, T., and Tsuji, M.: Studies on Human Pancreatic Lipase. I. Interconversion Between Low and High Molecular Forms of Human Pancreatic Lipase, *Biochim. Biophys. Acta*, **270**, 307 (1972).
36. Kimura, H., Mukai, M., and Kitamura, T.: Purification and Properties of a Protein Activator of Human Pancreatic Lipase, *J. Biochem. Japan*, **76**, 1287 (1974).
37. Kom, E.: Studies on Clearing Factor. A Lipoprotein Lipase, *Circulation*, **10**, 591 (1954).
38. Kreutzer, H., Pennings, A., Punt, J., and Verduin, P.: Further Studies on the Determination of Lipase Activity, *Clin. Chim. Acta*, **60**, 273 (1975).
39. Lagocki, J., Boyd, N., Law, J., and Kézdy, F.: Kinetics Analysis of the Action of Pancreatic Lipase on Lipid Monolayers, *J. Am. Chem. Soc.*, **92**, 2923 (1970).
40. Lagocki, J., Law, J., and Kézdy, F.: The Kinetic Study of Enzyme Action on Substrate Monolayers. Pancreatic Lipase Reactions, *J. Biol. Chem.*, **248**, 580 (1973).
41. Lieberman, R., and Ollis, D.: Hydrolysis of Particulate Tributyrin in A Fluidized Lipase Reactor, *Biotech. Bioengr.*, **17**, 1401 (1975).
42. Mani, V., and Lakshminarayana, G.: Comparative Rate of Lipolysis of Different Fatty Acids in Pancreatic Lipase Hydrolysis of Ethylene Glycol Mixed Diesters, *Biochim. Biophys. Acta*, **202**, 547 (1970).
43. Marchis-Mouren, G., Sarda, L., and Desnuelle, P.: Purification of Hog Pancreatic Lipase, *Arch. Biochem. Biophys.*, **83**, 309 (1959).
44. Mattson, F., and Beck, L.: The Specificity of Pancreatic Lipase for the Primary Hydroxyl Groups of Glycerides, *J. Biol. Chem.*, **219**, 735 (1956).

45. Mattson, F., and Latton, E.: The Specific Distribution of Fatty Acids in the Glycerides of Animal and Vegetable Fats, *J. Biol. Chem.*, **233**, 868 (1958).
46. Maylié, M., Charles, M., and Desnuelle, P.: Action of Organophosphates and Sulfonyl Halides on Pancreatic Lipase, *Biochem. Biophys. Acta*, **276**, 162 (1972).
47. Maylié, M., Charles, M., Gache, C., and Desnuelle, P.: Isolation and Partial Identification of a Pancreatic Colipase, *Biochim. Biophys. Acta*, **229**, 286 (1971).
48. Maylié, M., Charles, M., Sarda, L., and Desnuelle, P.: Action of Organophosphates on Pancreatic Lipase, *Biochim. Biophys. Acta*, **178**, 196 (1969).
49. Momsen, W., and Brockman, H.: Inhibition of Pancreatic Lipase B Activity by Taurodeoxycholate and Its Reversal by Colipase, Mechanism of Action, *J. Biol. Chem.*, **251**, 384 (1976b).
50. Momsen, W., and Brockman, H.: Effects of Colipase and Taurodeoxycholate on the Catalytic and Physical Properties of Pancreatic Lipase B at an Oil-Water Interface, *J. Biol. Chem.*, **251**, 378 (1976a).
51. Moreau, H., Moulin, A., Gargouri, Y., Noel, J., and Verger, R.: Inactivation of Gastric and Pancreatic Lipases by Diethyl Para-Nitrophenyl Phosphate, *Biochem. USA*, **30**, 1037 (1991).
52. Mullenax, D., and Lopez, A.: Swine Pancreatic Lipase Activity at Low Temperatures, *J. Food Sci.*, **40**, 310 (1975).
53. Overbeek, G.: Fat-Splitting Enzymes in Blood, *Clin. Chim. Acta*, **2**, 1 (1957).
54. Ramachandran, S., Yip, Y., and Wagle, S.: Studies on Lipase. Isolation and Purification of a High Molecular Weight Pancreatic Lipase, *Eur. J. Biochem.*, **12**, 201 (1970).
55. Rasco, B., Hultin, H.: A Comparison of Dogfish and Porcine Pancreatic Lipases, *Comp. Biochem. Physiol.*, **89**, 671 (1988).
56. Rovey, M., Bianchetta, J., and Guidoni, A.: Chemical Structure of Porcine Pancreatic Lipase. End Group Determination and Cyanogen Bromide Peptides, *Biochim. Biophys. Acta*, **328**, 391 (1973).
57. Safarik, I.: A Spectrophotometric Assay for Lipase Activity Utilizing Immobilized Triacylglycerols, *J. Biochem. Biophys. Meth.*, **23**, 249 (1991).
58. Santhanam, K., and Wagle, S.: Studies on In Vitro Activation of High Molecular Weight Pancreatic Lipase, *Biochem. Biophys. Res. Comm.*, **43**, 1369 (1971).
59. Sarda, L., Marchis-Mouren, G., Constantin, M., and Desnuelle, P.: Sur Quelques Essais de Purification de la Lipase Pancréatique, *Biochim. Biophys. Acta*, **23**, 264 (1957).
60. Sarda, L., Maylié, M., Roger, J., and Desnuelle, P.: Comportement de la Lipase Pancréatique sur Sephadex Application à la Purification et à la Détermination du Poids Moléculaire de cet Enzyme, *Biochim. Biophys. Acta*, **89**, 183 (1964).
61. Savary, P.: Action of Rat Pancreatic Juice and of Purified Pig Pancreatic Lipase Upon the Esters of Short-Chain Aliphatic Monoacids and Long-Chain Primary Monoalcohols, *Biochim. Biophys. Acta*, **270**, 463 (1972).
62. Savary, P., and Desnuelle, P.: Sur Quelques Elements du Specificité Pendant L'Hydrolyse Enzymatique des Triglycerides, *Biochim. Biophys. Acta*, **21**, 349 (1956).
63. Schandl, A., and Pittner, F.: The Role of Na⁺ and Ca²⁺ Ions on the Action of Pancreatic Lipase Studied with the Help of Immobilisation Techniques, *Eur. J. Biochem.*, **140**, 547 (1984).
64. Sémériva, M., and Dufour, C.: Further Studies on the Exocellular Lipase of *Rhizopus arrizus*, *Biochim. Biophys. Acta*, **260**, 373 (1972).
65. Sémériva, M., Dufour, C., and Desnuelle, P.: On the Probable Involvement of a Histidine Residue in the Active Site of Pancreatic Lipase, *Biochem.*, **10**, 2143 (1971).
66. Singer, T.: On the Mechanism of Enzyme Inhibition by Sulfhydryl Reagents, *J. Biol. Chem.*, **174**, 11 (1948).
67. Sugiura, M., Oikawa, T., Hirano, K., and Inukai, T.: Purification, Crystallization and Properties of Triacylglycerol Lipase from *Pseudomonas fluorescens*, *Biochim. Biophys. Acta*, **488**, 353 (1977).
68. Szafran, H., Popiela, T., and Szafran, Z.: Identity and Properties of Gastric Lipase, *Gastroenterology*, **64**, 140 (1973).
69. Tashiro, J., Kobayashi, J., Shirai, K., Saito, Y., Nakamura, H., Morimoto, Y., and Yoshida, S.: Trypsin Treatment May Impair the Interfacial Activation Action of Lipoprotein Lipase, *J. Biochem.*, **111**, 509 (1992).
70. Uzawa, H., Nishida, Y., Ohru, H., and Meguro, H.: A New Approach to Determine the Stereospecificity in Lipase Catalyzed Hydrolysis Using Circular Dichroism (CD): Lipases Produce Optically Active Diglycerides from Achiral Triglycerides, *Biochem. Biophys. Res. Comm.*, **168**, 506 (1990).
71. Vadehra, D., and Haromon, L.: Characterization of Purified Staphylococcal Lipase, *Appl. Microbiol.*, **15**, 480 (1967).
72. Vandemeers, A., Vandemeers-Piret, M., Rathe, J., and Christophe, J.: On Human Pancreatic Triacylglycerol Lipase: Isolation and Some Properties, *Biochim. Biophys. Acta*, **370**, 257 (1974).
73. Veeraragavan, K.: A Simple and Sensitive Method for the Estimation of Microbial Lipase Activity, *Anal. Biochem.*, **186**, 301 (1990).
74. Verduin, P., Punt, J., and Kreutzer, H.: Studies on the Determination of Lipase Activity, *Clin. Chim. Acta*, **46**, 11 (1973).
75. Verger, R., DeHass, R., Sarda, L., and Desnuelle, P.: Purification from Porcine Pancreas of Two Molecular Species with Lipase Activity, *Biochim. Biophys. Acta*, **188**, 272 (1969).
76. Verger, R., Sarda, L., and Desnuelle, P.: The Sulfhydryl Groups of Pancreatic Lipase, *Biochim. Biophys. Acta*, **207**, 377 (1970).
77. Verger, R., Sarda, L., and Desnuelle, P.: On the Sulfhydryl Groups of Porcine Pancreatic Lipase and Their Possible Role in the Activity of the Enzyme, *Biochim. Biophys. Acta*, **242**, 580 (1971).
78. Whitaker, J.: A Rapid and Specific Method for the Determination of Pancreatic Lipase in Serum and Urine, *Clin. Chim. Acta*, **44**, 133 (1973).
79. Wills, E.: The Relation of Metals and -SH Groups to the Activity of Pancreatic Lipase, *Biochim. Biophys. Acta*, **40**, 481 (1960).
80. Yagi, K., Osamura, M., and Ohishi, N.: Enzyme-Substrate Complex of Pancreatic Lipase and Its Artificial Substrates, Riboflavin Di- and Tributyrates, *J. Biochem. Japan*, **73**, 455 (1973).
81. Yip, Y., Ramachandran, S., and Wagle, S.: Studies on Catalytic Properties of Purified High Molecular Weight Pancreatic Lipase, *J. Exp. Biol. Med.*, **149**, 683 (1975).
82. Zieve, F.: A Rapid and Sensitive Lipase Assay Using Triethylaminoethyl Cellulose Columns, *Anal. Biochem.*, **51**, 436 (1973).
83. Zinterhofer, L., Wardlaw, S., Jatlow, P., and Seligson, D.: Nephelometric Determination of Pancreatic Enzymes. II. Lipase, *Clin. Chim. Acta*, **44**, 173 (1973).