UNITED STATES PATENT AND TRADEMARK OFFICE

BEFORE THE PATENT TRIAL AND APPEAL BOARD

Merck Sharp & Dohme LLC,
Petitioner,

v.

Halozyme Inc.,
Patent Owner.

Case No. PGR2025-00003
U.S. Patent No. 11,952,600

PETITIONER'S UPDATED EXHIBIT LIST

No.	Exhibit Description
1001	U.S. Patent No. 11,952,600
1002	File History of U.S. Patent No. 11,952,600
1003	Declaration of Dr. Michael Hecht
1004	Declaration of Dr. Sheldon Park
1005	U.S. Patent No. 7,767,429
1006	Chao et al., "Structure of Human Hyaluronidase-1, a Hyaluronan Hydrolyzing Enzyme Involved in Tumor Growth and Angiogenesis," Biochemistry, 46:6911-6920 (2007)
1007	WO 2010/077297, published 8 July 2010
1008	Stern et al., "The Hyaluronidases: Their Genomics, Structures, and Mechanisms of Action," Chem. Rev. 106:818-839 (2006)
1009	Jedzrejas et al., "Structures of Vertebrate Hyaluronidases and Their Unique Enzymatic Mechanism of Hydrolysis," Proteins: Structure, Function and Bioinformatics, 61:227-238 (2005)
1010	Zhang et al., "Hyaluronidase Activity of Human Hyal1 Requires Active Site Acidic and Tyrosine Residues," J. Biol. Chem., 284(14):9433-9442 (2009)
1011	Arming et al., "In vitro mutagenesis of PH-20 hyaluronidase from human sperm," Eur. J. Biochem., 247:810-814 (1997)
1012	Bordoli et al., "Protein structure homology modeling using SWISS-MODEL workspace," Nature Protocols, 4(1):1-13 (2008)
1013	Frost, "Recombinant human hyaluronidase (rHuPH20): an enabling platform for subcutaneous drug and fluid administration," Expert Opinion on Drug Delivery, 4(4):427-440 (2007)
1014	Brandon & Tooze, "Introduction to Protein Structure," Second Ed., Chapters 1-6, 11-12, 17-18 (1999)
1015	Table Associating Citations from the '600 Patent (EX1001) to Corresponding Citations in the '731 Application (EX1026)
1016	Steipe, "Consensus-Based Engineering of Protein Stability: From Intrabodies to Thermostable Enzymes," Methods in Enzymology, 388:176-186 (2004)

No.	Exhibit Description
1017	Green, "Computer Graphics, Homology Modeling, and Bioinformatics," Protein Eng'g & Design, Ch. 10, 223-237 (2010)
1018	Chica et al., "Semi-rational approaches to engineering enzyme activity: combining the benefits of directed evolution and rational design," Curr. Opin. Biotechnol., (4):378-384 (2005)
1019	Hardy et al., "Assessment of contraceptive vaccines based on recombinant mouse sperm protein PH20," Reprod., 127:325-334 (2004)
1020	Pomering et al., "Restricted Entry of IgG into Male and Female Rabbit Reproductive Ducts Following Immunization with Recombinant Rabbit PH-20," Am. J. Reprod. Immunol., (3):174-82 (2002)
1021	Baba et al., "Mouse Sperm Lacking Cell Surface Hyaluronidase PH-20 Can Pass through the Layer of Cumulus Cells and Fertilize the Egg," J. Biol. Chem., 277(33):30310-4 (2002)
1022	Primakoff et al., "Reversible Contraceptive Effect of PH-20 Immunization in Male Guinea Pigs," Biol Reprod., 56(5):1142-6 (1997)
1023	Tung et al., "Mechanism of Infertility in Male Guinea Pigs Immunized with Sperm PH-20," Biol. Reprod., 56(5):1133-41 (1997)
1024	Rosengren et al., "Recombinant Human PH20: Baseline Analysis of the Reactive Antibody Prevalence in the General Population Using Healthy Subjects," BioDrugs, 32(1):83-89 (2018)
1025	U.S. Patent No. 9,447,401
1026	U.S. Patent Application No. 13/694,731
1027	2024 Chemistry Nobel Prize Background
1028	[Reserved]
1029	Gmachl et al., "The human sperm protein PH-20 has hyaluronidase activity," FEBS Letters, 3:545-548 (1993)
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No.	Exhibit Description
1031	Yue et al., "Loss of Protein Structure Stability as a Major Causative Factor in Monogenic Disease," J. Mol. Biol., 353:459-473 (2005)
1032	Wang & Moult, "SNPs, Protein Structure, and Disease," Hum. Mutation, 17:263-270 (2001)
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1037	Benkert et al., "Toward the Estimation of the Absolute Quality of Individual Protein Structure Models," Bioinformatics, 27:343-350 (2010)
1038	Schwede et al., "SWISS-MODEL: An Automated Protein Homology-Modeling Server," Nucleic Acids Rsch., 31:3381-3385 (2003)
1039	Alberts, "Molecular Biology of the Cell," Fifth Edition, Chapter 3 (2007).
1040	He et al., "NMR Structures of Two Designed Proteins with High Sequence Identity but Different Fold and Function," PNAS, 105:14412-14417 (2008)
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1042	Ruan et al., "Design and Characterization of a Protein Fold Switching Network," Nature Comm., 14 (2023)
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1044	Mihel, "PSAIA – Protein Structure and Interaction Analyzer," BMC Structural Biology, 8:21 (2008)
1045	Redline Comparison of the '731 and '600 Specifications

No.	Exhibit Description
1046	Beasley & Hecht, "Protein Design: The Choice of *de Novo* Sequences," J. Biological Chemistry, 272:2031-2034 (1997)
1047	Xiong et al., "Periodicity of Polar and Nonpolar Amino Acids is the Major Determinant of Secondary Structure in Self-Assembling Oligomeric Peptides," PNAS, 92: 6349-6353 (1995)
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1050	Pace & Scholtz, "A Helix Propensity Scale Based on Experimental Studies of Peptides and Proteins," Biophysical J. 75:422-427 (1998)
1051	U.S. Patent Application No. 61/631,313
1052	U.S. Patent Application No. 61/796,208
1053	Hom_pre2011
1054	Hom_pre2011_header
1055	Hom_pre2011_header_clean
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1058	Ph20_pre2011 Alignment html
1059	Leisola & Turunen, "Protein Engineering: Opportunities and Challenges," Appl. Microbiol. Biotechnol. 75:1225-1232 (2007)
1060	Hecht et al., "De Novo Proteins from Designed Combinatorial Libraries," Protein Sci., 13:1711-1723 (2004)
1061	Rosengren et al., "Clinical Immunogenicity of rHuPH20, a Hyaluronidase Enabling Subcutaneous Drug Administration," AAPS J., 17:1144-1156 (2015)
1062-1063	[Reserved]
1064	Collection of BLAST Webpages from the Internet Archive, navigable from:

No.	Exhibit Description
	https://web.archive.org/web/20111022151531/http://www.clustal.org/omega/
1065	Collection of Clustal Omega Webpages from the Internet Archive, navigable from: https://web.archive.org/web/20111022151531/http://www.clustal.org/omega/
1066	Collection of SWISS-MODEL Webpages from the Internet Archive, navigable from: https://web.archive.org/web/20110519141121/http://swissmodel.expasy.org/?pid=smh01&uid=&token=
1067	Collection of PyMol Webpages from the Internet Archive, navigable from: https://web.archive.org/web/20110701072314/http://pymol.org/
1068	Declaration of Jeffrey P. Kushan
1069	Swiss Model Printout of PH20 Model
1070	Swiss Model Printout of PH20 Model with D320K Mutation
1071	Swiss Model Printout of PH20 Model with D320H Mutation
1072	Swiss Model Printout of PH20 Model with D320R Mutation
1073	Swiss Model Printout of PH20 Model with D320S Mutation
1074	Declaration of Leif E. Peterson
1075-1112	[Reserved]
1113	Declaration of Sue Wang
1114	Declaration of Brian M. Goldberg
1115	Declaration of Katherine A. Helm
1116	Declaration of Christine M. Engen
1117	Declaration of Amit Bhatla
1118	[Reserved]
1119	Csoka, A. et al., “The Six Hyaluronidase-Like Genes in the Human and Mouse Genomes” Matrix Biology 20:499-508 (2001)

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1120	Huo, Z. et al., "Systemic and Mucosal Immune Responses to Sublingual or Intramuscular Human Papilloma Virus Antigens in Healthy Female Volunteers," PLoS ONE 7(3):e33736 (2012)
1121	[Reserved]
1122	Printz, M. et al., "Risk Factors, Hyaluronidase Expression, and Clinical Immunogenicity of Recombinant Human Hyaluronidase PH20, an Enzyme Enabling Subcutaneous Drug Administration," AAPS 24(6):110 (2022)
1123-1124	[Reserved]
1125	Emails re: AutoDock Vina (Sept. 9, 2025 - Oct. 27, 2025)
1126	Im, S. et al., "Cryo-EM Structure of Human Hyaluronidase PH-20," Proteins: Structure, Function, and Bioinformatics 93:1067-1073 (2025)
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1129	Transcript of the Deposition of Dr. Gary Cherr, Ph.D., November 12, 2025
1130	Melanie Ann Simpson, Ph.D. Deposition Transcript
1131	Gregory Petsko, Ph.D. Deposition Transcript
1132	Transcript of the Deposition of Dr. James Moon, Ph.D., November 18, 2025
1133	Declaration of James Naismith, Ph.D.
1134	Declaration of Dr. Garnett Kelsoe
1135	Dr. Naismith Analysis Spreadsheet
1136	Dobson, "Principles of protein folding, misfolding and aggregation" Semin. Cell. Dev. Biol. 15(1):3-16 (2004)
1137	Dobson, "Protein folding and misfolding" Nature. 18;426(6968):884-90 (2003)
1138	Dobson, "Experimental investigation of protein folding and misfolding" Methods 34(1):4-14 (2004)
1139	Chiti & Dobson, "Protein Misfolding, Functional Amyloid, and Human Disease" Annu. Rev. Biochem. 75:333–66 (2006)

No.	Exhibit Description
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1141	Goldsmith & Tawfik, "Directed enzyme evolution: beyond the low-hanging fruit" Curr. Opin. Struct. Biol. 22(4):406-12 (2012)
1142	Farber & Petsko, "The evolution of alpha/beta barrel enzymes" Trends Biochem. Sci. 15(6):228-34 (1990)
1143	Nagano et al., "One Fold with Many Functions: The Evolutionary Relationships between TIM Barrel Families Based on their Sequences, Structures and Functions" J. Mol. Biol. 30;321(5):741-65 (2002)
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1152	Ringe & Petsko, "Q&A: What are pharmacological chaperones and why are they interesting?" J. Biol. 8:80 (2009)
1153	Petsko, "Herding CATS" Sci. Transl. Med. 24;3(97):97 (2011)
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1155	Calloni et al., "Investigating the Effects of Mutations on Protein Aggregation in the Cell" J. Biol. Chem. 10607-13 (2005)
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1159	Brachmann et al., "Genetic selection of intragenic suppressor mutations that reverse the effect of common p53 cancer mutations" EMBO Journal 17:1847–59 (1998)
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No.	Exhibit Description
1166	MERCK_PGR00462 - RE_ 2_25-cv-03179-ES-JRA_ HALOZYME, INC. v. MERCK SHARP & DOHME CORP.
1167	MERCK_PGR00094 - DNJ-2-25-cv-03179 Stipulation to Substitute Parties
1168	MERCK_PGR00090 - DNJ-2-25-cv-03179 2025-07-14 [019] Corporate Disclosure Statement
1169	Declaration of Jeffrey P. Kushan (Petitioner's Reply)
1170	Murphy, K., "Janeway's Immuno Biology," (8th 2012), Chapters 1, 4, 8, 10, 11, 12, 15, and Appx I
1171	Yang, X. et al., "An Introduction to Epitope Prediction Methods and Software," Rev. Med. Virol., 19: 77–96 (2009)
1172	Bolland, S., "An Innate Path to Human B Cell Tolerance," Immunity, 29(5):667-9 (2008)
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1189	Martin-Deleon, "Epididymal SPAM1 and its Impact on Sperm Function" Molecular and Cellular Endocrinology 250:114-121 (2006)
1190	Nolan et al., "The Safety of Recombinant Human Hyaluronidase PH20 in Nonclinical Models: An Overview of Toxicology,

No.	Exhibit Description
	Pharmacology, and Impact of Anti-PH20 Antibodies," Journal of Controlled Release, 374:369-383 (2024)
1191	McCutcheon, J. et al., "HLA-B*0702 Antibody Epitopes are Affected Indirectly by Distant Antigen Residues," Hum Immunol., 36(2):69-75 (1993)
1192	Nobusawa, E. et al., "Comparison of Complete Amino Acid Sequences and Receptor-Binding Properties among 13 Serotypes of Hemagglutinins of Influenza A Viruses," Virology, 182(2):475-85 (1991)
1193	CDC on H5N1 CDC Press Release: "Highly Pathogenic Avian Influenza A (H5N1) Virus Infection Reported in a Person in the U.S." (Apr. 1, 2024)
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1195	Zhang, H. et al., "Mouse Spam1 (PH-20) Is a Multifunctional Protein: Evidence for Its Expression in the Female Reproductive Tract," Biol Reprod., 69(2):446-54 (2003)
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1201	Rodriguez, L. et al., "Foot and mouth disease virus vaccines," Vaccine 27:D90-D94 (2009)
1202	Redwood, A. et al., "Viral Vectored Immunocontraception: Screening of Multiple Fertility Antigens using Murine

No.	Exhibit Description
	Cytomegalovirus as a Vaccine Vector," Vaccine, 25(4):698-708 (2007)
1203	Russell, M. et al., "Complement-Fixing Properties of Human IgA Antibodies. Alternative Pathway Complement Activation by Plastic-Bound, but not Specific Antigen-Bound, IgA," Scand J Immunol., 30(2):175-83 (1989)
1204	Nikolova E. et al., "All Forms of Human IgA Antibodies Bound to Antigen Interfere with Complement (C3) Fixation Induced by IgG or by Antigen Alone," Scand J Immunol., 39(3):275-80 (1994)
1205	Goodey, N. et al., "Allosteric Regulation and Catalysis Emerge via a Common Route," Nat Chem Biol. 4(8):474-82 (2008)
1206-1212	[Reserved]
1213	Supplemental Material to Exhibit 1120 [NOT FILED]
1214	Supplemental Material to Exhibit 1122 [NOT FILED]
1215	Supplemental Material to Exhibit 1144 [NOT FILED]
1216	Supplemental Material to Exhibit 1151 [NOT FILED]
1217	Supplemental Material to Exhibit 1155 [NOT FILED]
1218	Supplemental Material to Exhibit 1157 [NOT FILED]
1219	Supplemental Material to Exhibit 1173 [NOT FILED]
1220	Supplemental Material to Exhibit 1175 [NOT FILED]
1221	Supplemental Material to Exhibit 1190 [NOT FILED]
1222	Supplemental Material to Exhibit 1202 [NOT FILED]

No.	Exhibit Description
1223	Declaration of Jeffrey P. Kushan [NOT FILED]
1224-1229	[Reserved]
1230 [NEW]	[PROTECTIVE ORDER MATERIAL - UNREDACTED FOR IN CAMERA REVIEW ONLY]
1231 [NEW]	[PROTECTIVE ORDER MATERIAL - UNREDACTED FOR IN CAMERA REVIEW ONLY]
1232 [NEW]	[PROTECTIVE ORDER MATERIAL - UNREDACTED FOR IN CAMERA REVIEW ONLY]
1233 [NEW]	[PROTECTIVE ORDER MATERIAL - UNREDACTED FOR IN CAMERA REVIEW ONLY]

Dated: February 11, 2026

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CERTIFICATE OF SERVICE

Pursuant to 37 C.F.R. § 42.6(e), I hereby certify that on this 11th day of February, 2025, I caused to be served a true and correct copy of the foregoing and any accompanying exhibits by e-mail on the following counsel of record for Patent Owner:

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