

CURRICULUM VITAE

PART I: General Information

Name: Mason Wright Freeman

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FAX: 617-726-2879

Place of Birth: Arlington, Virginia

Spouse: Gale Sherrard Haydock, M.D.

Children: James Harrington (1/13/89), Sarah Wright (2/5/91)

Education:

1973 B.A., Harvard College, Cambridge, MA *cum laude* in History & Literature

1979 M.D., University of California, San Francisco

Postdoctoral Training:

Internship and Residencies:

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| 1979-1980 | Intern in Medicine | Massachusetts General Hospital |
| 1980-1981 | Junior Assistant Resident in Medicine | Massachusetts General Hospital |
| 1981-1982 | Senior Assistant Resident in Medicine | Massachusetts General Hospital |
| 1985 | Chief Resident in Medicine | Massachusetts General Hospital |

Fellowships:

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| 1982-1984 | Clinical and Research Fellow in Medicine (Endocrinology), Massachusetts General Hospital |
| 1986-1990 | Research Fellow in Medicine, Harvard Medical School, Boston, MA |
| 1989-1990 | Postdoctoral Research Fellow, Department of Biology, Massachusetts Institute of Technology, Cambridge, MA |

Licensure and Certification:

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| 1981 | Massachusetts Board of Registration |
| 1983 | American Board of Internal Medicine |
| 1985 | American Board of Internal Medicine, Subspecialty Board in Endocrinology and Metabolism |

Academic Appointments:

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| 1985-1990 | Instructor in Medicine | Harvard Medical School, Boston, MA |
| 1990-2000 | Assistant Professor | Harvard Medical School, Boston, MA |
| 2001-present | Associate Professor | Harvard Medical School, Boston, MA |

Hospital Appointments: (all at the Massachusetts General Hospital)

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| 1985-1990 | Assistant in Medicine |
| 1990-1994 | Assistant Physician |
| 1994-2001 | Associate Physician |
| 2001-present | Physician |
| 1990-1992 | Director, Cardiovascular Health Center |
| 1992-present | Chief, Lipid Metabolism Unit |
| 1994- | Director, Melvin and Barbara Nessel Gene Therapy Center |
| 2004 - | Assoc. Program Director for Academic Career Development Department of Medicine, Internal Medicine Residency Program |

Other Professional Positions and Major Visiting Appointments:

- 2004-present Director, Genetics and Genomics Unit
Clinical Research Program Massachusetts General Hospital
- 2004-present Associate Program Director for Academic Development
Internal Medicine Residency Program Massachusetts General Hospital
- 2004-present Clinical Director of Adult Medical Genetics
Partner's Healthcare Massachusetts General Hospital

Major Committee Assignments:

National and Regional:

- 1986-1988 Voluntary Hospitals of America, Lipid Council, member
- 1991-1995 American Heart Association, Massachusetts Affiliate, Cholesterol Educational Task Force
Senior council member
- 1991-1995 American Heart Association, Research Peer Review Committee, reviewer
- 1997-2000 National Institutes of Health, Metabolism Study section member
- 2002-present The Endocrine Society, Annual Meeting Steering Committee
- 2002 Site Visit team, National Institutes of Health,
Program Project in Diabetes, University of Washington, Seattle, WA.
- 2002-present The Endocrine Society, Cardiovascular Task Force
- 2003 Co-chair, Programs in Genomics Application symposium organizing committee, NHLBI
- 2003-present American Heart Association, program/abstract review member
- 2003-present National Institutes of Health, ZRG1-F06 Study section, Member

Harvard Medical School:

- 1985-1986 Medicine Curriculum Committee, New Pathway Program, member
- 1993-1999 Harvard University Committee on Microbiologic Safety, Human Gene Therapy
Subcommittee, member
- 1999-2004 Harvard University Committee on Microbiologic Safety, Human Gene Therapy
Subcommittee, Chairman

Massachusetts General Hospital/Partner's:

- 1984-present Intern Selection Committee (periodic sabbaticals), member
- 1989-1990 Committee on Research, member
- 1993-1994 Co-Chairman, Committee on Research, Subcommittee on Gene Therapy
- 1994-1995 Transgenic Animal Planning Committee, member
- 1994-1996 Committee on Research, Subcommittee on Review of Research Proposals, member
- 1994-1995 Partner's Subcommittee on Gene Therapy, Co-Chairman
- 1994-1996 Committee on Research, Subcommittee on Office of Technology Affairs, member
- 1995-1996 Research Computer Advisory Group, member
- 1996-1997 Committee on Research, Subcommittee on Genetic Research Guidelines, member
- 1995-present Clinical Research Council, member
- 1997-1999 Partner's Council on Atherosclerosis, Co-chairman
- 2000 Partner's Genetics Advisory Panel, member
- 2004-present Executive Committee on Research (elected by MGH scientific community)
- 2004-present Partner's Steering Committee on Genetics and Genomics

Professional Societies

- 2002-present The Endocrine Society
- 2000-present Atherosclerosis, Thrombosis, and Vascular Biology Council of the Am Heart Association

Community Service:

- 1996-2004 Established lipid clinic in outpatient internal medicine practice located in New Bedford, MA to serve the needs of local residents.

Editorial Boards

American Journal of Medicine
UpToDate (Section Editor for Lipid disorders)

Reviewer

Arteriosclerosis, Thrombosis, and Vascular Biology
Atherosclerosis
Blood

Circulation
Circulation Research
Genomics
Journal of Biological Chemistry
Journal of Clinical Endocrinology and Metabolism
Journal of Clinical Investigation
Journal of Experimental Medicine
Journal of Lipid Research
Nature Medicine
New England Journal of Medicine
Proceedings of the National Academy of Sciences
Science

Awards and Honors:

- 1974-1979 Regents Scholar, University of California, San Francisco
- 1979 Alpha Omega Alpha
- 1985-1990 Physician-Scientist Awardee, NIH
- 1990-1993 American Heart Association, National Grant-in-Aid award

PART II: Research, Teaching and Clinical Contributions

Narrative Report:

My major effort at both MGH and Harvard Medical School focuses on disorders of lipid metabolism and their contribution to atherosclerosis, with an increasing emphasis on designing novel genetic strategies to explore those disorders and treat them. I created the Lipid Clinic at MGH in 1986, ushering in the use of HMG CoA reductase inhibitors to treat cholesterol disorders before their FDA approval in 1987. This was followed by my creation of a basic research laboratory and then a clinical lipid laboratory in 1990-2. The clinic continues to serve as the major site for evaluating and managing patients with lipid disorders at the MGH as well as one of the major teaching venues for lipid disorders at Harvard Medical School. Medical students and residents on the Endocrine service participate with Endocrine fellows in this weekly clinic. An off-site clinic in New Bedford, MA served as a teaching site for fellows and provided a community service to an under-served population in that region of the state from 1996-2004. Other teaching responsibilities include annual service as an attending physician on the Bigelow Medical and Endocrine Services, as well as

the lipid lecturer in the Harvard Medical School human physiology course for second year students and a lecturer in the new Pasteur program at HMS. In the current academic year, I taught in a new Harvard Science and Technology course (HST-146), “Human Biochemistry & Metabolic Diseases” and expect to participate in that annually. In 2004, I became the Associate Program Director for the MGH Internal Medicine Residency program and have co-organized a new curriculum component for the housestaff which focuses on teaching about the tools needed to perform modern clinical investigation.

My research laboratory’s initial work focused on the movement of cholesterol into macrophages and the impact of that event on inflammation. This work began by studying the structure and function of the macrophage scavenger receptors. I cloned the first macrophage scavenger receptor (SR-A) ever identified while a post-doctoral fellow at MIT in Monty Krieger’s lab (Nature, 1990). The work has evolved into broader studies of macrophage function in atherosclerosis using both transgenic and gene deletion animal models. My lab established that PPAR γ was not required for macrophage differentiation as had been thought and that ligands for that transcription factor did not exacerbate lipid accumulation and foam cell formation, a concern that had been raised in the treatment of diabetics with the thiazolidinedione class of oral hypoglycemics (Nature Medicine, 2001). Our latest work on scavenger receptors (J Clin Invest 2005, in press) actually challenges the major paradigm of lipid-induced foam cell formation in atherosclerosis and is causing a substantial revision in thinking about the importance of oxidation of LDL in atherogenesis.

In addition to studying the lipid uptake processes in macrophages, we have also examined cholesterol movement out of macrophages via efflux pathways involving the ABCA class of ABC transporters. We originally mapped the gene causing Tangier disease to chromosome 9 and were one of several groups to identify mutations in ABCA1 as the cause of the disease (JLR, 2000). My lab subsequently determined the major topological features of ABCA1 in the plasma membrane, a topology which has proven to be the model for the entire A class of ABC transporters (JBC 2001). We continue to work on the cell biology of this transporter, with our major focus concentrating on how the transporter interacts with the apolipoprotein acceptors to which it transfers cholesterol. This work has established that a direct protein-protein contact between the acceptor apoprotein, apo A-I, and ABCA1 is required for cholesterol efflux to occur in cells

(JBC, 2002). We recently identified a novel protein interaction domain in the transporter and have utilized a purification method developed in our lab to isolate binding partners required for cholesterol efflux to occur (JBC 2004). Currently, we are generating mice lacking expression of the other members of the A class of transporters to determine their presumptive role in lipid trafficking, which ABCA7 representing the first in this group of novel knockouts (JBC, 2005).

Our work has also led us to study the inflammatory signaling cascades activated by lipid interactions with macrophages and this work has in turn led us into studies of the macrophage receptor signal transduction pathways engaged by microbial lipids. We are currently examining whether these pathogen pathways can also be activated by endogenous lipids, which would provide one of the critical missing links in our understanding of the pathogenesis of atherosclerosis. Our study (Nature Medicine, 2004) showing that a deletion of MyD88, a central adaptor protein in the toll receptor signaling cascade, profoundly reduces atherosclerosis has opened up an entirely new understanding of inflammatory signaling pathways in atherosclerosis and a several other laboratories have now followed up on our seminal finding by confirming that Toll receptors do contribute to atherogenesis.

Another interest of the laboratory has been the development of expression profiling as a tool for characterizing gene expression in macrophages and atheroma. I direct the microarray facility at MGH that is seeking to generate low-cost, high-density oligonucleotide spotted microarrays as part of my responsibilities as the Director of our NHLBI funded Program in Genomics center. This center is currently providing microarray expression profiling services for a large number of NHLBI funded investigators in Boston and around the country.

B. Research Funding Information:

Past:

- 1985-1990 Physician Scientist Award, National Institutes of Health
- 1986-1988 Lovastatin treatment of patients with moderate hypercholesterolemia; a multicenter clinical trial. (Principal Investigator, Massachusetts General Hospital trial center)
- 1987-1989 Lovastatin effects on the eye; a multicenter clinical trial.
(Principal Investigator, Massachusetts General Hospital trial center)
- 1991-1994 American Heart Association/National Grant-in-Aid, Principal Investigator
Rabbit macrophage scavenger receptors in atherosclerosis.
- 1994-1999 NHLBI/RO1HL 53694 Co-investigator
Gene Therapy of Hemophilias A and B
- 1997-2000 Genome Therapeutics Principal Investigator
Functional studies of the Tangier disease gene.
- 1997-2002 NHLBI Specialized Center on Research in Molecular Medicine and Atherosclerosis
Critical Transitions in Atherosclerosis; PI, Project II MGH subcontract
Transgenic strategies to explore plaque rupture in atherosclerotic mice

Current:

- 1990-2006 NHLBI/RO1 HL 45098 Principal Investigator
Scavenger Receptors in Atherosclerosis.
- 1995-2005 NIHPO1DK 50305 Key Investigator
The Phagocyte Membrane
Project 2: Macrophage lipid receptors in atherosclerosis
- 2000-2005 NIHPO1DK 50305 Core Director: Molecular and Cell Biology Core
The Phagocyte Membrane
- 2000-2005 NHLBI UO1 HL66678 Co- Director and Director
Program in Genomics Applications : Genomic Analysis of Stress and Inflammation
- 2001-2006 NCRR/R24 RR14466 Principal Investigator
Development and Characterization of CD14 deficient mice
- 2002-2007 NHLBI/RO1 HL 68988 Principal Investigator
Cell biology of ABCA1
- 2002-2006 NHLBI/RO1 HL072358 Principal Investigator
Shared Microarray Facility

C. Report of Current Research Activities:

Atherosclerosis-transgenic and gene-deletion models aimed at examining macrophage function in the mouse, including the role of pathogen activated inflammatory pathways in the genesis of atheroma

Macrophage lipid receptors structure/function studies and examination of their role in atherosclerosis

Disorders of lipid metabolism- studies of the cell biology of the ABCA class of ABC transporters. Mutations in ABCA1 cause Tangier disease, which is characterized by the near absence of circulating HDL.

D. Report of Teaching:

1. Local contributions

Harvard Medical School

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| 1991-1993 | Pharmacology & Clinical Therapeutics (HST 350) Lecturer/discussion leader 50-75 medical students 2 hours/yr |
| 1992 | Enteric Biology Course for Clinical and Research Fellows Gastrointestinal fellows from Harvard and other teaching hospitals 30 students Lecturer (2 hours) |
| 1995-present | HMS Human Systems-Lipids Second year HMS students 130 students Lecture 1-2 hrs/yr |
| 1996-1999 | HMS Gene Therapy Course Graduate students at HMS and undergraduates at Harvard College 20 students Lecture 2 hrs/yr |
| 1995-2000 | Primed course (HMS Primary care postgraduate course) Practicing internists 1500-2500 physicians Lecture 1hr/yr |
| 2000-present | Pasteur Course Medical students 20-39 students Lecture 1hr/yr |
| 2001 | HST 0.90s Cardiovascular Pathophysiology for Engineers and Scientists 20 students |
| 2000- | Intellihealth- on-line consultant editor |
| 2002-present | Genetics Review course-Partners program in Human genetics Genetic counseling students |

10 students
Lecture 1.5 hr/yr

2004-present HST-146 HST , Human Biochemistry & Metabolic Diseases
30 Medical Students
Lecture 2/hr yr

Massachusetts General Hospital

- 1985 Chief Resident in Medicine, Massachusetts General Hospital
Conducted residents report and attending rounds daily for year
Taught 20 medical students two times per week
Preparation time was 30 hrs/week for 50 weeks
- 1986-2002 Attending Physician, Bigelow Medical Service, MGH
2 medical students, 1 resident, three interns
1 month per year (3-5 hrs/day)
- 1986-present Attending Physician, Endocrine Service, MGH
1-4 medical students, 1-2 residents, and 3-5 Endocrine fellows
1 month/year (2-3 hrs/day)
- 1986-present Housestaff lecturer for inpatient and ambulatory care rotations
50-75 interns, residents, and medical students
Lecturer, 2-4 times per year; 1 hour lectures
- 1986-present Attending Physician and Director of Lipid Clinic
1-2 medical students, 1-2 residents, and 3 Endocrine fellows
Usually 1-3 trainees present each week in the clinic; clinic meets 1 day/week for 4 hours
- 1997-present Attending Physician, New Bedford Lipid Clinic
1 medical student or 1 fellow (both Endocrine and Cardiac fellows)
1 day per month for the entire day (8hrs)
- 2004-present Director, Tools for Human Investigation
Two-week daily course for internal medicine residents provided four times/yr
4 hours per day
15-20 residents
- 1988-present Continuing Medical Education lecturer in several Medical and Surgical Service courses including: Cardiology, Endocrinology, Internal Medicine, Reproductive Endocrinology, and Women's Health, Primary Care for Subspecialists, Vascular Surgery. Lectures have been given every year since 1988 in Cardiology, Endocrinology, and Internal Medicine courses with an occasional hiatus due to scheduling conflicts. Topics vary from management of hyperlipidemia to use of molecular biologic techniques in reproductive endocrinology. Typically 100-300 physicians
Lecture, 1hr/yr for each course and usually an additional hour involving meet the professor sessions.

The lecture given in the internal medicine course has consistently been one of the top-rated talks in the course every year for the past decade

- 1987 Medical Grand Rounds
New approaches to the treatment of hyperlipidemia
- 1990 Medical Grand Rounds
The role of modified lipoproteins and scavenger receptors in atherosclerosis
- 1993 Medical Grand Rounds
A tale of two E's; apo E and Vitamin E in Alzheimer's Disease and Coronary Heart Disease
- 1997 Medical Grand Rounds
Gene therapy
- 2001 Medical Grand Rounds
Trafficking in cholesterol: one Island's export product
- 2003 Endocrine Grand Rounds
The Ins and Outs of Cholesterol transport
- 2004 Medical Grand Rounds
Cholesterol: New genes, new insights, new therapies.

Advising Responsibilities

Post-doctoral trainees in my laboratory (1 pre-doctoral)

| <u>Duration of Training</u> | <u>Name</u> | <u>Current Position</u> |
|-----------------------------|--------------------------|--|
| 1990-1992 | Perry E. Bickel, M.D. | Assistant Professor, Washington University, St. Louis, MO |
| 1990-1996 | Paul Aftring, M.D., Ph.D | Senior Director, Metabolic Research GlaxoSmithKline, Philadelphia, PA |
| 1995-1999 | Gretchen Eberhart, M.D. | Instructor in Medicine University of Pittsburgh, Pittsburgh, PA |
| 1996-2001 | Kathryn Moore, Ph.D | Assistant Professor Harvard Medical School |
| 1997-2001 | Xinzhong Wang, Ph.D. | Research staff scientist Biogen, Cambridge, MA |
| 1997-1998 | Rosalind Fabunmi, Ph.D. | Staff Scientist American Heart Association, Dallas, TX |
| 1999-2001 | M. Murakawa, MD PhD | Staff physician Kyushu University Hospital Fukuoka, Japan |

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| 2000-2001 | Yuri Bobryshev, Ph.D. | Senior Research Officer Univ of New South Wales Sydney, Australia |
| 1997-2002 | Vidya Kunjathoor, Ph.D. | Senior staff scientist, Atherosclerosis Research Novartis Cambridge, MA |
| 1999-2003 | Michael Fitzgerald, Ph.D. | Instructor in Medicine Harvard Medical School |
| 2001-2004 | Harry Bjorkbacka, Ph.D | Asst. Professor Experimentell kardiovaskulär forskning Universitetssjukhuset MAS Malmö, Sweden |
| 2001-present | W S Kim, Ph.D | Research Fellow Harvard Medical School |
| 2002-2003 | Francois Huet, Ph.D. | Staff scientist, Acambis, Cambridge, MA |
| 2003-present | Keichiro Okuhira, Ph.D. | Research Fellow, Harvard Medical School |
| 2004-present | Marc Laberge | HST 3 rd year medical student Harvard-MIT HST program |

2. Regional, National, and International Contributions

Invited Presentations (selected and most recent samples)

- 1989 IXth International Symposium on Drugs Affecting Lipid Metabolism, Houston, Texas
Cloning of the macrophage scavenger receptor
- 1990 31st International Conference on the Biochemistry of Lipids, Munster, Germany
Macrophage scavenger receptors
- 1991 Annual Session of the American College of Physicians, New Orleans, LA
Managing Hyperlipidemia

- 1991 Deuel Lipid Conference, Monterey, CA
Macrophage scavenger receptor: role in foam cell formation
- 1992 Atherosclerosis Forum, Meribel, France
Macrophage scavenger receptors
- 1993 Cellular and Molecular Biology of the Cardiovascular System, Oviedo, Spain
Molecular pathogenesis of atherosclerosis
- 1994 Colorado Diabetes/Endocrine Institute, Snowmass, CO
Oxidized lipids and atherosclerosis
- 1995 Mid-Atlantic Lipid Research Symposium, Atlantic City, NJ
Scavenger receptors in atherosclerosis
- 1995 Gordon research conference: atherosclerosis, invited speaker, Meriden, N.H.
Scavenger receptor promoter activity
- 1996 Ministry of Science, Education, and Culture of Japan; Tokyo, Japan
Macrophages and Atherosclerosis
- 1998 NIH Specialized Ctrs of Research in Molecular Medicine & Atherosclerosis, Bethesda, MD
Transgenic strategies for exploring plaque rupture
- 1999 NIH conference on Niemann-Pick C disease, Bethesda, MD
Adenoviral-mediated complementation of genetic mutations
- 2001 NHLBI Program in Genomics Applications, Milwaukee, WI
Genetic studies in human subjects-what are the rules?
- 2002 The Endocrine Society Annual Meeting, San Francisco, CA
Speaker and Symposium Chair, Genetics and Endocrinology
- 2002 The American Heart Association, Chicago Illinois
Speaker and Plenary Chair, Macrophage lipid accumulation and atherosclerosis
- 2002 NIDDK symposium on Microarrays, Bethesda, MD
The use of spotted oligonucleotide arrays
- 2003 Harvard School of Public Health, Boston, MA
The Ins and Outs of Cholesterol Trafficking
- 2003 The Endocrine Society: Meet the Professor. Philadelphia, PA
Statin Therapies: Treating the difficult-to-treat patient
- 2003 NIH Program in Genomics Applications symposium, Bethesda, MD
The development of spotted oligonucleotide microarrays for gene expression profiling
- 2003 Clinical Endocrinology Update: The Endocrine Society Board Review Course- Miami, FL
Lipid disorders- top-rated lecture in the Board review course for endocrinologists

- 2003 Boston University Medical Center, Endocrine Grand Rounds,
Cholesterol trafficking and its role in atherosclerosis
- 2003 University of Kentucky, Gill Heart Center Cardiovascular Symposium
Innate Immunity and Atherosclerosis
- 2004 5th Annual Conference on Arteriosclerosis, Thrombosis and Vascular Biology,
San Francisco, CA
Innate immune signaling and atherosclerosis
- 2004 Endocrine Society: Meet the Professor. New Orleans, LA
The treatment of hyperlipidemia: beyond statins
- 2004 International Atherosclerosis Society, HDL Workshop, Heraklion, Crete
ABCA transporters
- 2004 Wake Forest Medical Center, Visiting Professor and lecturer
Lipid trafficking in macrophages
- 2004 American Heart Association, New Orleans, LA
Gene Expression Profiling for Complex Diseases
- 2005 Cell Biology of Atherosclerosis, Keystone Conference
Macrophages in Atherosclerosis
- 2005 Deuel Lipid Conference, Los Angeles, CA
ABCA transporters
- 1986-present Medical or Endocrine Grand Rounds or Departmental Seminars on topics in lipid
metabolism or atherosclerosis at universities and hospitals including Dartmouth College,
Harvard Biological Chemistry Department, Brigham and Women's Hospital, Boston
University, Rhode Island Hospital, Beth Israel Hospital, East Carolina University, Maine
Medical Center, University of Rochester, University of Vermont, University of Kentucky,
University of Colorado, and Univ. of California, San Francisco, and Yale Univ.

E. Report of Clinical Activities:

My clinical activities involve directing the Lipid Clinic at MGH and, from 1996 until early 2004, I also directed a community lipid clinic practice in New Bedford, MA. The MGH clinic is held every week while the New Bedford practice was conducted once per month. The MGH practice sees approximately 20 patients per session, with most patients having complex lipid disorders that have been referred by internists or cardiologists for specialty evaluation. The introduction of HMG-CoA reductase inhibitors to patients at the MGH was first accomplished in my Lipid Clinic during phase 2-3 trials and the clinic was involved in a critical investigation of lovastatin that established the safety of this class of drugs. The Lipid Clinic has served as a major training site for other physicians and nurse trainees, including the current director of the Brigham and Women's lipid clinic, as well as multiple nurse practitioners who have helped establish lipid clinics in the region.

I also serve as the attending physician on the Endocrine Division service for approximately one month a year and continue to attend on the general medical service (Bigelow) of the MGH Dept of Medicine for 2-4 weeks a year.

PART III: Bibliography

Original reports

1. **Freeman MW**, Spring-Mills E, and AL Jones. The effect of oxandrolone on low and high density lipoprotein profiles in retired breeder rats. *J Gerontol.* 1980;35:31-38.
2. Vasicek TJ, MeDevitt BE, **Freeman MW**, Hendy GN, Potts JT Jr, Rich A, and HM Kronenberg. Nucleotide Sequence of the human parathyroid hormone gene. *Proc Natl Acad Sci USA.* 1983;80:2127-2131.
3. Kronenberg HM, Igarashi T, **Freeman MW**, Okazaki T, Brand SJ, Wiren KM, and JT Potts Jr. Structure and expression of the human parathyroid hormone gene. *Rec Prog Horm Res.* 1986;42:641-663.
4. Born W, **Freeman MW**, Bornstein W, Rapoport A, Klein RD, Hendy GN, Khorana HG, Rich A, Potts JT Jr, and HM Kronenberg. Signal sequence of human preproparathyroid hormone is inactive in yeast. *J Bone Min Res.* 1987;2:353-361.
5. Born W, **Freeman MW**, Hendy GN, Rapoport A, Rich A, Potts JT Jr, and HM Kronenberg. Human preproparathyroid hormone synthesized in *E. coli* is transported to the surface of the bacterial inner membrane but not processed to the mature hormone. *Mol Endocrinology* 1987;1:5-14.
6. **Freeman MW**, Wiren KM, Rapoport A, Lazar M, Potts JT Jr, and HM Kronenberg. Consequences of amino-terminal deletions of preproparathyroid hormone signal sequence. *Mol Endocrinology* 1987;9:628-638.
7. Wiren KM, Ivashkiv L, Ma P, **Freeman MW**, Potts JT Jr, and HM Kronenberg. Mutations in signal sequence cleavage domain of preproparathyroid hormone alter protein translocation, signal sequence cleavage, and membrane-binding properties. *Mol Endocrinology* 1989;3:240-250
8. Abou-Samra AB, **Freeman MW**, Jueppner H, Uneno S, and GV Segre . Characterization of fully active biotinylated parathyroid hormone analogs: application to fluorescence activate cell sorting of parathyroid hormone receptor bearing cell. *J Biol Chem.* 1990;265:58-62.
9. Kodama T, **Freeman M**, Rohrer L, Zabrezky J, Matsudaira P, and M Krieger. Type I macrophage scavenger receptor contains alpha-helical and collagen-like coiled coils. *Nature* 1990;343:531-535.
10. Rohrer L, **Freeman M**, Kodama T, Penman M, and M Krieger. Coiled coil fibrous domains mediate ligand binding by macrophage scavenger receptor type II. *Nature* 1990;343:570-572.
11. Atkinson MJ, **Freeman MW**, and HMKronenberg. Thymosin β_4 is expressed in ROS 17/2.8 osteosarcoma cells in a regulated manner. *Mol Endocrinology* 1990;4:69-74.
12. **Freeman M**, Ashkenas J, Rees DJ, Kingsley DM, Copeland NG, Jenkins NA, and M Krieger. An ancient, highly conserved family of cysteine-rich protein domains revealed by cloning the type I and II murine macrophage scavenger receptors. *Proc Natl Acad Sci USA.* 1990;87:8810-8814.
13. **Freeman M**, Ekkel L, Rohrer L, Penman M, Freedman NJ, Chisholm GM, and M Krieger . Expression of type I and type II bovine scavenger receptors in Chinese hamster ovary cells: Lipid

droplet accumulation and nonreciprocal cross competition by acetylated and oxidized low density lipoprotein. *Proc Natl Acad Sci USA*. 1991;88:4931-4935.

14. Juppner H, Abou-Samra AB, **Freeman M**, Kong XF, Schipani E, Richards J, Kolakowski LF, Kronenberg HM, GV Segre. A G-protein linked receptor for parathyroid hormone and parathyroid hormone related peptide. *Science* 1991;254:1024-1025.
15. Segre GV, Abou-Samra AB, Juppner H, Schipani E, Force T, Urena P, **Freeman M**, Kong XF, Kolawshi LF Jr, Hock J, Bonventre J, Potts JT Jr, HM Kronenberg. Characterization of cloned PTH/PTHrP receptors. *Journal of Endocrinological Investigation* 1992;15:11-17.
16. Abou-Samra AB, Juppner H, Force T, **Freeman M**, Kong XF, Schipani E, Urena P, Richards J, Bonventre JV, Potts JT Jr, Kronenberg HM, and GV Segre. Expression cloning of a PTH/PTHrP receptor from rat osteoblasts-like cells: a single receptor stimulates intracellular accumulation of both cAMP and inositol triphosphates and increases intracellular free calcium. *Proc Natl Acad Sci USA*. 1992;89:2732-2736.
17. Bickel P and **MW Freeman**. Rabbit smooth muscle cells express an inducible scavenger receptor mRNA that is absent from endothelial cells. *J Clin Invest*. 1992;90:1450-1457.
18. Retsky KL, **Freeman MW**, and B. Frei. Ascorbic acid and dehydro ascorbic acid prevent oxidative modification of human low density lipoproteins. *J Biol Chem*. 1993;268:1304-1309.
19. Acton S, Resnick D, **Freeman MW**, Ekkel Y, Ashkenas J, and M Krieger. The collagenous domains of macrophage scavenger receptors and complement factor C1q mediate their similar, but not identical, binding specificities for polyanionic ligands. *J Biol Chem*. 1993;268:3530-3537.
20. Ashkenas J, Penman M, Vasile E, Acton S, **Freeman M**, and M Krieger. Structures and high and low affinity ligand binding properties of murine type I and type II macrophage scavenger receptors. *J Lipid Res*. 1993;34:983-1000.
21. Zoeller RA, Liu Y, Millham F, **Freeman MW**, and DT Golenbock. Surface expression of human CD 14 in Chinese Hamster Ovary fibroblasts transfers responsiveness to bacterial endotoxin. *J Biol Chem*. 1993;268:22055
22. Li H, **Freeman MW**, and P. Libby. Regulation of smooth muscle cell scavenger receptor expression *in vivo* by atherogenic diets and *in vitro* by cytokines. *J Clin Invest*. 1995;95:122-133.
23. Aftring RP and **MW Freeman**. Structure of the murine macrophage scavenger receptor gene and evaluation of sequences which regulate expression in the macrophage cell line, P388D1. *J Lipid Res*. 1995;36:1305-1314.
24. Christie R, **Freeman M**, and BT Hyman. Expression of the macrophage scavenger receptor, a multifunctional lipoprotein receptor, in microglia associated with senile plaques in Alzheimer disease. *Am.J.Pathol*. 1996;148:399-403.
25. Eberhart, G.P., Mendez, A.J., and **MW Freeman**. Decreased cholesterol efflux from fibroblasts of a patient without Tangier Disease, but with markedly reduced high density lipoprotein cholesterol levels. *J. Clin. Endo. Metabolism* 1998;83:836-846.

26. Andersson LA and **MW Freeman**. Functional changes in scavenger receptor binding conformation are induced by charge mutants spanning the entire collagen domain. *J. Biol. Chem.* 1998;273:19592-19601.
27. Moore KJ, Fabunmi RP, Andersson LP and **MW Freeman**. In vitro differentiated embryonic stem cell macrophages: a model system for studying atherosclerosis-associated macrophage functions. *Arterioscler Thromb Vasc Biol* 1998;18:1647-1654.
28. Chiang N, Gronert K, Clish, CB, O'Brien JA, **Freeman MW** and C.S. Serhan. Leukotriene B4 receptor transgenic mice reveal novel protective roles for lipoxins and aspirin-triggered lipoxins in reperfusion *J Clin Invest* 1999; 309-316.
29. Fabunmi RP, Moore KJ, Libby P, and **MW Freeman**. Stromelysin-1 (MMP-3) expression driven by a macrophage-specific promoter results in reduced viability in transgenic mice. *Atherosclerosis* 1999; 148:375-386.
30. Fitzgerald M, Moore KJ, **MW Freeman**, GL Reed. Lipopolysaccharide induces scavenger receptor-A expression in murine macrophages via a non-transcriptional mechanism. *J Immunol.* 2000; 164:2692-2700.
31. Brousseau ME, Schaefer EJ, Dupuis J, Eustace B, Van Eerdewegh P, Yasek-McKenna D, O'Neill G, Eberhart GP, Weiffenbach B, **Freeman MW**, Brown RH, JZ Gu. Novel mutations in the gene encoding ATP-binding cassette 1 in four Tangier Disease kindreds. *J Lipid Res* 2000; 41:433-441.
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