

BIOGRAPHICAL SKETCH

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NAME Xiao Xiao	POSITION TITLE Professor		
eRA COMMONS USER NAME XiaoXiao			
EDUCATION/TRAINING <i>(Begin with baccalaureate or other initial professional education, such as nursing, and include postdoctoral training.)</i>			
INSTITUTION AND LOCATION	DEGREE <i>(if applicable)</i>	YEAR(s)	FIELD OF STUDY
Shanghai Medical University, China	B.S.	1982	Pharmaceutical Chem.
Wuhan University, China	M.S.	1985	Biochemistry
University of Pittsburgh	Ph.D.	1992	Molecular Biology

A. Positions and Honors

- 2006 Eshelman Distinguished Professor, Division of Molecular Pharmaceutics, School of Pharmacy and Gene Therapy Center, Univ. of North Carolina at Chapel Hill.
- 2003-2006 Associate Professor with tenure, Dept. of Molecular Genetics & Biochemistry and Dept. of Orthopedic Surgery, Univ. of Pittsburgh
- 1998-2002 Tenure-track Assistant Professor, Dept. of Molecular Genetics & Biochemistry and Dept. of Orthopedic Surgery, Univ. of Pittsburgh
- 1997 Research Associate Professor, School of Pharmacy and Gene Therapy Center, Univ. of North Carolina at Chapel Hill.
- 1994-1997 Senior Scientist, Somatix Therapy Corporation, California, Visiting Scientist, Univ. of North Carolina at Chapel Hill, Gene Therapy Center.
- 1993 Associate Scientist and Scientist, Avigen Inc., California.
- 1989 Teaching Assistant for Genetics, Dept of Biological Sciences, Univ. of Pittsburgh.
- 1988 Teaching Assistant, Biology Lab Course, Dept of Biological Sciences, Univ. of Pittsburgh.
- 1986-1987 Research Assistant, Department of Molecular Biology, Vanderbilt University.
- 1985-1986 Instructor, taught Lab Molecular Biology, Dept. of Biology, Wuhan University, China.

B. Selected publications (from a total of > 100)

- 1 **Xiao, X.**, Li, J. and Samulski, R.J.(1998) Production of high titer recombinant AAV vectors in the absence of helper adenovirus. *J. Virol.* 72: 2224-2232.
- 2 During MJ, Samulski RJ, Elsworth JD, Kaplitt MG, Leone P, **Xiao X**, Li J, Freese A, Taylor JR, Roth RH, Sladek Jr. JR, O'Malley KL, and Redmond Jr. DE.(1998) In vivo expression of therapeutic human genes for dopamine production in the caudates of MPTP-treated monkeys using an AAV vector., *Gene Therapy*, , 5: 820-827
- 3 Rabinowitz J E., **Xiao X.**, Samulski R.J. (1999) Adeno-associated virus: A new tool for cancer gene therapy *Hum Gene Ther* 10: 837
- 4 Li, J., D Dressman, YP Tsao, A Sakamoto, T Toyo-oka, EP. Hoffman and **X Xiao**. (1999) rAAV Vector Mediated sarcoglycan Gene Transfer In A Hamster Model For Limb Girdle Muscular Dystrophy, *Gene Therapy*. 6: 74-8
- 5 R-Y. Pan, **X Xiao**, S-L Chen, J Li, L-CLin, H-J Wang, Y-P Tsao, (1999) Disease-inducible transgene expression from a recombinant adeno-associated viral vector in a rat arthritis model, *J. Virol.* 73:3410-17
- 6 **Xiao, X**, Li, J., Tsao, YP., Dressman, D., Hoffman, EP. and Watchko, J. (2000) Full functional rescue of a complete muscle (TA) in dystrophic hamsters by adeno-associated virus vector-directed gene therapy, *J. Virol.*74:1436-1442

- 7 Pruchnic, R., Cao, BH., Qu, Z., **Xiao, X.**, Li, J., Samulski, RJ., and Huard, J. (2000) The use of adeno-associated virus to circumvent the maturation dependent viral transduction of muscle fibers, *Human Gene Therapy*, 11:521-536.
- 8 Keir, SD., House, SB., Li, J., **Xiao, X.**, and Gainer, H. (1999) Gene Transfer into hypothalamic organotypic cultures using an adeno-associated virus vector, *Exp. Neurol*, 160(2):313-6
- 9 Pan, RY., Chen, SL., **Xiao, X.**, Liu, DW., Peng, HJ., Tsao, YP., (2000) Therapy and prevention of arthritis by recombinant adeno-associated virus vector with delivery of interleukin-1 receptor antagonist, *Arthritis and Rheumatism*, 43:289-97
- 10 Liu, DW., Tsao, YP., Kung, JT., Ding, YA., Sytwu, HK., **Xiao, X.**, and Chen, SL. (2000) Recombinant adeno-associated virus expressing human papillomavirus 16 E7 peptide DNA intramurally fused with heat shock protein DNA as a tumor vaccine for cervical cancer, *J. Virol.* 74(6):2888-94
- 11 Watchko, J., Li, J., Hoffman, EP., and **Xiao, X.** (2000) Contractile properties of Bio14.6 delta-sarcoglycan deficient hamster tibialis anterior muscle, *Muscle & Nerve*, in revision.
- 12 Sun, LW., Li, J. and **Xiao, X.** (2000) Overcoming the size limitation of adeno-associated viral vectors by heterodimerization, *Nature Medicine*, 6:599-602.
- 13 Wang, B., Li, J. and **Xiao, X.** (2000) AAV vector carrying novel human mini-dystrophin genes effectively ameliorates muscular dystrophy in mdx mouse model, *Proc. Natl. Acad. Sci.* 97:13714-1371
- 14 Jiang, XC., S Qin, CP Qiao, A Skold, K Kawano, M Lin, **X Xiao** & AR. Tall (2001) Decreased ApoB Secretion and Atherosclerosis in Mice with Phospholipid Transfer Protein Deficiency, *Nature Medicine*, 7:847-852
- 15 David L. Silver, Nan Wang, **Xiao Xiao** and Alan R. Tall. (2001) HDL particle uptake mediated by SRBI results in selective sorting of HDL cholesterol from protein and polarized cholesterol secretion. *J. Biol. Chem.* 276:25287-93
- 16 Keir SD, **Xiao X**, Li J, Kennedy PG. (2001) Adeno-associated virus-mediated delivery of glial cell line-derived neurotrophic factor protects motor neuron-like cells from apoptosis. *J. Neurovirol* 7:437-46
- 17 Chen., Z., Lu, L., Li, J., Li, W., Fung, J.J., **Xiao, X.**, Qian S. (2001) Transfection with genes encoding CTLA4-Ig mediated by adeno-associated virus vectors prolongs survival of heart allografts. *Transplant Proc.* 33(1-2):604
- 18 Cao, B., R. Pruchnic, M. Ikezawa, **X. Xiao**, Juan Li, T. J Wickham, I. Kovesdi, W. A. Rudert, and J. Huard (2001) The role of receptors in the maturation-dependent adenoviral transduction of myofibers, *Gene Therapy*, 11(4):521-36
- 19 Wang, Z., Qiu SJ., Ye, SL., Tang, ZY, and **Xiao X** (2001) Combined IL-12 and GM-CSF Gene Therapy for Murine Hepatocellular Carcinoma, *Cancer Gene Therapy*, 8:751-758
- 20 Chao, H., Sun, L., Liu, Y., Bruce, A., **Xiao, X.**, and Walsh, CE (2002) Dimerization of rAAV vectors: a novel method for human factor VIII packaging and expression, *Molecular Therapy* 11(4):521-36
- 21 DeMiguel, F., Lou W., Skold, A., **Xiao, X.**, Gao, A. (2002) Stat3 enhances androgen receptor ligand independent activation, *Prostate* 52:123-9
- 22 Ma, H., Lin, SZ., Tsao, YP., Li, J., Chiang, YC., Chen, SL. **Xiao, X.**, (2002) Intratumoral antiangiogenesis gene therapy for glioma using an AAV vector carrying angiostatin cDNA, *Gene Therapy*, 9:2-11
- 23 Qiao, CP., Skold, A., Li, J., and **Xiao, X.** (2002) inducible packaging cell lines for adeno-associated viral (AAV) vectors, *J. Virol.*, 9: 2-11
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- 26 Watchko J., T. O'Day, B. Wang, L. Zhou, Y. Tang, J. Li, and Xiao X. (2002) AAV Vector-Mediated Mini-Dystrophin Gene Therapy Improves Dystrophic Muscle Contractile Function in mdx Mice *Human Gene Therapy*, 13:1451-1460
- 27 CP. Qiao, B. Wang, XD. Zhu, Juan Li and **Xiao X.** (2002) A novel gene expression control system and its use in high-titer and stable 293 cell-based adeno-associated virus packaging cell lines, *J. Virology* 76: 13015-13027

- 28 K. Yuasa, M. Sakamoto, Y. Suzuki, A. Tanouchi, H. Yamamoto, Juan Li, JS. Chamberlain, **X. Xiao** and S. Takeda (2002) Adeno-associated virus vector-mediated gene transfer into dystrophin-deficient skeletal muscles encounters elevated immune response against the transgene product, *Gene Therapy*, 9:1576-1588
- 29 Tsai TH, Chen SL, **Xiao X**, Liu DW, Tsao YP (2002) Gene therapy for treatment of cerebral ischemia using defective recombinant adeno-associated virus vectors. *Methods*. 28:253-258
- 30 Tsai TH, Chen SL, **Xiao X**, Chiang YH, Lin SZ, Kuo SW, Liu DW, Tsao YP. (2003) Related Articles, Links Abstract Gene treatment of cerebral stroke by rAAV vector delivering IL-1ra in a rat model. *Neuroreport*. 14:803-7.
- 31 Yang XP, Yan D, Qiao C, Liu RJ, Chen JG, Li J, Schneider M, Lagrost L, **Xiao X**, Jiang XC (2003) Increased Atherosclerotic Lesions and Lipoprotein Oxidizability in Apolipoprotein E-Null Mice Overexpressing Plasma Phospholipid Transfer Protein. *Arterioscler Thromb Vasc Biol*. Jul y 10 issue
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- 43 Zhu T., L. Zhou, S. Mori, Z. Wang¹, CF. McTiernan, C. Qiao, C. Chen, D. Wang, J. Li, & **X Xiao** (2005) Sustained whole-body functional rescue by gene transfer in congestive heart failure and muscular dystrophy hamsters, *Circulation*, 112:2650-9
- 44 X. Ye, T. Zhu, S. Bastacky, T. McHale, J. Li, **X. Xiao** (2005) Prevention and reversal of lupus in NZB/W mice by costimulatory inhibitor gene transfer with adeno-associated viral (AAV) vectors, *Rheumatism and Arthritis*, 52:3975-86
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- 46 Z. Wang, T. Zhu, KK. Rehman, S. Bertera, J. Zhang, CL. Chen, G. Papworth, S. Watkins, M. Trucco, PD. Robbins, J. Li, and **X. Xiao** (2006) Widespread and stable pancreatic gene transfer by AAV vectors via different routes, *Diabetes*, 5(4):875-84

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- 48 Draviam RA, Wang B, Shand SH, **Xiao X**, Watkins SC. (2006) Alpha-sarcoglycan is recycled from the plasma membrane in the absence of sarcoglycan complex assembly. *Traffic*. 7(7):793-810.
- 49 Chen CC, Ko TM, Ma HI, Wu HL, **Xiao X**, Li J, Chang CM, Wu PY, Chen CH, Han JM, Yu CP, Jeng KS, Hu CP, Tao MH. (2006) Long-term inhibition of hepatitis B virus in transgenic mice by double-stranded adeno-associated virus 8-delivered short hairpin RNA. *Gene Ther*. 14:11-9
- 50 Lai LJ, **Xiao X**, Wu JH. (2007) Inhibition of corneal neovascularization with endostatin delivered by adeno-associated viral (AAV) vector in a mouse corneal injury model. *J Biomed Sci*. 2007 Mar 21; [Epub ahead of print]
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- 52 Yuan G, Deng J, Wang T, Zhao C, Xu X, Wang P, Voltz JW, Edin ML, **Xiao X**, Chao L, Chao J, Zhang XA, Zeldin DC, Wang DW. (2007) Tissue Kallikrein Reverses Insulin Resistance and Attenuates Nephropathy in Diabetic Rats by Activation of PI3 kinase/Akt and AMPK Signaling Pathways. *Endocrinology*. 2007 Feb 1.
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RESEARCH SUPPORT

Active funding:

1U01 NS 46546 (Xiao) NIH/NINDS 1.8 cal. mos. 09/01/04-5/31/08

AAV mini-dystrophin for DMD gene therapy (in no-cost extension)

The overall goal of this proposal is to design , optimize and produce a recombinant AAV vector carrying a mini-dystrophin gene which will subsequently be tested for efficacy and safety studies as required by the FDA for future clinical studies.

1R01AR050595 (Xiao) NIH/NIAMS 1.8 cal. mos. 4/1/04-3/31/09

Gene therapy for a severe DMD mouse model

The major goal of this project is to use transgenic mouse technology and gene transfer technology to test the functions of mini-dystrophin genes and its ability to rescue a severe DMD mouse model, which has both utrophin and dystrophin gene mutations, the double KO model.

2R01AR45967 (Xiao) NIH/NIAMS 1.8 cal. mos. 10/1/04-7/31/09

AAV vectors for heart and muscle gene therapy

The major goal of this project is to use AAV vector to treat heart failure and limb-girdle muscular dystrophy in Bio14.6 and TO-2 hamster models, which have a mutation in the delta-sarcoglycan gene.