

Wanling YANG

Associate Professor (With Tenure)

Department of Paediatrics and Adolescent Medicine
 LKS Faculty of Medicine, The University of Hong Kong
 L7-55 Laboratory Block, 21 Sassoon Road, Hong Kong
 Email: yangwl@hku.hk; Tel: (852)283-15077; Fax: (852)2819-8142
 Lab webpage: <http://paed.hku.hk/genome>

EDUCATION:

- Ph.D., Pharmacology (minor in Bioinformatics), University of Minnesota, Minneapolis, USA (2002)
- Master of Medicine, Jiamusi Medical College, China (1989)

Research interest:

1. Association studies of complex diseases, especially autoimmune diseases
2. Genetic studies of Mendelian disorders using next generation sequencing (NGS) tools
3. Comprehensive and integrative analyses of various types of omics data and databases to understand the disturbances of signal transduction pathways and interaction networks underlying human diseases

RESEARCH AND TEACHING EXPERIENCE:

July 2014 – Present	Associate Professor, Department of Paediatrics and Adolescent Medicine, The University of Hong Kong
Sep. 2008 – June 2014	Assistant Professor, Department of Paediatrics and Adolescent Medicine, The University of Hong Kong
Jan. 2006 – Aug. 2008	Research Assistant Professor and Honorary Assistant Professor, Department of Paediatrics and Adolescent Medicine, The University of Hong Kong
Jul. 2002 – Jan. 2006	Postdoctoral Research Fellow (with John D. Hildebrandt), Department of Pharmacology, Medical University of South Carolina, Charleston, South Carolina, USA
May 1995 – Jan. 1997	WHO Fellow in Neurobiology, Department of Pharmacology, University of Minnesota, Minneapolis, Minnesota, USA
Jul. 1989 – Apr. 1995	Lecturer, Jiamusi Medical College, China

PUBLICATIONS:

Book chapter:

Yang W, and Hildebrandt, J.D. (2013) Gq Family. In: Lennarz, W.J. and Lane, M.D. (eds.) The Encyclopedia of Biological Chemistry, Vol. 2, pp. 496-500. Waltham, MA: Academic Press, invited and editor-reviewed.

Journal publications: (from Pubmed: <http://paed.hku.hk/genome>)

(*: Corresponding or co-corresponding author. #: Equal contribution/co-first author.)

1. Yeung KS, Chung BH, Choufani S, Mok MY, Wong WL, Mak CC, **Yang W**, Lee PP, Wong WH, Chen YA, Grafodatskaya D, Wong RW, Lau CS, Chan DT, Weksberg R, Lau YL. Genome-Wide DNA Methylation Analysis of Chinese Patients with Systemic Lupus Erythematosus Identified Hypomethylation in Genes Related to the Type I Interferon Pathway. **PLoS One**. 2017 Jan 13;12(1):e0169553.
2. Zhou XJ, Nath SK, Qi YY, Sun C, Hou P, Zhang YM, Lv JC, Shi SF, Liu LJ, Chen R, **Yang W**, He KZ, Li Y, Zhang H. Novel identified associations of RGS1 and RASGRP1 variants in IgA Nephropathy. **Sci Rep**. 2016 Nov 2;6:35781.
3. Kwok J, Guo M, **Yang W**, Lee CK, Ho J, Tang WH, Chan YS, Middleton D, Lu LW, Chan GC. HLA-A, -B, -C, and -DRB1 genotyping and haplotype frequencies for a Hong Kong Chinese population of 7595 individuals. **Hum Immunol**. 2016 Dec;77(12):1111-1112.
4. Kwok J, Guo M, **Yang W**, Lee CK, Chan NK, Ho J, Tang WH, Chan YS, Middleton D, Lu LW, Chan GC. HLA-A, -B and -DRB1 genotyping and haplotype frequencies of 3892 cord blood units in the Hong Kong Chinese Cord Blood Registry. **Hum Immunol**. 2016 Dec;77(12):1109-1110.
5. Chen R, Lau YL, Zhang Y, **Yang W**. SRinversion: a tool for detecting short inversions by splitting and re-aligning poorly mapped and unmapped sequencing reads. **Bioinformatics**. 2016 Dec 1;32(23):3559-3565.
6. Zhang F, Wu L, Qian J, Qu B, Xia S, La T, Wu Y, Ma J, Zeng J, Guo Q, Cui Y, **Yang W**, Huang J, Zhu W, Yao Y, Shen N, Tang Y. Identification of the long noncoding RNA NEAT1 as a novel inflammatory regulator acting through MAPK pathway in human lupus. **J Autoimmun**. 2016 Dec;75:96-104.
7. Jiang S, Tang L, Zhao N, **Yang W**, Qiu Y, Chen HZ. A Systems View of the Differences between APOE ε4 Carriers and Non-carriers in Alzheimer's Disease. **Front Aging Neurosci**. 2016 Jul 12;8:171.
8. #Morris, D.L., #Sheng, Y., #Zhang, Y., #Wang, Y.F., #Zhu, Z., #Tomblason, P., Chen, L., Graham D.S.C., Bentham, J., Chen, R., Zuo, X., Wen, L., Yang, C., Liu, L., Yang, L., Li, F., Huang, Y., Yang, S., Rönnblom, L., Fürtrohr, B.G., Voll, R.E., Schett, G., Costedoat-Chalumeau, N., Gaffney P.M., Lau, Y.L., Zhang, X., **Yang, W.**, #Cui, Y., & #Vyse, T.J. Meta-analysis of Chinese and European GWAS identifies 10 novel SLE associated loci and provides evidence for increased genetic risk of disease in Chinese. **Nature Genetics** (in press 2016) (co-corresponding author)

9. Chiu YT, Wong JK, Choi SW, Sze KM, Ho DW, Chan LK, Lee JM, Man K, Cherny S, **Yang W**, Wong CM, Sham PC, Ng IO. Novel pre-mRNA splicing of intronically integrated HBV generates oncogenic chimera in hepatocellular carcinoma. **J Hepatol.** 2016 Feb 8. pii: S0168-8278(16)00084-2. doi: 10.1016/j.jhep.2016.02.005.

10. Leng RX, Pan HF, Liu J, Yang XK, Zhang C, Tao SS, Wang DG, Li XM, Li XP, **Yang W**, Ye DQ. Evidence for genetic association of TBX21 and IFNG with systemic lupus erythematosus in a Chinese Han population. **Sci Rep.** 2016 Feb 26;6:22081.

11. Zhu Z, Liang Z, Liany H, Yang C, Wen L, Lin Z, Sheng Y, Lin Y, Ye L, Cheng Y, Chang Y, Liu L, Yang L, Shi Y, Shen C, Zhou F, Zheng X, Zhu J, Liang B, Ding Y, Zhou Y, Yin X, Tang H, Zuo X, Sun L, Bei JX, Liu J, Yang S, ***Yang W**, ***Cui Y**, ***Zhang X**. Discovery of a novel genetic susceptibility locus on X chromosome for systemic lupus erythematosus. **Arthritis Res Ther.** 2015 Dec 3;17:349.

12. Jing Zhang, Lu Zhang, Yan Zhang, Jing Yang, Mengbiao Guo, Liangdan Sun, Hai-Feng Pan, Nattiya Hirankarn, Dingge Ying, Shuai Zeng, Tsz Leung Lee, Chak Sing Lau, Tak Mao Chan, Alexander Moon Ho Leung, Chi Chiu Mok, Sik Nin Wong, Ka Wing Lee, Marco Hok Kung Ho, Pamela Pui Wah Lee, Brian Hon-Yin Chung, Chun Yin Chong, Raymond Woon Sing Wong, Mo Yin Mok, Wilfred Hing Sang Wong, Kwok Lung Tong, Niko Kei Chiu Tse, Xiang-Pei Li, Yingyos Avihingsanon, Pornpimol Rianthavorn, Thavatchai Deekajorndej, Kanya Suphapeetiporn, Vorasuk Shotelersuk, Shirley King Yee Ying, Samuel Ka Shun Fung, Wai Ming Lai, Maria-Mercè Garcia-Barceló, Stacey S. Cherny, Pak Chung Sham, Yong Cui, Sen Yang, Dong Qing Ye, Xue-Jun Zhang, ***Yu Lung Lau**, ***Yang W**, Gene-based meta-analysis of GWAS data identifies independent SNPs in ANXA6 as associated with SLE in Asian populations, **Arthritis Rheumatol.** 2015 Nov;67(11):2966-77. doi: 10.1002/art.39275.

13. Tam RC, Lee AL, **Yang W**, Lau CS, Chan VS. Systemic Lupus Erythematosus Patients Exhibit Reduced Expression of CLEC16A Isoforms in Peripheral Leukocytes. *Int J Mol Sci.* 2015 Jun 25;16(7):14428-40. doi: 10.3390/ijms160714428. PubMed PMID: 26121298.

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PubMed Central PMCID: PMC4198120.

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32. **Yang W**, Tang H, Zhang Y, Tang X, Zhang J, Sun L, Yang J, Cui Y, Zhang L, Hirankarn N, Cheng H, Pan HF, Gao J, Lee TL, Sheng Y, Lau CS, Li Y, Chan TM, Yin X, Ying D, Lu Q, Leung AM, Zuo X, Chen X, Tong KL, Zhou F, Diao Q, Tse NK, Xie H, Mok CC, Hao F, Wong SN, Shi B, Lee KW, Hui Y, Ho MH, Liang B, Lee PP, Cui H, Guo Q, Chung BH, Pu X, Liu Q, Zhang X, Zhang C, Chong CY, Fang H, Wong RW, Sun Y, Mok MY, Li XP, Avihingsanon Y, Zhai Z, Rianthavorn P, Deekajorndej T, Suphapeetiporn K, Gao F, Shotelersuk V, Kang X, Ying SK, Zhang L, Wong WH, Zhu D, Fung SK, Zeng F, Lai WM, Wong CM, Ng IO, Garcia-Barceló MM, Cherny SS, Shen N, Tam PK, Sham PC, Ye DQ, Yang S, Zhang X, Lau YL. Meta-analysis followed by replication identifies loci in or near CDKN1B, TET3, CD80, DRAM1, and ARID5B as associated with systemic lupus erythematosus in Asians. *Am J Hum Genet.* 2013 Jan 10;92(1):41-51. doi:10.1016/j.ajhg.2012.11.018. Epub 2012 Dec 27. PubMed PMID: 23273568; PubMed Central PMCID: PMC3542470.
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SUMMARY and Research Plan

Research:

My research is on genetic studies of complex diseases and Mendelian disorders, and development of methodologies in analyzing genetics and genomics data, two parallel tracks involving genomics and bioinformatics.

Research on Complex diseases:

I have been working on the genetics of systemic lupus erythematosus (SLE), an autoimmune disease that mainly affects women of child-bearing age. We have performed genome-wide association study (GWAS) on SLE and uncovered 8 novel susceptibility loci for this disease, some of which are likely to be population-specific. I have published more than a dozen papers on lupus, depicting genetic factors particularly relevant to Asian populations. I have on-going collaborations with renowned researchers world-wide including Dr. Mark Lathrop from McGill, and Dr. Tim Vyse from KCL. I would like to continue this research on complex diseases including but not limited to SLE depending on collaborations.

Rare diseases and NGS data analysis:

I have also worked on single gene disorders diagnosed in the Department of Paediatrics and Adolescent Medicine and other departments in Queen Mary Hospital, Hong Kong, such as primary immunodeficiency, using next-generation sequencing (NGS). We are among the first to identify IL10RA mutations causal to infant-onset Crohn's disease and have made molecular diagnosis for a number of patients using this cutting-edge technology. We also published tools in facilitating whole exome sequencing data analysis (PriVar, Zhang et al, Bioinformatics). I would like to continue to work on perfecting analysis for NGS data, combining genetics, molecular biology, population databases and population genetics, pathway analysis and gene expression profiles etc. to help make a comprehensive analysis and understanding of the data. This is very much a multidisciplinary field and my expertise in genetics and genomics and experience in collaborating with statisticians and computer scientists will definitely help to make headways in this field.

Development of bioinformatics tools

My laboratory has developed two programs, HRRA (*Hum Mut*, 2011) and HaploShare (in revision with *Genome Biology*), to help detect recent founder mutations that are difficult to detect using linkage analysis or association studies. We also developed software packages such as EFIN and PriVar to facilitate analysis of whole exome sequencing data. Cancer genomics is also an interest for me if suitable collaborators can be found in this area.

Applicational genomics:

We also work to apply genomics to clinical diagnosis, testing and screening and have developed a barcoding system that is particularly suitable to screen a number of genes on thousands of samples using NGS technology. We made progress on calling HLA types using NGS data and are developing a novel method on HLA typing using NGS sequencing. Genomics is both for research and clinical application. I will continue to work on applying genomics to clinical diagnosis and population screening, to make genomics really translational.

I have limited bioinformatics training myself but have been able to assemble a laboratory composed of a number of bioinformatics students, and we are able to use/modify tools for genetic and genomic data analysis and to develop tools ourselves in this field. I have been collaborating with research scientists from computer science and statistics in my own research and tool development. I also have a broad range of collaborations with molecular biologists and clinician scientists to help their data analysis and research. I believe I am in the best position for genomics, bioinformatics and interdisciplinary research, being a biologist with in-depth understanding of the questions but also with a good understanding of different languages, and am able to collaborate and interact with scientists from different disciplines.

Research interest:

Dr. Yang's research focuses on genetic studies of complex diseases such as SLE using GWAS and of single gene disorders using whole exome sequencing (WES), and on developing bioinformatics tools to improve analysis of genomic data. His laboratory, in collaboration with others, identified *ETS1*, *WDFY4*, *CDKN1B*, *CD80*, *ARID5B*, *TET3*, *DRAMI* and *ELF1* as associated with SLE, and many of these susceptibility genes are believed to be specific to Asians, pointing to the necessity of developing population-based approaches in risk prediction and personalized treatment of complex diseases. His lab also developed tools for detecting recent founder mutations based on GWAS data (HRRA, HaploShare) and facilitating mutation identification from whole exome sequencing (PriVar, EFIN). His recent focus is on developing tools for applying genomics to clinical diagnosis and population screening, such as HLA typing using next generation sequencing.

Editorship and Referee of scholarly journals

Served as *ad hoc* referee for the following international peer-review journals in the last 4 years (*ad hoc*, multiple times for most of the listed journals):

1. **American Journal of Human Genetics**
2. Annals of Human Genetics
3. Arthritis Research and Therapy
4. Artificial intelligence in Medicine
5. Autoimmunity
6. Briefings in Bioinformatics
7. Behavior Genetics
8. BMC Bioinformatics
9. Clinica Chimica Acta
10. DNA and Cell Biology
11. **Human Molecular Genetics**
12. Human Genetics
13. Immunologic Research (IMRE)
14. International Journal of Immunogenetics
15. International Journal of Rheumatic Diseases
16. Journal of Cellular and Molecular Medicine
17. Journal of Human Genetics
18. Journal of Rheumatology
19. Molecular Biology Reports
20. Mutation Research/Fundamental and Molecular Mechanisms of Mutagenesis
21. **Nature Genetics**
22. **PLoS Genetics**
23. PLoS One
24. Tissue Antigens
25. Tumor Biology

Invited Presentations:

1. Seoul 2015 invited speaker, 2015 International Symposium on Statistical Genetics, Seoul, May 27-28, 2015 Integrative Analysis of GWAS data to Understand Disease Mechanism for Systemic Lupus Erythematosus
2. Kyoto Course and Symposium on Bioinformatics for NGS with Applications in Human Genetics “Target sequencing analysis of HLA loci (course)” and “GWAS and Beyond: understanding the disease mechanisms of SLE through integrative data analysis (symposium)”
3. Susceptibility genes for SLE: what do we know and what do they mean. 4th International Symposium on Frontiers in Cardiovascular Diseases, June 4th, 2013 Wuhan, China
4. Lupus genetics. International Dermatogenetics Workshop, June 19-20, 2013 Chengdu, China
5. SLE: what do we know and what do they mean. NPG Asia-Pacific: From GWAS to Precision Medicine Workshop, May 16-18, 2013, Shanghai, China
6. Genomics and Bioinformatics in Clinical Diagnostics: the Present and the Future, May 15, KinMed Diagnostics, Guangzhou, China
7. Teaching week: Genetics, Genomics, and Bioinformatics and the Future of the Personalized Medicine. College of Life Sciences, Fujian University of Agriculture and Forestry, April 14-19, 2013
8. Data analysis in a geneticist’s eye. International Society on Computational Biology (ISCB)-Asia 2012, Shenzhen, China, Dec 19th, 2012
9. Data analysis in modern genetic studies. Shandong University Medical School, Oct 18-19th, 2012
10. Medical Genetics and Genetic data analysis. Southeastern University, Nanjing China, Oct. 10th-11st, 2012
11. Bioinformatics, genomics and the future of genomic medicine. 5th International Workshop and Summer School on Crops, Chips and Computers, Sep 9th, 2012, Fuzhou, China
12. Invited Lecture Series: Application of new sequencing technologies and bioinformatic data analysis to clinical genetics, Dec 2011-jan. 2012 (Invited teacher of a commissioned training course), Clinical Genetics Service, Department of Health, the Government of Hong Kong
13. The 12th International Congress of Human Genetics, Montreal Canada, Nov 2011, Selected for Platform Oral Presentation: Finding rare variants in GWAS and exome sequencing data by making use of recent common founder information
14. Asia Pacific Conference in Medical Genetics, Hong Kong, Nov 4th, 2010: Identification of compound heterozygous mutations in an early onset Crohn’s disease patient using whole exome sequencing technology, invited presentation
15. Frontiers in Biomedical Research HKU2010, Mendelian diseases, complex diseases and those in between—and their study approaches, invited presentation
16. The International Conference on BioMedical Engineering and Informatics; 2008, Sanya, Hainan, China, Biomarker identification for early tumor detection aided by bioinformatics gene expression analysis, invited presentation
17. Joint Symposium on “New paradigms in Medical Genetics”, Nov. 24-26, 2006, Hong Kong, Linkage analysis with dense SNP genotyping, invited presentation
18. Genomics Symposium at Experimental Biology 2005 by FASEB MARC program on April 4th, 2005, Characterization of a recombination hotspot in the GNAS locus previously associated with hypertension, invited presentation

Memberships:

- Member, Strategic Research Theme of University Research Committee, HKU, Genomics
- Member, American Society of Human Genetics

Honors and Awards:

- World Health Organization Fellowship, Neurobiology (1995);
- LM. Nutter Research Award, Dept of Pharmacology, University of Minnesota (2001)
- Buccaneer Research Award, Minnesota Medical Foundation (2002).

Research Grants:**General Research Fund (major funding form in Hong Kong, externally peer reviewed, in a PI's capacity):**

- 17146616 Unravelling the genetic puzzle of systemic lupus erythematosus (SLE) through comprehensive analyses of genetic association by functional annotation of genes, variants, and genomic regions, \$1,016,800, Oct 2016-Oct 2019
- 17125114 Meta-Analysis of three GWAS datasets from Hong Kong, Anhui China and UK followed by replication in independent cohorts in Asia and UK for identification of novel susceptibility genes associated with SLE, \$1,098,000
- Mapping Susceptibility Genes for Systemic Lupus Erythematosus (SLE) through a Genome-Wide Association Study, 2009-2011, HKU781709 M, \$755,760.
- Meta-Analysis of two Asian GWAS datasets for identification of novel genetic variants associated with systemic lupus erythematosus, 2011-2013, HKU784611M, \$650,000.
- Identifying susceptibility genes in the human MHC region independently associated with systemic lupus erythematosus (SLE) in Chinese populations. HKU783813M, 2013-2015, \$ 822,731

General Research Fund in a co-I's capacity:

- HKU 770411- Identification of coding variants associated with early-onset and familial systemic lupus erythematosus through whole exome sequencing and replication, 2011-2013, \$747,500.
- HKU777511M-Method and Software for Personal Risk Profiling of Complex Diseases, 2011-2013, \$1086, 750. This is a study that proposes to make use of available GWAS findings to predict disease risks for the general population. (PI: Prof Pak Sham).
- HKU 765311M, Genome-wide DNA methylation profiling in Chinese patients with Systemic Lupus Erythematosus (SLE), 2011-2013, \$805,000. This project proposes to study the role of methylation on disease risk of SLE, trying to bridge the effect of genetics and environmental factors. (PI: Dr. Brian Chung).
- Design and Analysis of Algorithms for Finding the Maximum Quasi-Biclique, Perfect Duplication History and Identification of Linked Regions, HK\$ 497,084 2009-2011 (PI: Prof. Wang LS, Department of Computer science, City U HK).
- HKU 776412M -Bioinformatics tools for identifying disease loci from exome sequencing data, 2012-2014, HK\$1,098,000 (PI: Prof Pak Sham).
- Cell-lineage specific differential DNA methylation in Chinese patients with early-onset systemic lupus erythematosus (SLE): implications for disease susceptibility. 2013-2015 (PI: Dr. Brian Chung)

Other Peer Reviewed Grants as PI:

- HMRF Molecular diagnosis for severe combined immunodeficiencies (SCID) using whole exome sequencing, PI, External, peer-reviewed grant. (project no.: 01120846), \$980,000, 01/02/2014—31/01/2016
- 12133701 HMRF, Meta-analysis of SLE GWAS followed by replication on X chromosome in cross-ethnic populations \$395,000.00 1/03/2015----28/02/2017

CURRICULUM DEVELOPMENT:

- Genome Bioinformatics, Bioinformatics Program 2009 and 2010
- Common Core Course: Genetics and Human Nature
- BBMS2003 Human Genetics, 2014, 2015, 2016, 2017
- BBMS3009 Genome Science, 2015, 2016, 2017