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### Present position

2012-present Associate Research Scientist, Institute of Biomedical Science, Academia Sinica

2005-present Assistant Director, National Center for Genome Medicine at Academia Sinica

2005-2011 Assistant Research Scientist, Institute of Biomedical Science, Academia Sinica

### Education and Training

Education / Training:			
Institution and Location	Degree	Year	Field of study
Graduate program of Molecular and Cellular Biology, SUNY at Stony Brook, USA	Ph.D.	1993-1999	Developmental Biology
Institute of Genetics, National Yang-Ming University, Taiwan	Post-doctoral fellow	1999-2000	Cancer Genetics
Division of Molecular and Genomic Medicine, National Health Research Institutes, Taiwan	Post-doctoral fellow	2001-2005	Cancer Genomics
Institute of Biomedical Science Academia Sinica, Taiwan	Post-doctoral fellow	2005	Human Molecular Genetics

**Specialty:** Molecular biology, Human molecular genetics, Cancer genomics

### Research Description

Our lab is primarily focusing on using Affymetrix SNP genotyping platform to investigate genomic variants in general population and psychological diseases, and chromosomal aberrations in patients with congenital malformation syndromes and cancers. We have constructed a copy number variation (CNV) map for the Han Chinese population, which is an important reference for identifying pathogenic or susceptible CNV in disease groups. We have identified *de novo* CNVs which could be responsible for the phenotypes in several patients with congenital malformation syndromes. In addition, we integrate chromosomal aberrations and differential expression of microRNA and protein-coding genes to identify cancer-related genes. Currently we are working on colorectal cancer. We have identified and validated several candidate tumor-related genes by integrative analysis. In the future, we will investigate the underlying mechanisms of these genes contributing to tumorigenesis. We will also evaluate whether the candidate genes can serve as biomarkers for diagnosis and/or disease monitoring.

**Research Grant:**

08/01/2007-07/31/2009 (PI) Construction of genome-wide copy number variation map in Han Chinese and its application in disease gene identification. (Grant #: NSC 96-2320-B-001-023-MY2, supported by National Science Council)

01/01/2009-12/31/2010 (PI) Identification and characterization of oncogenic miRNAs and genes involved in initiation and progression of colorectal cancer by genome-wide SNP scan. (A sub-project of an IBMS New Project Grant; supported by Institute of Biomedical Sciences, Academia Sinica)

08/01/2011-07/31/2014 (Co-PI) Study population genomics and human disease genomics using genome-wide single nucleotide polymorphism arrays, gene expression arrays and deep sequencing data. (Grant #: 100-2314-B-001-005-MY3, supported by National Science Council)

01/01/2016-12/31/2016 (PI) Identification of Genes and Oncogenic miRNAs Involved in Colorectal Cancer Susceptibility and Progression (I). (Grant #: VTA105-A-1-1, supported by Taichung Veteran Hospital and Academia Sinica)

01/01/2017-12/31/2017 (PI) Identification of Genes and Oncogenic miRNAs Involved in Colorectal Cancer Susceptibility and Progression (II). (Grant #: VTA106-A-1-1, supported by Taichung Veteran Hospital and Academia Sinica)

01/01/2018-12/31/2018 (co-PI) Developing a gut microbiota-based platform for guiding colorectal cancer screening and treatment-from animal study to pre-clinical trial (supported by Ministry of Science and Technology)

01/01/2019-12/31/2019 (co-PI) Developing a gut microbiota-based platform for guiding colorectal cancer screening and treatment-from animal study to pre-clinical trial (supported by Ministry of Science and Technology)

**National Core Grant:**

05/01/2012-04/30/2013 (Co-PI) National Center for Genome Medicine (II). (Grant #: NSC101-2319-B-001-001, supported by National Science Council)

05/01/2013-04/30/2014 (Co-PI) National Center for Genome Medicine (III). (Grant #: NSC102-2319-B-001-001, supported by National Science Council)

05/01/2014-04/30/2015 (Co-PI) National Center for Genome Medicine (IV). (Grant #: NSC103-2319-B-001-001, supported by National Science Council)

05/01/2015-04/30/2016 (Co-PI) National Center for Genome Medicine (I). (Grant #: MOST 104-2319-B-001-002, supported by Ministry of Science and Technology)

05/01/2016-04/30/2017 (Co-PI) National Center for Genome Medicine (II). (Grant #: MOST 105-2319-B-001-001, supported by Ministry of Science and Technology)

05/01/2017-04/30/2018 (Co-PI) National Center for Genome Medicine (II). (Grant #: MOST 106-2319-B-001-001, supported by Ministry of Science and Technology)

05/01/2018-04/30/2019 (Co-PI) National Center for Genome Medicine (II). (Grant #: MOST 107-2319-B-001-001, supported by Ministry of Science and Technology)

## Academic Service

- 2012-2018    Committee Member of Institutional Review Board at Academia Sinica  
2016-2018    Chief Executive Officer of Institutional Review Board at Academia Sinica

## Publications

1. Chen, C. H., Chen, H. I., Chien, W. H., Li, L. H., Wu, Y. Y., Chiu, Y. N., Tsai, W. C., Gau, S. S. (2017) High resolution analysis of rare copy number variants in patients with autism spectrum disorder from Taiwan. *Scientific Reports*, 7:11919.
2. Chuang, T.P., Wang, J.Y., Jao, S.W., Wu, C.C., Chen, J.H., Hsiao, K.H., Lin, C.Y., Chen, S.H., Su, S.Y., Chen, Y.J., Chen, Y.T., Wu, D.C.\*, Li, L.H.\* Over-expression of AURKA, SKA3 and DSN1 contributes to colorectal adenoma to carcinoma progression. *Oncotarget* 2016; 7(29):45803-45818. (Co-corresponding author)
3. Lin, Y.-F., Li, L.-H., Lin, C.-H., Tsou, M.-H., Chuang, M.-T.K., Wu, K.-M., Liao, T.-L., Li, J.-C., Wang, W.-J., Tomita, A., Tomita, B., Huang, S.-F., and Tsai, S.-F. (2016). Selective Retention of an Inactive Allele of the DKK2 Tumor Suppressor Gene in Hepatocellular Carcinoma. *PLoS Genetics* 12, e1006051
4. Yin, C.-L.<sup>#</sup>, Chen, H.-I.<sup>#</sup>, Li, L.-H.<sup>#</sup>, Chien, Y.-L., Liao, H.-M., Chou, M.C., Chou, W.-J., Tsai, W.-C., Chiu, Y.-N., Wu, Y.-Y., Lo, C.-Z., Wu, J.-Y., Chen, Y.-T., and Gau, S.S.-F. (2016). Genome-wide analysis of copy number variations identifies PARK2 as a candidate gene for autism spectrum disorder. *Molecular Autism* 7, 23. (Co-first authors)
5. Huang, M.-C., Chuang, T.-P., Chen, C.-H., Wu, J.-Y., Chen, Y.-T., Li, L.-H.\*, and Yang, H.-C\*. (2016). An integrated analysis tool for analyzing hybridization intensities and genotypes using new-generation population-optimized human arrays. *BMC Genomics* 17, 266. (Co-corresponding authors)
6. Lin, D.S., Chuang, T.P., Chiang, M.F., Ho, C.S., Hsiao, C.D., Huang, Y.W., Wu, T.Y., Wu, J.Y., Chen, Y.T., Chen, T.C., and Li, L.H.\* (2014). De novo MECP2 duplication derived from paternal germ line result in dysmorphism and developmental delay. *Gene* 533, 78-85.
7. Chiu, L.-Y., Kishnani, P.S., Chuang, T.-P., Tang, C.-Y., Liu, C.-Y., Bali, D., Koeberl, D., Austin, S., Boyette, K., Weinstein, D.A., Murphy, E., Yao, A., Chen, Y.-T., and Li, L.-H.\* (2014). Identification of differentially expressed microRNAs in human hepatocellular adenoma associated with type I glycogen storage disease: a potential utility as biomarkers. *Journal of gastroenterology* 49, 1274-1284.
8. Lin, Y.C., Lee, Y.C., Li, L.H., Cheng, C.J., and Yang, R.B. (2013). Tumor suppressor SCUBE2 inhibits breast-cancer cell migration and invasion through the reversal of epithelial-mesenchymal transition. *Journal of cell science*.
9. Yang, H.C., Lin, H.C., Kang, M., Chen, C.H., Lin, C.W., Li, L.H., Wu, J.Y., Chen, Y.T., and Pan, W.H. (2011). SAQC: SNP Array Quality Control. *BMC Bioinformatics* 12, 100.
10. Lin, C.H., Lin, J.K., Chang, S.C., Chang, Y.H., Chang, H.M., Liu, J.H., Li, L.H., Chen, Y.T., Tsai, S.F., and Chen, W.S. (2011). Molecular profile and copy number analysis of sporadic colorectal cancer in Taiwan. *Journal of biomedical science* 18, 36.
11. Yang, H.C., Lin, H.C., Huang, M.C., Li, L.H., Pan, W.H., Wu, J.Y., and Chen, Y.T. (2010). A new analysis tool for individual-level allele frequency for genomic studies. *BMC Genomics*

12. Wu, I.C., Wu, D.C., Huang, C.C., Lin, H.S., Chen, Y.K., Tsai, H.J., Lu, C.Y., Chou, S.H., Chou, Y.P., Li, L.H., Tai, S.Y., and Wu, M.T. (2010). Plasma decorin predicts the presence of esophageal squamous cell carcinoma. *Int J Cancer* 127, 2138-2146.
13. Shiao, Y.M., Lee, C.C., Hsu, Y.H., Huang, S.F., Lin, C.Y., Li, L.H., Fann, C.S., Tsai, C.Y., Tsai, S.F., and Chiu, H.C. (2010). Ectopic and high CXCL13 chemokine expression in myasthenia gravis with thymic lymphoid hyperplasia. *J Neuroimmunol* 221, 101-106.
14. Chien, S.C., Li, Y.C., Li, L.H., Wu, J.Y., Hsu, P.C., Shi, S.L., Tsai, F.J., and Lin, C.C. (2010). A new familial insertion, ins(18;9)(q12.2;q33.1q31.1) with a 9q31.1-9q33.1 deletion in a girl with a cleft lip and palate. *Am J Med Genet A* 152A, 1862-1867.
15. Chen, Y.K., Chang, W.S., Wu, I.C., Li, L.H., Yang, S.F., Chen, J.Y., Hsu, M.C., Chen, S.H., Wu, D.C., Lee, J.M., Huang, C.H., Goan, Y.G., Chou, S.H., Huang, C.T., and Wu, M.T. (2010). Molecular characterization of invasive subpopulations from an esophageal squamous cell carcinoma cell line. *Anticancer Res* 30, 727-736.
16. Chang, C.F., Li, L.H., Wang, C.H., Tsai, F.J., Chen, T.C., Wu, J.Y., Chen, Y.T., and Tsai, A.C. (2010). Identification of a submicroscopic 3.2 Mb chromosomal 16q12.2-13 deletion in a child with short stature, mild developmental delay, and craniofacial anomalies, by high-density oligonucleotide array-a recognizable syndrome. *Am J Med Genet A* 152A, 2365-2371.
17. Wu, K.M., Li, L.H., Yan, J.J., Tsao, N., Liao, T.L., Tsai, H.C., Fung, C.P., Chen, H.J., Liu, Y.M., Wang, J.T., Fang, C.T., Chang, S.C., Shu, H.Y., Liu, T.T., Chen, Y.T., Shiao, Y.R., Lauderdale, T.L., Su, I.J., Kirby, R., and Tsai, S.F. (2009). Genome sequencing and comparative analysis of *Klebsiella pneumoniae* NTUH-K2044, a strain causing liver abscess and meningitis. *J Bacteriol* 191, 4492-4501.
18. Kishnani, P.S., Chuang, T.P., Bali, D., Koeberl, D., Austin, S., Weinstein, D.A., Murphy, E., Chen, Y.T., Boyette, K., Liu, C.H., Chen, Y.T., and Li, L.H.\* (2009). Chromosomal and genetic alterations in human hepatocellular adenomas associated with type Ia glycogen storage disease. *Hum Mol Genet* 18, 4781-4790.
19. Yang, H.C., Huang, M.C., Li, L.H., Lin, C.H., Yu, A.L., Diccianni, M.B., Wu, J.Y., Chen, Y.T., and Fann, C.S. (2008). MPDA: microarray pooled DNA analyzer. *BMC Bioinformatics* 9, 196.
20. Lin, C.H., Li, L.H., Ho, S.F., Chuang, T.P., Wu, J.Y., Chen, Y.T., and Fann, C.S. (2008). A large-scale survey of genetic copy number variations among Han Chinese residing in Taiwan. *BMC Genet* 9, 92.
21. Lin, C.H., Huang, M.C., Li, L.H., Wu, J.Y., Chen, Y.T., and Fann, C.S. (2008). Genome-wide copy number analysis using copy number inferring tool (CNIT) and DNA pooling. *Hum Mutat* 29, 1055-1062.
22. Huggins, R., Li, L.H., Lin, Y.C., Yu, A.L., and Yang, H.C. (2008). Nonparametric estimation of LOH using Affymetrix SNP genotyping arrays for unpaired samples. *J Hum Genet* 53, 983-990.
23. Yang, H.C., Liang, Y.J., Huang, M.C., Li, L.H., Lin, C.H., Wu, J.Y., Chen, Y.T., and Fann, C.S. (2006). A genome-wide study of preferential amplification/hybridization in

microarray-based pooled DNA experiments. *Nucleic Acids Res* 34, e106.

24. Li, L.H., Ho, S.F., Chen, C.H., Wei, C.Y., Wong, W.C., Li, L.Y., Hung, S.I., Chung, W.H., Pan, W.H., Lee, M.T., Tsai, F.J., Chang, C.F., Wu, J.Y., and Chen, Y.T. (2006). Long contiguous stretches of homozygosity in the human genome. *Hum Mutat* 27, 1115-1121.
25. Chen, Y.T., Shu, H.Y., Li, L.H., Liao, T.L., Wu, K.M., Shiau, Y.R., Yan, J.J., Su, I.J., Tsai, S.F., and Lauderdale, T.L. (2006). Complete Nucleotide Sequence of pK245, a 98-Kilobase Plasmid Conferring Quinolone Resistance and Extended-Spectrum- $\beta$ -Lactamase Activity in a Clinical *Klebsiella pneumoniae* Isolate. *Antimicrob Agents Chemother* 50, 3861-3866.
26. Liu, Y.F., Chen, W.M., Lin, Y.F., Yang, R.C., Lin, M.W., Li, L.H., Chang, Y.H., Jou, Y.S., Lin, P.Y., Su, J.S., Huang, S.F., Hsiao, K.J., Fann, C.S., Hwang, H.W., Chen, Y.T., and Tsai, S.F. (2005). Type II collagen gene variants and inherited osteonecrosis of the femoral head. *N Engl J Med* 352, 2294-2301.
27. Chou, T.Y., Chiu, C.H., Li, L.H., Hsiao, C.Y., Tzen, C.Y., Chang, K.T., Chen, Y.M., Perng, R.P., Tsai, S.F., and Tsai, C.M. (2005). Mutation in the tyrosine kinase domain of epidermal growth factor receptor is a predictive and prognostic factor for gefitinib treatment in patients with non-small cell lung cancer. *Clin Cancer Res* 11, 3750-3757.
28. Li, L.H., Li, J.C., Lin, Y.F., Lin, C.Y., Chen, C.Y., and Tsai, S.F. (2004). Genomic shotgun array: a procedure linking large-scale DNA sequencing with regional transcript mapping. *Nucleic Acids Res* 32, e27.
29. Jong, Y.J., Li, L.H., Tsou, M.H., Chen, Y.J., Cheng, S.H., Wang-Wuu, S., Tsai, S.F., Chen, C.M., Huang, A.T., Hsu, M.T., and Lin, C.H. (2004). Chromosomal comparative genomic hybridization abnormalities in early- and late-onset human breast cancers: correlation with disease progression and TP53 mutations. *Cancer Genet Cytogenet* 148, 55-65.
30. Huang, S.F., Liu, H.P., Li, L.H., Ku, Y.C., Fu, Y.N., Tsai, H.Y., Chen, Y.T., Lin, Y.F., Chang, W.C., Kuo, H.P., Wu, Y.C., Chen, Y.R., and Tsai, S.F. (2004). High frequency of epidermal growth factor receptor mutations with complex patterns in non-small cell lung cancers related to gefitinib responsiveness in Taiwan. *Clin Cancer Res* 10, 8195-8203.
31. Chang, Y.T., Shiao, Y.M., Chin, P.J., Liu, Y.L., Chou, F.C., Wu, S., Lin, Y.F., Li, L.H., Lin, M.W., Liu, H.N., and Tsai, S.F. (2004). Genetic polymorphisms of the HCR gene and a genomic segment in close proximity to HLA-C are associated with patients with psoriasis in Taiwan. *Br J Dermatol* 150, 1104-1111.
32. Vander Zwan, C.J., Wheeler, J.C., Li, L.H., Tracey, W.D., and Gergen, J.P. (2003). A DNA-binding-independent pathway of repression by the *Drosophila* Runt protein. *Blood Cells Mol Dis* 30, 207-222.
33. Li, L.H., and Gergen, J.P. (1999). Differential interactions between Brother proteins and Runt domain proteins in the *Drosophila* embryo and eye. *Development* 126, 3313-3322.
34. Aimes, R.T., Li, L.H., Weaver, B., Hawkes, S., Hahn-Dantona, E.A., and Quigley, J.P. (1998). Cloning, expression, and characterization of chicken tissue inhibitor of metalloproteinase-2 (TIMP-2) in normal and transformed chicken embryo fibroblasts. *J Cell Physiol* 174, 342-352.
35. Golling, G., Li, L., Pepling, M., Stebbins, M., and Gergen, J.P. (1996). *Drosophila* homologs of

the proto-oncogene product PEBP2/CBF beta regulate the DNA-binding properties of Runt.  
Mol Cell Biol 16, 932-942.