# Yuanyuan Li

Staff Scientist, Biostatistics and Computational Biology Branch National Institute of Environmental Health Sciences / National Institutes of Health (NIEHS/NIH), Durham, NC Email: yuanyuan.li@nih.gov

URL: https://yuanyuanli66.github.io/

## Education:

2010	Ph.D., Electrical Engineering and Computer Science, University of Tennessee, Knoxville
2006	M.S., Electrical Engineering and Computer Science, UTK
2001	B.S., Computer Information Science, Minnesota State University, Mankato

## Awards and Honors:

2017, 2014	NIH Fellows Award for Research Excellence (FARE)
2013	NIEHS Science Day Best Poster Presentation Award
2011-2014	NIH Intramural Research Training Award (IRTA) Postdoctoral Fellowship
2008	Upsilon Pi Epsilon (UPE) International Honor Society for the Computing Sciences
2007, 2004	Scholarly and Research Incentive Funds (SARIF) Scholarship, University of Tennessee, Knoxville
2006	Best Graduate Teaching Assistant Award, University of Tennessee, Knoxville
2001	Graduate with honors (cum laude), Minnesota State University, Mankato
1999-2001	International Student Endowment Fund Scholarship, Minnesota State University, Mankato
1998-2001	Dean's List, Minnesota State University, Mankato

## **Research Experience:**

Staff Scientist, Biostatistics & Computational Biology Branch, NIH/NIEHS 10/2018 - present

Method development and mining pan-cancer genomic data using machine learning (ML)

- Use bagging and boosting to identify biomarkers that associates with various clinical outcomes from The Cancer Genome Atlas (TCGA) RNA-seq and clinical data
- Develop in-house stochastic gradient boosted machine (GBM) packages using R and Java
- Extend logistic regression trees to handle pair-matched (case-control) data
- Develop RNA-seq pipelines to detect differential expressed genes from raw signals
- Key ML techniques include: GBMs with various loss functions (logloss, mean-squared-error and learningto-rank), rank aggregation, permutation test, T-test and Wilcoxon rank-sum test with False Discovery Rate (FDR) correction, Kaplan–Meier estimator and Cox proportional-hazards model with left truncation

Collaboration: mining various -omics, diet, clinical chemistry/hormone data, and histopathology findings

- Use various supervised and unsupervised ML techniques to recognize patterns and identify features associated with outcomes of collaborators' interests
- Collaborators include: within BCBB, epidemiology branch, clinical research unit, signal transduction branch, genome integrity and structural biology, National Toxicology Program (NTP) and Centers for Disease Control and Prevention (CDC)
- Develop various supervised and unsupervised ML pipelines to detect biological signals
- Incorporate nested sampling in source data as weights into tree learning
- Apply developed T-KDE toolbox on ChIP-seq data (see below)
- Key ML techniques include: Classification and regression trees (CART), GBMs, boosted logistic regression tree stumps, cost-sensitive learning, various clustering analysis, topic models, Kernel Density Estimation (KDE), Principal Component Analysis (PCA), T-distributed Stochastic Neighbor Embedding (t-SNE), survival analysis, and various statistical hypothesis tests

#### Research Fellow, Biostatistics & Computational Biology Branch, NIH/NIEHS

05/2014 - 10/2018

Pan-cancer classification using in-house Genetic Algorithm / K-nearest neighbors (GA/KNN)

- Developed a parallel version of GA/KNN using POSIX Threads (p-threads)
- Proposed to use Latent Semantic Indexing (LSI) to speedup searches in GA
- Proposed to use GBMs improve GA/KNN's performances
- Associated identified driver genes to their corresponding tumor types by various cluster analysis

### IRTA Postdoctoral Fellow, Biostatistics Branch, NIH/NIEHS

Method for analyzing genome-wide protein binding patterns from ChIP-seq data

- Proposed and implemented T-KDE toolbox, to identify the locations of constitutive protein binding sites using ChIP-seg data
- T-KDE, which combines a binary range tree with a kernel density estimator to quickly identify constitutive protein binding sites from multiple cell lines.
- T-KDE can also identify genomic "hot spots" where several different proteins bind and, conversely, cellspecific sites bound by a given protein

Identify functional relevance of CCCTC-binding factor (CTCF) protein

Analyzed CTCF's genomic distributions, transcriptional environment, and epigenomic environment

#### Postdoctoral Researcher, Biomedical Engineering, UTK

Machine learning-based approach for immune system and drug design

 Developed ML approach to model immune system and drug interaction using Fuzzy-clustering combined with variable length Markov model implemented in forms of Probabilistic Suffix Tree (PST)

Immune-inspired computational model

· Developed immune-inspired game theory for irregular warfare

Plant-based sensor network for nanoparticles toxicity study

- Developed and submitted a NSF proposal
- Developed a plant-based sensor-network for characterizing, monitoring, and understanding the environmental impact of both naturally occurring and man-made nanoparticles

Graduate Research Assistant, Distributed Intelligence Laboratory, **EECS**, UTK 05/2004 - 08/2010

Anomaly detection in unknown environments using wireless sensor networks (WSNs) and a mobile robot (Partly funded by Oak Ridge National Laboratory)

- Designed and implemented a variety of distributed machine learning algorithms on a hierarchical resourceconstrained sensor network (MICA2 and MICA2dots)
- Video demonstration: (Physical robots) Sensor network detects abnormal situation, with mobile robot (Pioneer 3) responding to location of anomaly (2007)
- Proposed a novel multiple missing data imputation technique that uses KD-tree with Mahalanobis distance for WSNs
- Key ML techniques include: PSTs, KD-trees, Fuzzy Adaptive Resonance Theory (Fuzzy-ART) neural network, Lempel-Ziv-Welch (LZW) algorithm for compression, likelihood-ratio test, autoregressive model and  $R^2$

Indoor wireless localization for mobile robots

 Designed and implemented wireless indoor positioning system to locate mobile robots using triangulation and fingerprinting

### Graduate Research Assistant, Computer Science, MNSU

Bluetooth network simulator (Individual study research project)

 Developed Bluetooth network simulation software that simulated the behavior of a Bluetooth PicoNet with 1 to 7 slaves by using JavaSimulation package (a Java package for process-based discrete event simulation)

Text-to-Speech Synthesis for Mandarin Chinese

Researched text-to-speech synthesis for Mandarin Chinese

## **Professional Experience:**

### Midwest Wireless Corporation (now Alltel Corporation), Mankato, MN

Software engineer co-op

 Developed a framework monitoring the SMSC (short messaging) server including short messages from phone-to-phone (NOKIA7160), phone-to-PC and PC-to-phone; and service messages push to the phone

05/2011 - 05/2014

2

# 01/2002 - 08/2003

05/2003 - 08/2003

09/2010 - 05/2011

## Yuanyuan Li

SpeechGear Inc (U.S. Naval Research funded project), Northfield, MN	07/2002 - 05/2003
Software engineer co-op	an DDAa (Windowa CC)
<ul> <li>Designed and developed multiple interfaces for voice-enabled dictionary running using Java and eMbedded Visual Basic/C++</li> </ul>	on PDAs (Windows CE)
DataPlanIT Consulting, Mankato, MN	01/2002 - 05/2002
Software engineer (part-time)	
<ul> <li>Web design and development for surrounding businesses in Mankato using Activ</li> </ul>	ve Server Pages
J.D. Edwards Company (now Oracle Corporation), Denver, CO	06/2001 - 08/2001
Software engineer intern	
<ul> <li>Wrote testing cases and suites for the MetaData software using JUnite</li> </ul>	
Hairs Supply Company, Chicago, IL	09/2000 - 12/2000
Software engineer part-time	
<ul> <li>Designed and developed an e-commerce website that sells hair supplies using A</li> </ul>	ctive Server Pages
Visible Edge Company, Mankato, MN	06/2000 - 08/2000
Software engineer intern	
<ul> <li>Upgraded and debugged the Performance Look Up System (PLUS) for Minnes Visual Basic</li> </ul>	ota high schools using
Teaching Experience:	
Instructor, Winter/Summer Biostatistics and Bioinformatics Short Courses, NIEHS	12/2017 - present
DNA microarray data analysis	
<ul> <li>Overview of microarray technology, experimental design, data preprocessing, sta ing, clustering, and classification</li> </ul>	atistical hypothesis test-
Teaching Assistant, Electrical Engineering and Computer Science, UTK	08/2003 - 09/2010
CS100: Introduction to Computer Science (for non-majors)	
<ul> <li>Programming: HTML, JavaScript and basic algorithms</li> </ul>	
CS102: Introduction to Computer Science (for majors)	
Programming: C++	
CS302: Fundamental Algorithms	
<ul><li>CS365: Programming Languages and Systems</li><li>Programming: Java, Python and Perl</li></ul>	
CS530: Computer Systems Organization	
CS594: Data Mining Practices and Principles	
Research Facilitator, The Oak Ridge Associated Universities (ORAU), TN	07/2008 - 08/2008
<ul> <li>Mentored a team of high school students on how to solve challenging navigation Robotics Kit; sponsored by Appalachian Regional Commission and Oak Ridge Asso</li> </ul>	
Teaching Assistant, Computer Science, MNSU	08/2001 - 2002
<ul><li>CS100: Introduction to Computer and Computing</li><li>Programming: Microsoft Office 2000, HTML and JavaScript</li></ul>	
Publications:	

#### **Refereed journal papers**

1. Y. Li, M. Li, I. Shats, J. M. Krahn, G. P. Flake, D. M. Umbach, X. Li, and L. Li, "Glypican 6 is a putative biomarker for metastatic progression of cutaneous melanoma", *PLoS One*, to appear, 2019

3

- T.T. Nguyen, S. A. Grimm, P. R. Bushel, J. Li, Y. Li, B.D. Bennett, D. C. Fargo, C. W. Anderson, L. Li, M. A. Resnick, and D. Menendez, "Revealing the human p53 universe", *Nucleic acids research*, 46 (16), 8153-8167, 2018
- 3. Y. Li, D. M. Umbach, and L. Li, "Putative genomic characteristics of BRAF V600K versus V600E cutaneous melanoma", *Melanoma Research*, 27 (6), 527-535, 2017
- Y. Li, J. M. Krahn, N. Croutwater, K. Lee, D. M. Umbach, and L. Li, "A comprehensive genomic pan-cancer analysis using The Cancer Genome Atlas gene expression data", *BMC genomics*, 18 (1), 508, 2017 (Cited by 19, source: Google)
- 5. Y. Li, J. M. Krahn, G. P. Flake, D. M. Umbach and L. Li, "Toward predicting metastatic progression of melanoma based on gene expression data", *Pigment Cell & Melanoma Research*, 28 (4), 453-463, 2015
- 6. Y. Li, D. M. Umbach, and L. Li, "T-KDE: A method for analyzing genome-wide protein binding patterns from ChIP-seq data sets", *BMC Genomics*, 15 (1), 27, 2014
- Y. Li, M. Thomason, and L. E. Parker, "Sequential anomaly detection using wireless sensor networks in unknown environments", *Human behavior understanding in networked sensing - Theory and Applications* of Networks of Sensors, 99-123, 2014
- 8. Y. Li, and L. E. Parker, "Nearest neighbor imputation using spatial-temporal correlations in wireless sensor networks", *Information fusion*, 15, 64-79, 2014 (Cited by 44, source: Google)
- Y. Li, W. Huang, L. Niu, S. Covo, D. M. Umbach, and L. Li, "Characterization of constitutive CTCF/Cohesin loci: a possible role in establishing topological domains in mammalian genomes", *BMC Genomics*, 14 (1), 553, 2013 (Cited by 61, source: Google)
- S. Lenaghan, Y. Li (co-first authors), H. Zhang, J. Burris, C. Stewart, L. E. Parker, and M. Zhang, "Monitoring the environmental impact of TiO2 nanoparticles using a Plant-based sensor-network", *IEEE Transactions on Nanotechnology*, 2 (2), 182-189, 2013
- 11. Y. Li, S. Lenaghan, and M. Zhang, "A data-driven predictive approach for drug delivery using machine learning techniques", *PLoS one*, 7(2): e31724, 2012

### **Refereed conference papers**

- 1. Y. Li, S. Lenaghan, J. Burris, C. N. Stewart, L. E. Parker, and M. Zhang, "Detecting the environmental impact of nanoparticles using plant-based biosensors", *The* 11<sup>th</sup> *IEEE Conference on Nanotechnology* (*IEEE-NANO*), pages 48-52, doi:10.1109/NANO.2011.6144505, August, 2011
- Y. Li, M. Thomason, and L. E. Parker, "Detecting time-related changes in wireless sensor networks using symbol compression and probabilistic suffix trees", *IEEE/RSJ International Conference on Intelligent Robots and Systems (IROS)*, pages 2946-2951, doi:10.1109/IROS.2010.5649660, October, 2010
- Y. Li, and L. E. Parker, "Detecting and monitoring time-related abnormal events using a wireless sensor network and mobile robot", *IROS*, pages 3292-3298, doi:10.1109/IROS.2008.4651031, October, 2008 (Cited by 30, source: Google)
- 4. Y. Li, and L. E. Parker, "A spatial-temporal imputation technique for classification with missing data in a wireless sensor network", *IROS*, pages 3272-3279, doi:10.1109/IROS.2008.4650774, October, 2008 (Cited by 37, source: Google)
- 5. Y. Li, and L. E. Parker, "Intruder detection using a wireless sensor network with an intelligent mobile robot response", *IEEE Southeastcon*, pages 37-42, doi:10.1109/SECON.2008.4494250, April, 2008, (Cited by 56, source: Google)
- 6. Y. Li, and L. E. Parker, "Classification with missing data in a wireless sensor network", *IEEE Southeastcon*, pages 533-538, doi:10.1109/SECON.2008.4494352, April, 2008 (Cited by 23, source: Google)
- 7. Y. Li, and S. Case, "Text-to-Speech synthesis for Mandarin Chinese", In Proceedings of the Midwest Instruction and Computing Symposium (MICS), April, 2003

#### Papers under review:

- 1. Y. Li, A. Bingham, D. M. Umbach, and L. Li, "Putative biomarkers for predicting tumor sample purity based on gene expression data", *Scientific Reports*, 2019
- 2. K. Kang, Q. Meng, I. Shats, D. M. Umbach, B. Papas, Y. Li, X. Li and L. Li, "A novel complete computational deconvolution method using RNA-seq data", *Bioinformatics*, 2019

C. M. Clinton, J. R. Bain, M. J. Muehlbauer, Y. Li, L. Li, S. K. O'Neal, B. L. Hughes, D. E. Cantonwine, T. F. McElrath, K. K. Ferguson, "Non-targeted urinary metabolomics in pregnancy and associations with fetal growth restriction", 2019

## Papers in preparation:

- 1. Y. Li, D. M. Umbach, and L. Li "Classification of breast cancer and sub-classification of triple-negative breast cancer samples based on TCGA gene and protein expression data"
- 2. Y. Li, and L. Li "Gene expression profiles to predict melanoma's target distant organs"
- 3. M. Shi, L. Li, A. Wise, D.M. Umbach, J. Krahn, Y. Li, C. R. Weinberg, "GA-KNN algorithm for detecting epistasis effects in case-parents triads"

## **Dissertation:**

1. "Anomaly detection in unknown environments using wireless sensor networks", Distributed Intelligence Laboratory, EECS, UTK, May 2010

## **Technical Reports:**

1. Y. Li, "Indoor positioning using 802.11b for mobile robots", DiLab, EECS, UTK, December 2005

## Posters:

- 1. Y. Li, D. M. Umbach, and L. Li, "Putative biomarkers for tumor sample purity prediction based on gene expression data" *American Association for Cancer Research (AACR) Annual Meeting*, March, 2019
- C. M. Clinton, J. R. Bain, M. J. Muehlbauer, Y. Li, L. Li, S. K. O'Neal, B. L. Hughes, D. E. Cantonwine, T. F. McElrath, and K. K. Ferguson, "Urinary metabolimic profiles in pregnancy and feta growth restriction", *The Pregnancy Meeting*, February, 2019
- K. Kang, Q. Meng, I. Shats, D. M. Umbach, M. Li, Y. Li, X. Li, and L. Li, "A novel computational complete deconvolution method using RNA-seq data", *The* 17<sup>th</sup> *European Conference on Computational Biology* (*ECCB*), September, 2018
- 4. Y. Li, A. Bingham, Q. Li, Y. Zhuang, D. M. Umbach and L. Li, "Using tumor sample gene expression data to infer tumor purity levels with stochastic gradient boosting machines", AACR Annual Meeting, March, 2018
- 5. Q. Xu, I. Shats, Y. Li, L. Li, and X. Li, "1HNF4A-mediated methionine metabolism confers sensitivity of human hepatocellular carcinoma to methionine restriction", *DIR Board of Scientific Counselors (BSC) Review*, NIEHS, July, 2018
- 6. Y. Li, A. Bingham, D. M. Umbach, and L. Li, "Using tumor sample gene expression to learn about tumor purity and the tumor microenvironment" *NIEHS Science Day*, 2017
- A. Bingham, Y. Li, D. M. Umbach, and L. Li, "Using tumor sample gene expression data to learn about tumor purity levels and the tumor microenvironment" *NIEHS summer intern poster competition*, Best Poster Presentation Award, June, 2017
- 8. Y. Li, J. Krahn, N. Croutwater, K. Lee, D. M. Umbach, and L. Li, "A comprehensive genomic pan-cancer analysis using The Cancer Genome Atlas gene expression data" *DIR BSC Review*, NIEHS, November, 2016
- 9. D. M. Umbach, M. Shi, A. Wise, J. Krahn, Y. Li, C. R. Weinberg, and L. Li, "A stochastic search algorithm for finding multi-SNP effects using nuclear families", *Joint Statistical Meeting (JSM)*, July, 2016
- 10. N. Croutwater, L. Li, and, Y. Li, "A comprehensive genomic pan-cancer classification using The Cancer Genome Atlas gene expression data", *NIEHS summer intern poster competition*, June, 2016
- 11. Y. Li, D. M. Umbach, L. Li, "A comprehensive genomic pan-cancer analysis comparing males and females using The Cancer Genome Atlas gene expression data" *AACR Precision Medicine Series: Targeting the Vulnerabilities of Cancer Conference*, May, 2016
- 12. Y. Li, J. M. Krahn, G. P. Flake, D. M. Umbach, and L. Li, "Glypican 6 is a putative biomarker for metastatic progression of cutaneous melanoma", *NIEHS Science Day*, 2015
- C. R. Weinberg, M. Shi, A. Wise, D. M. Umbach, J. Krahn, Y. Li, and L. Li, "A stochastic search algorithm for finding multi-SNP effects using nuclear families", *International Genetic Epidemiology Society Conference* (*IGES*), October, 2015

- 14. **Y. Li**, J. M. Krahn, and L. Li, "Putative biomarkers indicative of metastatic progression of skin cutaneous melanoma", *AACR Melanoma: From Biology to Target Conference*, 2014
- 15. A. Mateja, **Y. Li**, and L. Li, "Using T-KDE to discover novel loci that may be implicated in X-inactivation", *NIEHS summer intern poster competition*, June, 2014
- 16. Y. Li, D. M. Umbach, and L. Li, "T-KDE: A method for analyzing genome-wide protein binding patterns from ChIP-seq data", *NIEHS Science Day*, **Best Poster Presentation Award**, 2013
- 17. Y. Li, D. M. Umbach, and L. Li, "Analysis of genome-wide protein binding patterns using kernel density estimators", *the Biology of Genomes Conference*, May, 2013
- 18. Y. Li, W. Huang, D. M. Umbach, S. Covo, and L. Li, "Constitutive CTCF/Cohesin loci in a transcriptionally complex environment", *NIEHS Science Day*, 2012
- 19. Y. Li, J. Wu, S. C. Lenaghan, and M. Zhang, "An Immuno-Inspired Game Theoretic Computational Framework for Irregular Warfare", *Naval Science & Technology Partnership Conference*, 2010

## **Conference and Research Presentations:**

- "Learning about tumor microenvironment using tumor sample gene expression and purity data" *North Carolina Biotechnology Seminar Series (invited talk)*, RTP, NC, 2019
- "Tree learing for big omics data", NIEHS seminar, 2018
- "Learning with eXtreme Gradient Boosting a gradient boosting approach", BCBB seminar, NIEHS, 2017
- "A comprehensive genomic pan-cancer classification using The Cancer Genome Atlas gene expression data", BCBB retreat, NIEHS, 2016
- "Identifying constitutive binding sites using kernel approach", BCBB retreat, NIEHS, 2012
- "Detecting the environmental impact of nanoparticles using plant-based biosensors", *IEEE-NANO conference*, August, 2011
- "Study complex biological systems: network modeling and AI-based analysis", Guest lecture: Systems Biology and Complex System Theory, BME, UTK, October, 2010
- "Detecting time-related changes in wireless sensor networks using symbol compression and probabilistic suffix trees", *IROS conference*, October, 2010
- "Environment monitoring using Wireless Sensor Networks", *Guest lecture: Artificial Intelligence*, EECS, UTK, November, 2009
- "Detecting and monitoring time-related abnormal events using a wireless sensor network and mobile robot", *IROS conference*, October, 2008
- "A spatial-temporal imputation technique for classification with missing data in a wireless sensor network", *IROS conference*, October, 2008
- "Intruder detection using a wireless sensor network with an intelligent mobile robot response", the IEEE Southeast conference, April, 2008
- "Classification with missing data in a wireless sensor network", the IEEE Southeast conference, April, 2008, Alabama, Huntsville, USA
- "Exploring the impact of mobility in wireless sensor network", Oak Ridge National Laboratory, 2006
- "Text-to-Speech synthesis for Mandarin Chinese", MICS conference, April, 2003

# **Technical Skills:**

- **Programming languages:** Java, Javascript, C/C++, NesC, eMbeddedVB/C++, shell scripts, Visual Basic, JSP, ASP, SQL, PL/SQL, XML, and WML
- Applications: Player/Stage, Matlab, R, WEKA, Oracle, MySQL, Orion server, LATEX, BIBTEX, Microsoft Office, and other common productivity packages for Windows and Linux platforms
- Operating Systems: Microsoft Windows, Unix/Linux, Mac OS, and TinyOS

## Services:

## Paper reviewing:

- NIH internal paper review, 2012 present
- PLoS One Journal, 2011 present
- Program committee of Association for the Advancement of Artificial Intelligence (AAAI), 2018
- Program committee of International Joint Conference on Artificial Intelligence (IJCAI), 2016 2018

#### Yuanyuan Li

- Scientific Reports Nature, 2017 2018
- International Journal of Wireless Information Networks (IJWI), 2013 2017
- Bioinformatics Journal, 2017
- Soft Computing Journal, 2017
- NIH FARES award, 2017
- IEEE Sensors Journal, 2014 2017
- Neurocomputing Journal, 2016
- IEEE Systems Journal, 2014 2016
- International Journal of Computer Systems Science and Engineering (IJCSSE), 2013
- Sensors (ISSN 1424-8220), 2010
- IEEE/RSJ Intl. Conf. on Intelligent Robots and Systems (IROS), 2006 2010, 2018
- IEEE International Conference on Robotics and Automation (ICRA), 2006 2007
- IEEE Intelligent Systems, 2006
- Journals of Robotics and Autonomous Systems (JRAS), 2005

#### Mentoring:

- Special volunteer within the group, BCBB, NIEHS, 2016 present
- Summer undergraduate and graduate students within the group, BCBB, NIEHS, 2014 present
  - Summer student own the 2017 best graduate student presentation award

#### **Robotics activities:**

- Coach for the First LEGO League (FLL) robotics competition, 2015 2016, 2018 present
  - Two groups own robot design awards (2015 and 2019)
- Software judge for the FLL robotics competition in Tennessee and North Carolina, 2004 2014
- Tour guide for the Tennessee Junior Science & Humanities Symposium, 2010
- Robotic demonstration for the National Science Foundation (NSF) campus tour, UTK, 2008
- Robotic demonstration for local middle school and high school students, UTK, 2005 2010
- Maintained operating systems, software and hardware for Pionneer autonomous robots and Crossbow sensors for the Distributed Intelligence Laboratory, UTK, 2005 2010

#### Others:

- Biostatistics branch liaison for the NIEHS Trainees Assembly (NTA) steering committee, 2012 2014
- Judge for Science Fair for Triad Math and Science Academy, North Carolina, 2014
- Volunteer for the United Way fund allocation committee, Tennessee, 2005 to 2008
- President of the Chinese Student and Scholar Association, Minnesota State University, 2001
- Director of Activities of the International Student Association, Minnesota State University, 2000
- Peer Leader for the Intentional Student Orientation, Minnesota State University, 2000
- Vice President of the Taiwan Student Association, Minnesota State University, 1999