# RAUNAK SHRESTHA [Ph.D. Candidate (Bioinformatics), University of British Columbia]

Curriculum Vitae [Last Updated: June 5, 2018]

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#### RESEARCH STATEMENT

My primary research goals are directed towards understanding the complex nature of cancer by computationally modelling and integrating multi-omics cancer profiles so as to enable precision oncology. I develop computational algorithms and apply them to analyze big biological data to identify appropriate treatment for individual cancer patient.

Recent advances in high throughput technologies have enabled us to assess multi-dimensional view of cancer process. I believe, to integrate these "big data", the computational model must be complex enough to be accurate while being simple enough to portray the underlying biology. For this, network approaches helps to bring these multi-dimensional data into a single-analysis framework.

My current research focuses on exploring this idea analyzing the genotype-phenotype relationships in cancer to prioritize cancer driver genes and precision therapeutics. Most recently, my research led to define a novel molecular subtype as well as biomarker stratifying this subtype of a rare cancer of abdomen vulnerable to immunotherapy.

Furthermore, I would like to understand the spatial heterogeneity in cancer. I believe, communication between tumor and its micro-environment and vice versa is critical for cancer progression and therapeutic resistance. Using integrated single-cell multi-omics analysis, I would like to study how tumor-microenvironment communication shapes cancer evolution and how can we leverage this knowledge to develop novel cancer therapeutics.

#### POST-SECONDARY EDUCATION

SEPT 2012 - AUG 2018	Doctor of Philosophy in BIOINFORMATICS, University of British Columbia, Canada under Bioinformatics Training Program for Health Research Advisors: Dr. Colin C. Collins and Dr. S. Cenk Sahinalp Thesis Title: "Computational Prioritization of Cancer Driver Genes for Precision Oncology"
SEPT 2011 - AUG 2012	Master of Science in BIOINFORMATICS, <b>Simon Fraser University</b> , Canada under Bioinformatics Training Program for Health Research (Transferred to PhD program)
2005 - 2009	Bachelors of Technology (B.Tech.) in BIOTECHNOLOGY, Kathmandu University, Nepal

#### RESEARCH & TRAINING EXPERIENCE

SEPT 2012 - AUG 2018	PhD Candidate, Laboratory for Advanced Genome Analysis, Vancouver Prostate Centre, Vancouver, BC, Canada
May - Aug 2012	Graduate Research Student, Laboratory for Advanced Genome Analysis, Vancouver Prostate Centre, Vancouver, BC, Canada
SEPT 2011 - MAY 2012	Graduate Research Student, Simon Fraser University, Burnaby, BC, Canada
JULY 2009 - JULY 2011	Research Associate, Center for Molecular Dynamics Nepal, Kathmandu, Nepal
Jan - June 2009	Undergrad Research Student, Genetics Nepal, Kathmandu, Nepal

## SELECTED PUBLICATIONS

- Shrestha R, Nabavi N, Lin Y, Mo F, Anderson S, Volik S, Adomat HH, Lin D, Xue H, Dong X, Shukin R, Bell RH, McConeghy B, Haegert A, Brahmbhatt S, Li E, Oo HZ, Hurtado-Coll A, Fazli L, Zhou J, McConnell Y, McCart A, Lowy A, Morin GB, Daugaard M, Sahinalp SC, Hach F, LeBihan S, Gleave ME, Wang Y, Churg A, and CC Collins. (2018) Integrated Multi-omics Molecular Subtyping Predicts Therapeutic Vulnerability in Malignant Peritoneal Mesothelioma. (In submission) (Preprint bioRxiv doi:10.1101/24347).
- Hodzic E, Shrestha R<sup>†</sup>, Zhu K, Cheng K, Collins CC, and SC Sahinalp. (2018) **Combinatorial Detection of Conserved Alteration Patterns for Identifying Cancer Subnetworks**. (In submission) <sup>†</sup>As joint first author
- Shrestha R, Hodzic E, Sauerwald T, Dao P, Yeung J, Wang K, Anderson S, Haffari G, Collins CC, and Sahinalp SC. (2017) HIT'nDRIVE: Patient-Specific Multi-Driver Gene Prioritization for Precision Oncology. Genome Research. 27:1573-1588. doi:10.1101/gr.221218.117. PMID:28768687
- Shrestha R, Hodzic E, Yeung J, Wang K, Sauerwald T, Dao P, Anderson S, Beltran H, Rubin M., Collins C, Haffari G and SC Sahinalp. (2014) HIT'nDRIVE: Multi-Driver Gene Prioritization based on Hitting Time. Research in Computational Molecular Biology: 18<sup>th</sup> Annual International Conference, RECOMB 2014, Proceedings, pages 293–306. doi:10.1007/978-3-319-05269-4\_23.

## RELEVANT PUBLICATIONS

- Feng Y, Ren S, Shrestha R, Bell R, Volik S, Cox M, Sharifi-Noghabi H, Zhou J, Jun W, Li Y, Jian W, Wang Y, Sun Y, and CC Collins. (2018) Metagenomic and metatranscriptomic analysis of human prostate microbiota from patients with prostate cancer. (in submission)
- Sharifi-Noghabi H, Liu Y, Erho N, **Shrestha R**, Alshalalfa M, Davicioni E, Collins CC, and M Ester. (2018) **Deep Genomic Signature for early metastasis prediction in Prostate Cancer**. (in submission) (Preprint bioRxiv doi:10.1101/276055)
- Gill E, Chan L, Winsor G, Dobson N, Lo R, Sui SH, Dhillon B, Taylor P, Shrestha R, Spencer C, Hancock R, Unrau P, and Brinkman F. (2018) High-throughput detection of RNA processing in bacteria. BMC Genomics. 19:223. PMID:29587634
- Yamada M, Tang J, Lugo-Martinez J, Hodzic E, Shrestha R, Saha A, Ouyang H, Yin D, Mamitsuka H, Sahinalp C, Radivojac P, Menczer F, and Chang Y. (2018) Ultra High-Dimensional Nonlinear Feature Selection for Big Biological Data. IEEE Transactions on Knowledge and Data Engineering. 99. doi:10.1109/TKDE.2018.2789451
- Luk I, Shrestha R, Xue H, Wang Y, Zhang F, Lin D, Haegert A, Wu R, Dong X, Collins CC, Zoubeidi A, Gleave ME, Gout PW, and Wang Y. (2016) BIRC6-targeting as potential therapy for advanced, enzalutamide-resistant prostate cancer. Clinical Cancer Research. 2016 Sep 23. doi:10.1158/1078-0432.CCR-16-0718. PMID:27663589
- Wyatt A, Mo F, Wang K, McConeghy B, Brahmbhatt S, Jong L, Mitchell D, Johnston R, Haegert A, Li E, Liew J, Yeung J, Shrestha R, Lapuk A, McPherson A, Shukin R, Bell R, Anderson S, Bishop J, Hurtado-Coll A, Xiao H, Chinnaiyan A, Mehra R, Lin D, Wang Y, Fazli L, Gleave M, Volik S, and Collins CC. (2014) Heterogeneity in the inter-tumor transcriptome of high risk prostate cancer.

  Genome Biology. 15(8):426 doi:10.1186/s13059-014-0426-y. PMID:25155515
- Wang K, Shrestha R, Wyatt AW, Reddy A, Lehar J, Wang Y, Lapuk A and CC Collins. (2014) A Meta-Analysis Approach for Characterizing Pan-Cancer Mechanisms of Drug Sensitivity in Cell Lines. PLoS ONE 9(7). doi:10.1371/journal.pone.0103050. PMID:25036042
- Shrestha R, Joshi R., Joshi K, Poudel BH and BG Shrestha. (2011) Analysis of KatG Ser315Thr Mutation in Multidrug Resistant Mycobacterium tuberculosis and SLC11A1 Polymorphism in Multidrug Resis-

tance Tuberculosis in Central Development Region of Nepal Using PCR-RFLP Technique: A Pilot Study. Nepal Journal of Biotechnology 1 (1), 14-21.

#### THESIS

Shrestha R. Computational Prioritization of Cancer Driver Genes for Precision Oncology.
 Ph.D. Thesis. University of British Columbia. August 2018.
 Advisors: Dr. Colin C. Collins & Dr. S. Cenk Sahinalp

## **CONFERENCE PRESENTATIONS**

#### **Selected Oral Presentations**

- BAP1 Loss Predicts Therapeutic Vulnerability in Malignant Peritoneal Mesothelioma. 14<sup>th</sup> International Conference of the International Mesothelioma Interest Group (iMig2018). May 2-5, 2018, Ottawa, Canada
- HIT'nDRIVE: Patient-Specific Multi-Driver Gene Prioritization for Precision Oncology. 8<sup>th</sup> Annual Scientific Meeting, Terry Fox Research Institute. November 4, 2017, Vancouver, Canada
- Translating Big-Data to Precision Oncology. Terry Fox Seminar, Vancouver Prostate Centre. February 17, 2017, Vancouver General Hospital, Vancouver, Canada.
- HIT'nDRIVE: Patient-Specific Multi-Driver Gene Prioritization for Precision Oncology. Vancouver Bioinformatics User Group (VanBUG). November 3, 2016, Vancouver, Canada
- HIT'nDRIVE: Patient-Specific Multi-Driver Gene Prioritization for Precision Oncology. 10<sup>th</sup> Annual Lorne D. Sullivan Lectureship & Research Day. June 21, 2016, Vancouver General Hospital, Vancouver, Canada.
- HIT'nDRIVE: Patient-Specific Multi-Driver Gene Prioritization to Guide Precision Cancer Medicine. Workshop on Network Biology (Algorithmic Challenges in Genomics). April 11-15, 2016, Simons Institute for the Theory of Computing, UC Berkeley, Berkeley, CA, USA. https://simons.berkeley.edu/talks/raunak-shrestha-04-12-16
- Computational Detection and Prioritization of Driver Alterations in Cancer. Terry Fox Seminar, Vancouver Prostate Centre. November 3, 2014, Vancouver General Hospital, Vancouver, Canada.
- HIT'nDRIVE: Multi-Driver Gene Prioritization based on Hitting Time. 18<sup>th</sup> Annual International Conference on Research in Computational Molecular Biology (RECOMB). April 2-5, 2014, Pittsburgh, PA, USA.
- Analysis of KatG Ser315Thr Mutation in Multidrug Resistant Mycobacterium tuberculosis and SLC11A1
   Polymorphism in Multidrug Resistance Tuberculosis in Central Development Region of Nepal Using
   PCR-RFLP Technique: A Pilot Study. 2<sup>nd</sup> International Symposium on Biotechnology. June 29-30, 2010,
   Kathmandu, Nepal

#### Selected Poster Presentations

- Raunak Shrestha, Ermin Hodzic, Thomas Sauerwald, Phuong Dao, Kendric Wang, Jake Yeung, Shawn Anderson, Fabio Vandin, Gholamreza Haffari, Colin C. Collins and S. Cenk Sahinalp.(2017). HIT'nDRIVE: Patient-Specific Multi-Driver Gene Prioritization for Precision Oncology. 4th Canadian Cancer Research Conference, November 4-7, 2017, Vancouver, BC, Canada.
- Raunak Shrestha, Ermin Hodzic, Thomas Sauerwald, Phuong Dao, Kendric Wang, Jake Yeung, Shawn Anderson, Fabio Vandin, Gholamreza Haffari, Colin C. Collins and S. Cenk Sahinalp.(2017). HIT'nDRIVE:

Patient-Specific Multi-Driver Gene Prioritization for Precision Oncology. Genome Informatics, November 1-4, 2017, Cold Spring Harbor Laboratory, NY, USA.

- Raunak Shrestha, Ermin Hodzic, Thomas Sauerwald, Phuong Dao, Kendric Wang, Jake Yeung, Shawn Anderson, Fabio Vandin, Gholamreza Haffari, Colin C. Collins and S. Cenk Sahinalp.(2016). HIT'nDRIVE: Patient-Specific Multi-Driver Gene Prioritization for Precision Oncology. The 27<sup>th</sup> International Conference on Genome Informatics, October 3-5, 2016, Fudan University, Shanghai, China.
- Noushin Nabavi, **Raunak Shrestha**, Yuzhuo Wang and Colin C. Collins (2016). Characterization of intertumor heterogeneity in Malignant Mesothelioma. 13<sup>th</sup> International Conference of the International Mesothelioma Interest Group (iMIG 2016), May 1-4, 2016, Birmingham, UK. (As a first co-author).
- Raunak Shrestha, Ermin Hodzic, Fan Mo, Anna Lapuk, S. Cenk Sahinalp and Colin C. Collins (2015). Computational Detection and Prioritization of Driver Alterations in Prostate Cancer from Multi-Omics Data. Eighth Annual Prostate Cancer Program Retreat (SPORE), March 15-17, 2015, Fort Lauderdale, Florida, USA. (Selected among the top-eight poster of the conference.)
- Raunak Shrestha, Kendric Wang, S. Cenk Sahinalp, Anna Lapuk and Colin Collins (2013). Exploring the Biology of Prostate Cancer Progression using Systems Biology Approach. The Eleventh Asia Pacific Bioinformatics Conference, January 21-23, 2013, Vancouver, Canada.

## **AWARDS AND HONORS**

2018	UBC Translational Cancer Genomics Travel Award (iMig2018).
2016-2018	Mitacs Accelerate PhD Fellowship.
2015-2016	Prostate Cancer Foundation - British Columbia (PCF-BC) Research Awards - 2015.
2014-2016	Faculty of Science Graduate Award, University of British Columbia.
2014	International Society for Computational Biology (ISCB) Travel Fellowship (RECOMB-2014).
2011-2013	CIHR Bioinformatics Training Program for Health Research.

## PROGRAMMING LANGUAGE SKILLS

- R, perl (Expert)
- C, C++, python, php, sql (proficient)

## **SOFTWARE PACKAGES**

- HIT'nDRIVE (https://github.com/sfu-compbio/hitndrive2)
- cd-CAP (https://github.com/ehodzic/cd-CAP)

#### STUDENTS MENTORED

- Hossein Sharifi-Noghabi (Ph.D. Candidate, Computer Science. Simon Fraser University, Canada)
- Jingchao Wei (Ph.D. Candidate, Urology, Central South University, China)

#### ACADEMIC PAPERS PEER REVIEWED

(number of reviews parenthesized)

(3)
(2)
(1)
(1)
(3)
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## PROFESSIONAL MEMBERSHIPS

- Vancouver Bioinformatics User Group (VanBUG): Development Group
- Biotechnology Society of Nepal (BSN): Life time Member and Former Executive Member

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