

Professor of Bioinformatics
Laboratoire d'informatique (LIX)
École Polytechnique
IP Paris

email: sebastian.will@polytechnique.edu
<https://www.lix.polytechnique.fr/~will>

SUMMARY

Full-time Professor of Bioinformatics at École Polytechnique. HDR equiv. 80+ peer-reviewed publications in international journals and conference proceedings (among them various well-cited first author publications); H-index 27 (according to Google scholar). Developer of the widely-used software package LocARNA (the original publication alone was cited 441 times). Class lecturer at École Polytechnique, University of Vienna; formerly, University of Leipzig, University of Freiburg, and MIT.

MAIN RESEARCH INTERESTS

Algorithmic bioinformatics of RNA and protein structure; Structure prediction, comparative analysis, design, interaction, and folding kinetics of RNAs; Genome-wide de-novo prediction of non-coding RNAs; Downstream analysis of high-throughput NGS data; Declarative methods in bioinformatics

SELECTED PUBLICATIONS

H.T. Yao, B. Marchand, S.J. Berkemer, Y. Ponty, **S. Will***. Infrared: a declarative tree decomposition-powered framework for bioinformatics. *Algorithms Mol Biol* 19, 2024.

H. Jabbari, I. Wark, C. Montemagno, and **S. Will*** Knotty: efficient and accurate prediction of complex RNA pseudoknot structures. *Bioinformatics*, 34 no. 22 pp. 3849–3856, 2018.

S. Will, M. Yu, and B. Berger* Structure-based whole genome realignment reveals many novel non-coding RNAs. *Genome Res*, 23 no. 6 pp. 1018–27, 2013.

S. Will, T. Joshi, I. L. Hofacker, P. F. Stadler, and R. Backofen* LocARNA-P: Accurate boundary prediction and improved detection of structural RNAs. *RNA*, 18 no. 5 pp. 900–14, 2012.

S. Will, K. Reiche, I. Hofacker, P. Stadler, and R. Backofen*. Inferring non-coding RNA families and classes by means of genome-scale structure-based clustering. *PLOS Comp Biol*, 3(4):e65, 2007.

GRANTS AND HONORS

French National Research Agency (ANR). Integrated Sequencing and Structural Analysis of RNA Probing Experiments – INSSANE (grant ANR-21-CE45-0034)

German Research Foundation postdoctoral research fellowship: [grant WI 3628/1-1](#)

Doctorate “*summa cum laude*”

Doctoral fellowship of the graduate program “Logic in Computer Science” at LMU, Munich

CONTRIBUTIONS TO THE SCIENTIFIC COMMUNITY

- | | |
|------------|--|
| since 2024 | Co-Chief editor of <i>Algorithms in Bioinformatics</i> (ALMOB, Springer-Nature). |
| since 2014 | PC member of WABI'15, BIOINFORMATICS'17–19, RECOMB'18, '19, & '20, ISMB 2025, ... |
| since 2006 | Reviewer for scientific journals, e.g. <i>Bioinformatics</i> , <i>NAR</i> , <i>Algorithms</i> , and international conferences, e.g. <i>RECOMB</i> , <i>ISMB</i> , <i>WABI</i> |
| 2011 | Associate editor of ALMOB thematic series; 2007 special issue of <i>Constraints</i> |

- since 2006 **Co-chair** of Workshops on Constraints in Bioinformatics (WCB) collocated with international constraint and logic programming conferences CP and ICLP
- since 2004 **Supervision** of students for Bachelor's, Master's/Diploma, and PhD Theses
- since 2000 **Development, maintenance, and deployment** of free scientific software

TEACHING EXPERIENCE (SUMMARY)

- since 2020 **Classes** at École Polytechnique
- 2017–2019 **Classes** at University of Vienna (“Structure and Dynamics of Biomacromolecules”, “Algorithmic Bioinformatics”, “Algorithms and Program Development for Biological Chemistry (APBC)”); all classes taught several times
- 2012–2016 **Classes** at University of Leipzig, including undergrad class “Algorithms and Data Structures” (~ 300 students) and graduate classes on advanced methods in bioinformatics, graph theory, and theoretical biology
- 2011 **Class** 18.417 as instructor of the MIT Math Dept.: “Introduction to Computational Molecular Biology — Foundations of Structural Bioinformatics”; as TA, with Bonnie Berger: Seminar 18.418 “Topics in Computational Biology”
- 2006–2012 **Classes** at Department of Computer Science, University Freiburg, Germany, covering “RNA Bioinformatics”, “Protein Folding and Energy Landscapes”, “Bioinformatics I+II”, “Constraint Programming”, “Structure Prediction in Simplified Protein Models” (all classes in English)
- 2006 **Invited Class** at Summer School BCI (Biology, Computation, and Information) on “Optimal Constrained-Based Protein Structure Prediction”
- 2002–2005 **Classes** at Department of Computer Science, University Jena, Germany covering class “Simplified Protein Models”, seminar “Constraint-Satisfaction-Problems in Bioinformatics”, lab class “Data Mining & Sequence Analysis”

POSITIONS AND FELLOWSHIPS

- since 2016 **Researcher** (Universitätsassistent, post-doc) at University of Vienna with Prof. Ivo Hofacker
- 2012–2016 **Researcher** at University of Leipzig with Prof. Peter Stadler
- 2005–2012 **Akademischer Rat** at the Albert-Ludwigs-University in Freiburg with Prof. Rolf Backofen
- 2011 **Instructor** at Math Department, Massachusetts Institute of Technology (MIT)
- 2011 **Postdoctoral Associate** at Computer Science and Artificial Intelligence Lab (CSAIL), MIT, with Prof. Bonnie Berger
- 2010 **Postdoctoral Fellow**; Scholarship granted by the German Research Foundation, CSAIL, MIT, with Prof. Bonnie Berger
- 2002–2005 **Researcher** at University Jena, Bioinformatics Group
- 2000–2002 **Doctoral Fellow**; granted by PhD program “Logic in Computer Science”; LMU Munich; Advisers: Prof. Peter Clote, Rolf Backofen

DEGREES

- 2005 **Dr. rer. nat. in Computer Science** — *Summa Cum Laude*
Friedrich-Schiller-University Jena, Germany

2000 [Diplom\(≈ M.Sc.\) in Computer Science](#) — *best in class*
Ludwig-Maximilians-University Munich, Germany

SELECTED SOFTWARE

2024 [LocARNA 2.0](#) - Versatile simultaneous alignment and folding of RNAs (C++, Perl)
2022– [Infrared](#). A declarative tree decomposition-powered framework for bioinformatics
(C++, Python)
2018 [RNARedprint](#). FPT Sampling of RNA designs with multiple target structures (C++,
Python)
2018 [Knotty](#). Space-efficient prediction of CCJ pseudoknots (C++)
2016–2019 [GitCATS](#) Git-based multilingual continuous integration testing system (written for
teaching a programming class (APBC), where—using Github—students submit
and revise their programs, which the system tests automatically) (Python)
2015 [SparseMFEFold](#). Space-efficient RNA structure prediction (C++)
since 2012 [SPARSE](#). Very efficient structure-based alignment of RNAs; distributed as part of
the LocARNA package (C++)
2011–2013 [REAPR](#). A pipeline for fast structure-based realignment for de-novo prediction of
structural non-coding RNAs in whole eukaryotic genomes (C++, Python)
since 2010 [CARN](#). Constraint-based Alignment of RNAs with structures of unrestricted
complexity — a competitive tool for alignment of pseudoknots and multi-stable
RNAs (C++, Gecode)
since 2009 [LocARNA-P](#). Reliabilities of RNAs sequence-structure alignment — LocARNA-P
computes reliabilities and match probabilities of sequence-structure alignments
and enables advanced comparative RNA analysis. (C++, Perl)
since 2005 [LocARNA](#). Multiple alignment of RNAs — LocARNA is one of the most accurate
and fastest tools for pairwise and multiple alignment of RNA. LocARNA is enjoying
widespread, active use in RNA research, is constantly extended, and inspires
further algorithmic development. (C++, Perl)
2006 [CTE-Alignment](#). Constrained alignment by Cluster-Tree-Elimination — declara-
tive integration of complex constraints preserving efficiency (C++)
2000–2005 [PSP](#). Constrained-based Protein Structure Prediction — first exact prediction in
face-centered cubic and cubic models; PSP has set the foundations for the CPSP
package. (Mozart/Oz)

SELECTED TALKS AND VISITS

2024 [Talk](#) Benasque RNA meeting
2022 [Talks](#) Benasque RNA meeting
2019 [Talk](#) WABI'19 (algorithmic bioinf. conference), Niagara Falls NY
2019 [Keynote Talk](#) at meeting of the former DFG Priority Program SPP 1258, Freiburg
2019 [Invited Talk](#) and visit at École Polytechnique, Palaiseau
2018 [Talk](#) at RECOMB (major algorithmic bioinformatics conference), Paris
2018 [Talk](#) at the Vienna RNA meeting, Vienna
2018 [Talk](#) at the Benasque RNA meeting (E. Rivas, E. Westhof), Benasque
2017 [Invited Talk](#) at Ribonets workshop, Düsseldorf
2015 Two [Talks](#) at the Benasque RNA meeting (E. Rivas, E. Westhof), Benasque
2014/15 [Talks](#) at WABI'15, Atlanta and WABI'14, Wrocław
2012/13 [Talk](#) at RECOMB'12 in Vancouver and two [Talks](#) at RECOMB'13, Beijing

2010	Invited Talk Prof. Jérôme Waldispühl, McGill, Montreal
2010	Invited Talk Prof. Cenk Sahinalp, SFU, Vancouver
2009	Talk at the Benasque RNA meeting, Benasque
2008	Invited Talks Prof. Bonnie Berger, MIT, Cambridge MA

SELECTED SCIENTIFIC CONTRIBUTIONS

Structure-Based Comparative Analysis of RNAs

<i>Motivation</i>	Structural non-coding RNAs (ncRNAs) play an essential role in cells. Analysis of RNA requires fast and space efficient structure-based alignment methods. However, the gold-standard for this purpose has prohibitive complexity. Fast methods enable new biological applications.
<i>Results</i>	The LocARNA software package — time and space-efficient algorithms for multiple alignment of RNA by simultaneous alignment and folding. The tool LocARNA has been used for large scale clustering of structural RNAs. SPARSE improves ensemble-based sparsification over LocARNA, resulting in even more efficient alignment and folding. LocARNA-P extends the approach for computing reliabilities of sequence-structure RNA alignment. I have applied the tool in the modENCODE project and revealed many novel ncRNA candidates in <i>Drosophila melanogaster</i> based on a further algorithmic extension of LocARNA .

Prediction of RNA-RNA-Interaction, Pseudoknot Prediction and Alignment

<i>Motivation</i>	Folding and alignment of RNA molecules is indispensable for current biological research. However, for considering pseudoknots in the study of RNA-RNA-interaction, efficiency is still limiting.
<i>Results</i>	Applying a sparsification technique, I developed sparsified algorithms to predict RNA-RNA-interaction and RNA pseudoknots , which significantly improves performance and space consumption. Furthermore, I developed fixed parameter-tractable and polynomial algorithms for the alignment of RNA pseudoknots .

Structure Prediction in 3D Protein Models and Symmetry Breaking in Constraint Search

<i>Motivation</i>	Protein models are valuable tools for the study of protein folding and evolution, as well as for benchmarking heuristic optimization. Many of these applications require optimal structure prediction, however effective approaches for unrestricted models didn't exist. Symmetry breaking is often essential for the efficient solving of constraint models, but most specialized methods interfere with the search heuristic and complicate constraint modeling.
<i>Results</i>	I developed a fast approach for optimal prediction of structures in cubic and face-centred-cubic HP and HPNX protein models . The implementation CPSP is freely available and is used in protein studies. Motivated by the work on structure prediction, I devised a general symmetry breaking scheme that works by introducing constraints during the search . The scheme has been used in a number of constraint models, among them our own models for protein structure prediction. The idea contributed significantly to the advancement of the field.