

# Gianluca Della Vedova | CV

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## Current position

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Associate professor with tenure (SSD INF/01 — Informatica)  
Dipartimento di Informatica, Sistemistica e Comunicazione  
Università degli Studi di Milano – Bicocca

## Positions

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<b>Associate Professor</b> <i>Dept. of Informatics, Systems, and Communications</i>	<b>Univ. Milano – Bicocca</b> 10/2012–today
<b>Associate Professor</b> <i>School of Statistics</i>	<b>Univ. Milano – Bicocca</b> 10/2005–09/2012
<b>Assistant Professor</b> <i>School of Statistics</i>	<b>Univ. Milano – Bicocca</b> 05/2001–09/2005

## Habilitation

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**Abilitazione a Professore di I fascia, settore concorsuale 01/B1**  
*Italian habilitation to be Full professor of Computer Science* 08/2018–today

## Education

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**Ph.D. in Computer Science** **Università degli Studi di Milano**  
1996–2001  
Thesis: “Multiple Sequence Alignment and Phylogenetic Reconstruction: Theory and Methods in Biological Data Analysis”

**M.Sc. in Computer Science** **Università degli Studi di Milano**  
1990–1995  
Thesis: “Sequential and Parallel Algorithms for Graph Decomposition”

## Funding

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**Leading**.....

**2020–2023:** H2020-MSCA-RISE-2019 (Univ. Milano – Bicocca amount: 197 800€) Pan-genome Graph Algorithms and Data Integration (PANGAIA) — Horizon 2020 Marie Skłodowska-Curie Research and Innovation Staff Exchange programme.

This project is coordinated by a co-leader of my research lab and it involves a post-doc I supervise (Simone Ciccolella), another post-doc a co-supervise, and 2 PhD students (that I co-supervise).

The main goal of this project is to foster collaborations between different research groups in computational pangenomics. I have been one of the main participants in planning and writing the proposal. I am **leading WP5** Communication and Dissemination.

The project involves Universität Bielefeld (Germany), Centrum Wiskunde & Informatica (the Netherlands), The Institut Pasteur (France), Comenius University Bratislava (Slovakia), Geneton s.r.o. (Slovakia), Cornell University (USA), University of Tokyo (Japan), Simon Fraser University (Canada), Pennsylvania State University (USA), University of California (USA). I have developed an especially strong collaboration with Prof. Iman Hajirasouliha at Cornell University on tumor phylogeny inference and metagenomics where also Simone Ciccolella is actively working.

**2021–2024:** H2020-MSCA-ITN (Univ. Milano – Bicocca amount: 261 499,68€) ALgorithms for PAngenome Computational Analysis (ALPACA) — Horizon 2020 Marie Skłodowska-Curie Innovative Training Network.

As part of the activities of this project I am **co-supervising** a PhD student. The project involves Universität Bielefeld (Germany), Centrum Wiskunde & Informatica (the Netherlands), The Institut Pasteur (France), Comenius University Bratislava (Slovakia), Geneton s.r.o. (Slovakia), University of Helsinki (Finland), European Molecular Biology Laboratory — European Bioinformatics Institute, Heinrich-Heine-Universität Düsseldorf (Germany), Università di Pisa (Italy), CNRS (France), University of Cambridge (UK), INRIA (France), Pendulum Therapeutics, Inc. (USA), Vrije Universiteit Amsterdam (the Netherlands), Oxford Nanopore Technologies plc (UK), DNANexus (US), Sorbonne University (France), Deinove (France), Finnish Red Cross (Finland), BaseClear B.V. (the Netherlands), French Alternative Energies and Atomic Energy Commission (France), Cornell University (USA), University of Tokyo (Japan), Simon Fraser University (Canada), Pennsylvania State University (USA), University of California (USA).

**2013–2016:** Modulation of anti-cancer immune response by regulatory non-coding RNAs — Fondazione Cariplo 2013. I have been in **charge of the bioinformatics WP**. The project had two research units: the National Institute of Molecular Genetics and the Univ. Milano – Bicocca.

During this project, I have been the **scientific supervisor of two 1-year post-docs** and I have co-supervised a third postdoc.

**2011–2014:** MIUR/Regione Lombardia 2011 (Univ. Milano – Bicocca amount: 199 991€). Piattaforma di Analisi Traslazionale Integrata (PATRI). I have been **in charge of all bioinformatics aspects**.

This project has led to a 1-year postdoc (23 000€) and 3 short-term research contracts (each between 10 000€ and 12 000€). I have co-supervised the postdoc and I have been the only supervisors of all short-term contracts.

**2016:** Fondo di Ateneo 2016 (12 490€). Modelli computazionali e algoritmi: aspetti teorici e sperimentali, con applicazioni alla Bioinformatica. **PI**.

**2015:** Fondo di Ateneo 2015 (10 980€). Algoritmi combinatori e modelli di calcolo: aspetti teorici e applicazioni in Bioinformatica. **PI**.

**2014:** Fondo di Ateneo 2014 (12 186€). Algoritmi e modelli computazionali: aspetti teorici e applicazioni nelle scienze della vita. **PI**.

**2013:** Fondo di Ateneo 2013 (9 337€). Metodi algoritmici e modelli: aspetti teorici e applicazioni in bioinformatica. **PI**.

**2011:** Fondo di Ateneo 2011 (4 055€). Tecniche algoritmiche avanzate in Biologia Computazionale. **PI**.

**2006:** Grandi Attrezzature 2006 (40 000€). Laboratorio Virtuale Statistico-Territoriale — Virtual

Statistical–Territorial Laboratory. This was a joint project between the Statistics Department and the Dept. of Sociology and I have been one of the two people **in charge of the entire project** (and the only one from the Statistics Department). It has resulted in the acquisition of 2 servers that are used to provide services to research and teaching activities on territorial sociology and computational statistics. **co-PI**.

**2005–2008:** MIUR/PRIN 2005, Potenzialità e ottimizzazione delle banche dati automatizzate in epidemiologia. **In charge of all bioinformatics aspects.**

#### **As a Participant** .....

**2013–2016:** Regione Lombardia. SPAC3 — Servizi smart della nuova Pubblica amministrazione per la Citizen-Centricity in cloud.

**2011–2014:** MIUR/PRIN 2011 Automi e Linguaggi Formali: Aspetti Matematici e Applicativi

**2003:** MIUR/FIRB 2003. Bioinformatica per la Genomica e la Proteomica.

**2000–2001:** NSF CCR-9988353, ITR-0085910.

**1999–2001:** MURST COFIN 98 “Bioinformatica e ricerca genomica”.

**1994–1995:** MURST 40% “Algoritmi e strutture di calcolo”.

**1994–1995:** ESPRIT-BRA ASMICS 2 n. 6317.

## **Teaching**

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I have started teaching when I have been hired as an Assistant professor at the School of Statistics in 2001. I was the first Computer Science faculty of the school, therefore I had to design new courses tailored to students with a good mathematical background, but with no Computer Science experience. During my career I had to design a course several times, partly due to the evolution of the background of the students and to changes in the overall degree in Statistics and Statistical Economics, partly because I had to teach also to students majoring in Computer Science or in Data Science.

My teaching philosophy is based on active learning. This requires important and continuous interactions between teacher and students, requiring the development of problems that students have to attack, as a group or solo. Designing an active learning course is usually innovative, since existing courses are mostly designed for more traditional approaches, where students have a more passive role and acquiring problem solving skills is not a main focus.

#### **Ph.D. courses taught** .....

**2016, 2018, 2020:** *Advanced Algorithms*, PhD in Computer Science, Univ. Milano – Bicocca. I have designed the course, since it had never been taught before.

**2021, 2023:** *Graph Theory and Algorithms*, PhD in Computer Science, Univ. Milano – Bicocca. I have co-designed the course, since it had never been taught before.

#### **Courses taught — in charge of the course** .....

**2022–today:** *Large-Scale Graph Algorithms*, M.Sc. in Computer Science, Univ. Milano – Bicocca (6 ECTS). I have designed the course, since it had never been taught before.

**2021–today:** *Laboratorio di Informatica (Computer Science Lab)*, B.Sc. in Statistics, B.Sc. in Statistical Economics, Univ. Milano – Bicocca (3 ECTS). I had fully redesigned the course.

**2017–today:** *Foundations of Computer Science*, M.Sc. in Data Science, Univ. Milano – Bicocca. (6 ECTS). I have designed the course, since it had never been taught before.

**2014-today:** *Elementi di Bioinformatica (Elements of Bioinformatics)*, B.Sc. in Computer Science, Univ. Milano – Bicocca (8 ECTS). I have designed the course, since it had never been taught before.

**2001–2020:** *Basi di Dati (Databases)*, B.Sc. in Statistics, B.Sc. in Statistical Economics, Univ. Milano – Bicocca (6 ECTS).

**2007-2022:** *Bioinformatica (Bioinformatics)*, M.Sc. in Biostatistics, Univ. Milano – Bicocca (6 ECTS). I have designed the course, since it had never been taught before.

**2010–2013:** *Algoritmi su stringhe (Text Algorithms)*, B.Sc. in Computer Science, Univ. Milano – Bicocca. I have designed the course, since it had never been taught before.

**2008:** *Strumenti informatici per la statistica M (Computational Tools for Statistics)*, M.Sc. in Biostatistics, blended e-learning, Univ. Milano – Bicocca. I have designed the course, since it had never been taught before.

**2007–2009:** *Informatica Applicata S (Applied Computer Science)*, M.Sc. in Biostatistics (2 ECTS), Univ. Milano – Bicocca. I have designed the course, since it had never been taught before.

**2001–2008:** *Laboratorio Statistico-Informatico (Computational Statistics Lab)*, all B.Sc. programs of the School of Statistics (6 ECTS), Univ. Milano – Bicocca. I have designed the course, since it had never been taught before.

**2001–2008:** *Programmazione e Basi Dati (Programming and Databases)*, all B.Sc. programs of the School of Statistics (6 ECTS), Univ. Milano – Bicocca. I have designed the course, since it had never been taught before. The course has changed name to "Basi di Dati". It was the first database course in Italy designed for students in Statistics.

**2008:** *Fondamenti di Informatica (Fundamentals of Computer Science)*, M.Sc. in Biostatistics, Univ. Milano – Bicocca. I have designed the course, since it had never been taught before.

**2006:** *Laboratorio di Informatica (Computer Science Lab)*, all B.Sc. programs of the School of Statistics (6 ECTS), Univ. Milano – Bicocca. I have designed the course, since it had never been taught before.

### Postgraduate courses.....

**2021:** *Beyond Genome Assembly*, Master “qOmics: quantitative methods for Omics Data”, Univ. Milano – Bicocca and Univ. Pavia.

**2012, 2014:** *Data Base e Sistemi Informativi (Databases and Information Systems)*, Master di primo livello in Amministratore di Sistema per la Diagnostica per Immagini (Master in Image Diagnostics Systems Administrator), Univ. Milano – Bicocca.

**2007, 2010, 2011:** *Fondamenti di Informatica (Fundamentals of Computer Science)*, Master di primo livello in Amministratore di Sistema per la Diagnostica per Immagini (Master in Image Diagnostics Systems Administrator), Univ. Milano – Bicocca.

**2003:** *Fondamenti di Informatica e Elementi di Programmazione (Fundamentals of Computer Science and Elements of Programming)*, Master di primo livello in Bioinformatica (Master in Bioinformatics), Univ. Milano – Bicocca.

**2001–2002:** *Sistemi Informatici e Elementi di Programmazione (Introduction to Programming)*, Master di primo livello in Bioinformatica (Master in Bioinformatics), Univ. Milano – Bicocca.

## Supervision

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### Postdocs.....

I have been **scientific supervisor** of the following postdoctoral positions:

**2022–today:** Simone Ciccolella. 2-year postdoc position on *Computational Approaches for tumor and viral phylogenies*.

**2018–2019:** Murray Patterson. 2-year postdoc position on *Haplotype assembly from sequencing reads*. Currently Assistant Professor (Tenure-track) at Georgia State University.

**2015:** Hassan Mahmoud Mohamed Ramadan Mohamed. 1-year postdoc position on *Methodology for treatment and data analysis of NGS data for the detection of alternate splicing events*.

Moreover, I have co-supervised the following postdocs:

- Luca Denti
- Marco Previtali
- Stefano Beretta

### PhD students supervision.....

**Supervisor of**

**2021–2024 (expected):** Jorge Avila Cartes — PhD in Computer Science

**2017–2021:** Simone Ciccolella, “*Algorithms for cancer phylogeny inference*” — PhD in Computer Science. Currently Post-doc at Università di Milano – Bicocca.

**2013–2017:** Marco Previtali, “*Self-indexing for de novo assembly*” — PhD in Computer Science. Currently at Bloomberg

**2008–2012:** Stefano Beretta, “*Algorithms for Next Generation Sequencing Data Analysis*” — PhD in Computer Science. Currently at San Raffaele Telethon Institute for Gene Therapy

**2001–2004:** Riccardo Dondi, “*Computational Problems in the Study of Genomic Variations*” — PhD in Computer Science. Currently Associate Professor at Università di Bergamo

Moreover, I have actively contributed to the training of the following PhD students:

- Giulia Bernardini — PhD in Computer Science. Currently Assistant Professor at Univ. Trieste.
- Simone Zaccaria — PhD in Computer Science. Currently Group Leader presso Department of Oncology, Univ. College London
- Anna Paola Carrieri — PhD in Computer Science. Currently Research Staff Member presso IBM Research Lab, UK

### M.Sc. students supervision.....

I have supervised or co-supervised the thesis of more than 10 M.Sc. students in Computer Science, Statistical Economics, Biostatistics, Data Science at University of Milano–Bicocca and at the University of Milan. I am currently supervising the thesis of a M.Sc. in Bioinformatics at the Pwani University Biosciences Research Centre, Kilifi (Kenya), as the result of a long-standing collaboration with Pjotr Prins (University of Tennessee).

I have supervised the activities related to the **Exchange mobility EXTRA UE** program of:

- Ramesh Rajaby ha spent 6 months with the research group led by prof. Jesper Jansson (Kyoto University, Japan). Ramesh Rajaby is currently postdoc at National University of Singapore.
- Simone Ciccolella has spent 3 months with the research group led by prof. Iman Hajirasouliha (Weill Cornell Medicine, New York). Simone Ciccolella is currently a postdoc under my supervision.

## B.Sc. students supervision.....

I have supervised the final projects of more than 50 B.Sc. students in Computer Science, Statistics, and Statistical Economics.

## Awards

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### Best Poster Awards.....

**Recomb 2018:** Luca Denti, Raffaella Rizzi, Stefano Beretta, Gianluca Della Vedova, Marco Previtali, and Paola Bonizzoni. *ASGAL: Aligning RNA-Seq Data to a Splicing Graph to Detect Novel Alternative Splicing Events*

## Professional activities

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### Steering Committee Member.....

**2018-today: Steering Committee member** of Computability in Europe (CiE). During the same period I have also been the Executive Officer of the Steering Committee.

### Program Committee Chair.....

**2023: Program Committee Chair** of Computability in Europe (CiE) 2023. This international conference will take place in Batumi (Georgia) in July 24-28 and is the flagship conference of the Association Computability in Europe that has more than 1000 members and focuses on computability science in all its multidisciplinary facets. The conference proceedings will be published in the LNCS series.

**2021: Chair** of the Data Structure in Bioinformatics Workshop (DSB 2021), February 11-12, 2021. The workshop had 19 research talks given from international scholars.

**2021: Organizer**, of Pangenome Bio Hacking (PGBH) 2021 (co-organized with Erik Garrison, Enza Colonna, Pjotr Prins). This online conference has taken place on December 9-10, 2021, and we have invited 9 speakers from Europe and the US. It has been the first world-wide conference for researchers who have a focus on pangenomics free and open source software development.

**2020: Organizer**, of the Computability in Europe special session “*Large Scale Bioinformatics and Computational Sciences*” (co-organized with Iman Hajirasouliha, Weill-Cornell Medicine, USA). We have selected and invited four speakers who have given an invited talk.

### PhD schools Board.....

**2022: Supervisory Board Member**, PhD Summer School “Introduction to Pangenomics”, held at the Lake Como School of Advanced Studies, July 4-8, 2022. The school has been attended by 25 international PhD students and had 12 teachers from Europe and the US. The Supervisory Board had 3 members who have decided the full program of the school.

### Program Committee Member.....

#### Program Committee Member:

- Computability in Europe (CiE) 2013, 2019, 2020, 2022, 2023
- Workshop on Algorithms in Bioinformatics (WABI) 2020, 2022, 2023
- Italian Conference on Theoretical Computer Science (ICTCS) 2023
- Combinatorial Pattern Matching (CPM) 2019
- Symposium on String Processing and Information Retrieval (SPIRE) 2017
- Bioinformatics Open Source Conference (BOSC) 2016–2021, 2023

**2020:** **Editor** of the proceedings of the conference Computability in Europe 2020. The proceedings have been published in the LNCS 12098 volume, Springer-Verlag.

#### **Editorial Board Member**.....

**2016–2019:** **Editorial Board** member of the journal “Advances in Bioinformatics”.

**2004:** **special issue editor**, Journal of Computer Science and Technology.

#### **PhD defence committee member**.....

**2023:** Evaluation committee member, PhD thesis in Computer Science, Univ. Milano.

**2018:** Evaluation committee member, PhD thesis in Computer Science, Univ. Roma la Sapienza.

**2018:** Evaluation committee member, PhD thesis in Computer Science, Univ. Milano.

#### **Competition evaluation member**.....

**2023:** Evaluation committee member, PhD positions, Technical University of Denmark.

**2019:** Evaluation committee member, PhD positions, Univ. Milano–Bicocca.

**2017:** Evaluation committee member, competition for an Assistant Professor position (valutazione comparativa per RTDb), Univ. Milano

**2013:** Evaluation committee member, PhD positions, Univ. Milano–Bicocca.

**2006:** Evaluation committee member, competition for an Assistant Professor position (valutazione comparativa per ricercatore universitario), Univ. Milano

#### **Grant proposal evaluation**.....

**2023:** **National Science Centre Poland.** Review of a grant proposal under the Funding scheme OPUS.

**2021:** **National Science Centre Poland.** Review of a grant proposal under the Funding scheme SONATA.

**2014:** **University of Catania.** Review of a grant proposal under the Funding scheme FIR.

#### **Incoming visits**.....

**2018:** I have been in charge of hosting Prof. Vladimir Filipovic (Associate Professor at University of Belgrade, Faculty of Mathematics, Department of Computer Science) during his sabbatical from 20/01/2018 to 30/09/2018. During his visit, we have collaborated on the development of metaheuristics in Bioinformatics

#### **Reviewer Service**.....

I have reviewed papers for the following journals:

- ACM/IEEE Transactions on Computational Biology and Bioinformatics
- Algorithmica
- Algorithms
- Bioinformatics
- Briefings in Bioinformatics
- Graphs and Combinatorics
- Information Processing Letters
- Journal of Computational Biology
- INFORMS J. Computing
- Journal of Computer Science and Technology
- Theoretical Computer Science



○ Theory of Computing Systems

**University Service.....**

**2021-today: Scientific Committee Member** Postgraduate Master Programme “quantitative methods for Omics Data” (EQF level 8), University of Milano – Bicocca

**2020-today: Representative of the Università degli Studi di Milano – Bicocca** Joint Research Unit ELIXIR IIB General Assembly, Italian Node of Elixir Europe, the main intergovernmental European organization in Bioinformatics.

**2019-today: Deputy coordinator of the PhD program** in Computer Science, Univ. Milano – Bicocca

**2018-today: Quality Assurance**, M.Sc. in Data Science.

**2019-today: President of the Teaching Committee (Commissione Didattica)**, M.Sc. in Data Science.

**2018-today: PI** of Università degli Studi di Milano – Bicocca of the MoU (Convenzione quadro) with Istituto Nazionale di Genetica Molecolare (National Institute of Molecular Genetics).

**2020-today: Scientific Committee Member**, Bicocca Ambiente Società Economia, by decree of the rector.

**2018-today: Member of the Students-Professors Committee (Commissione Paritetica Docenti-Studenti)**, Dipartimento di Informatica, Sistemistica e Comunicazione

**2013-today: Board Member**, PhD program in Computer Science, Univ. Milano – Bicocca

**2011-today: Executive Committee (Comitato direttivo) Member**, Centro di Produzione Multimediale di Ateneo (Multimedia Production Center), by decree of the Academic Senate.

**2015–2020: Local Technical Coordinator**, Univ. Milano – Bicocca, Elixir IIB

**2016-today: In charge of Elixir IIB training activities** in Univ. Milano – Bicocca. I have organized course on “Genome Assembly and Annotation”, “Data Carpentry Workshop”, “Software Carpentry Workshop”, “Docker Advanced”, and “Exome analysis with Galaxy”.

**2002–2012: Representative of the Computer Science Area**, School of Statistics.

**2004–2012: Representative of the School of Statistics**, University Committee on Computer Science.

**2010–2012: Delegate of the Dean on e-learning** School of Statistics.

**2007–2012: In charge of the Computer Labs** School of Statistics.

**2010–2012: Member of the e-learning committee**, School of Statistics.

**2002–2004: Representative of the Computer Science Area**, School of Statistics Committee in charge of designing the B.Sc. program in Statistics (Statistica e Gestione delle Informazioni). The program is still active.

**2004: Representative** of the School of Statistics for the short course “Information Technology For Problem Solving (IT4PS)”, organized by the CRUI Foundation.

**2003: In charge** of the course 167388 “Laboratorio Complementare di Informatica per Statistici (Elective Computer Science Lab for Statistics)”, part of the FSE 156165 project “Progetto Quadro Università degli Studi di Milano – Bicocca”. The course focused on intermediate Computer Science skills for statistics students. The course has been fully funded by the European Commission.



## Speaker

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**GGI Seminar Series, March 2022:** Invited by Department of Genetics, Genomics and Informatics, University of Tennessee. Title of the talk: “Computational Aspects of Models of Evolution”

**invited speaker:** Computability in Europe 2017 special session “Algorithmics for Biology”

**speaker at ISMB 2001:** I have presented a paper as first author — and the only author of my University — to the conference Intelligent Systems for Molecular Biology 2001: the main Bioinformatics scientific conference (**CORE: A, Microsoft Academic: A+**)

## Research activity

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I have published more than **50 journal papers** with more than **1800 citations** and **h-index 20** (from Google Scholar). According to Google Scholar, 7 of my papers have at least 100 citations each, and I have 166 coauthors.

From 2016 to 2019 I have been the founder and the leader of the Research Lab “AlgoLab — Experimental Algorithmics Lab”, at the Università di Milano – Bicocca. The research lab has several international collaborations and it focuses on algorithm design, implementation, and experimental analysis on large datasets. In fact, the research activities regards both methodological and experimental aspects, leading to the development of software tools to attack bioinformatics problems. In 2019 the lab has merged with the Bioinformatics Lab, resulting in the Bioinformatics and Experimental Algorithmics (BIAS) Lab which I am effectively co-leading with Paola Bonizzoni.

My personal research activity has focused on the development of combinatorial algorithms in Bioinformatics, with a strong attention on foundational aspects. Moreover, I have paid special attention to algorithm implementation and validation, from both a theoretical and an experimental point of view.

Bioinformatics algorithmics has gained relevance with the establishment of the Human Genome project whose main goal is to determine the influence that biological sequences (such as DNA, RNA) have on living beings. Completing the sequencing of the human genome was only the first fundamental step; finding which proteins are expressed by each gene and determining the interaction among the various DNA sites are among the most important open problems. The huge quantity of data to analyze (remember that human DNA contains 3 billions nucleotides) makes computers and efficient algorithms a needed cornerstone of the field. My research has been mainly devoted to the design of such efficient algorithms and can be detailed in a few subfields.

### Sequence comparison.....

The central dogma in Computational Biology states that sequence homology leads to functional homologies (that is similarity among the effects performed by such sequences), therefore it is of the utmost importance to have some computational tools for comparing sequences. In this direction the multiple sequence alignment problem has been formally introduced in (Altschul, Lipman SIAM J. Appl. Math. 1989), even though it had already been studied previously, and some of its variant are intractable. My focus is on the study of the computational and approximation complexity and on the development of efficient algorithms for the classical notions of longest common subsequence and shortest common supersequence and for sequence alignment.

Most recently, my focus is on computational pangenomics, where a large set of genomes is considered and represented as a (pangeome) graph. In this case, my main contribution has been the introduction of a formal notion of the computational problem of pangenome graph construction, which in turn allows to analyze existing and new algorithms for the problem. Moreover, I studied efficient and practical sequence-to-graph alignment algorithms that fully exploit the topology of the graphs to infer recombination events — this problem was not previously attacked with approaches based on sequence alignment.

#### **Journal papers**

[J3] [J6] [J38] [J41] [J45] [J54] [J55]

#### **Software**

○ RecGraph <https://github.com/AlgoLab/RecGraph>

#### **Collaborations**

- Universitaet Bielefeld (Alexander Schoenhuth)
- University of California at Santa Cruz (Jouni Siren)

### **Phylogeny reconstruction and comparison**.....

Reconstructing phylogenies is another problem that has great relevance in Computational Biology, as a phylogeny is an intuitive representation of a common evolutionary history of a set of extant species. In this setting I have studied the quartet-based reconstruction technique (a quartet is the optimal solution over four species), developing new algorithms to clean some of the error that inherent in the use of such technique.

More recently, I have studied the algorithmic implications of new models of evolution, especially for tumor and viral phylogeny. In fact, most of the literature focuses on a very restrictive model of evolution where mutations can only be acquired. Cancer evolution is too complex for that model, in fact losses of entire regions of the genome is common. Unfortunately, this makes the computational problem much harder, since the solution space explodes. My contributions on these aspects spans from mathematical exploration of combinatorial properties of phylogenies to efficient and practical algorithms to infer phylogenies, leading their design, development, and analysis.

The phylogeny comparison problem is fundamental when you have to compare the results of various experiments on the same set of species. I have extensively studied the problem In (Amir e Keselman, SIAM J. Comp. 1997) some formulations of the problem have been introduced. My initial contributions have been on the approximability of the maximum common isomorphic subtree, and on efficient algorithms to reconcile gene and species trees. Most recently, I have shifted my focus on practical similarity measures between tumor phylogenies. In fact, there are several tools for inferring tumor phylogenies, but the research community needs a measure to understand which of the phylogenies are more similar then others.

#### **Journal papers**

[J5] [J7] [J10] [J11] [J12] [J18] [J21] [J26] [J42] [J50] [J53]

#### **Software**

- SASC <https://github.com/sciccolella/sasc>
- MP3 <https://github.com/AlgoLab/mp3treesim>
- ggpff <https://github.com/AlgoLab/gppf>
- ggps <https://github.com/AlgoLab/gpps>

#### **Collaborations**

- Georgia State University (Murray Patterson)
- Weill Cornell Medicine (Iman Hajirasouliha)

## Clustering

The problem of classifying data in similar sets is one of the most important problems in Computer Science; it is common to have a similarity measure between pairs of elements and to aim at computing a partition of the elements so that elements in a common sets are similar while elements in different sets are not similar. In this field I have studied the correlation clustering problem on weighted graphs, which has the important property that the number of clusters is not fixed a priori, but depends on the dataset analyzed. My contribution has been to develop the connections with another version of clustering, called consensus clustering, showing that an interesting restriction of the problem is NP-complete and providing two polynomial-time approximation schemes for a different formulation of the problem.

Moreover I have studied a different clustering problem, closely related to the analysis of microarray data. In this case, data are represented as vectors on  $\{0, 1, N\}$  alphabet, where  $N$  stands for missing or undecided data. His contributions consist of proving that some restrictions of the problem are APX-hard and designing a polynomial-time constant-factor approximation algorithm. Moreover, I have designed an efficient algorithm for a different restriction of the problem.

Another research topic related to Clustering is the  $k$ -anonymity problem, where we want to cluster the rows of a matrix so that each cluster has identical entries, except for  $k$  columns, where I have designed some algorithms that are efficient for small values of  $k$ . This formulation has originally appeared in data privacy, but I am investigating its applicability to pangeneome privacy.

Tumor phylogeny inference methods are especially computationally expensive. For this reason, I have developed some clustering techniques that are tailored for clustering single-cell tumoral data. These new approaches are based on community detection, and I have proved experimentally that they result in faster and more precise phylogenies.

### Journal papers

[J1] [J2] [J9] [J29] [J31] [J35] [J36] [J40]

### Software

- plastic <https://github.com/plastic-phy/plastic>
- CouGaR-g <https://github.com/AlgoLab/CouGaR-g>
- MALVIRUS <https://algolab.github.io/MALVIRUS/>
- celluloid <https://github.com/AlgoLab/celluloid>

### Collaborations

- Georgia State University (Murray Patterson)

## Alternative Splicing

Alternative splicing is the biological mechanism that allows a gene to encode and produce more than one protein and it is correlated to the onset of several diseases. In this field I have focused on efficient algorithms for detecting novel (that is not previously known) alternative splicing events.

I have supervised the development of two approaches to infer novel alternative splicing events, based on the alignment of reads against a transcriptome. In the second approach, the alignment is against a graph transcriptome which has been one of the very first papers introducing graph-based representation of a set of transcripts. I have supervised the algorithm design and the experimental analysis.

The sheer amount of data requires some efficient filtering. I have developed a fast and accurate filter, based on Bloom filters, that is able to extract from a sample on the much smaller reads

that originate from a given gene, allowing to use a more precise, albeit slower, downstream tool. I have developed the algorithm and supervised the experimental analysis.

#### **Software**

- Pintron <https://github.com/AlgoLab/PIntron>
- ASGAL <https://github.com/AlgoLab/galig>
- Shark

#### **Main collaborations**

- CNR (Graziano Pesole)

#### **Journal papers**

[J6] [J16] [J24] [J33]

### **DNA microarray design**.....

The introduction of DNA microarray have greatly increased the throughput of experimental data in Molecular Biology. Such technology (Drmanac et. al. Science 1991) has given relevance to some computational problems on the optimal microarray synthesis or on the experimental data analysis.

More precisely, classifying microbial communities can be performed only exploiting microarrays, as the microbial external aspect is hard to study, due to their extremely small size. A fundamental computational problem in this field is computing the minimum set of substrings that are able to distinguish a set of strings (Probe Selection). The importance of this problem is due to the fact that it formalizes the search for the cheapest experiment obtaining the desired result.

I have proposed, implemented and analyzed some algorithms, while supervising the use of the implementation on some biological data that were previously impossible to analyze. This effort has led to a new protocol for microbial communities analysis.

#### **Journal papers**

[J34] [J48] [J49] [J51]

#### **Software**

- ProbeSelection

#### **Main collaborations**

- University of California at Riverside (Tao Jiang, James Borneman)

### **Haplotyping**.....

Several species, including human beings, are diploid, that is each chromosome consists of two distinct copies called haplotypes. Current technological limitations do not allow to cheaply compute those haplotypes, but only genotypes (that is the two nucleotides that are in the same position in those haplotypes). Since it is important to know the actual haplotypes, a number of related computational problems have been recently introduced; those problems compose the field of haplotyping.

I have designed and analyzed an algorithm to complete haplotypes on incomplete data and under the coalescent model (such models forbids some otherwise possible recombinations), where the objective function is the entropy of the solution. The algorithm belongs to the class of Kernighan-Lin heuristics and it has been empirically and favorably compared to the greedy algorithm that was previously routinely employed.

Moreover I have studied the xor-genotyping problem, where the input data contains even less information than in most other haplotyping problems, as it is known only the positions where the two haplotypes differ, and not their contents. This formulation is a faithful model of the results that can be obtained with a recent and economically viable technology. I have

obtained some preliminary results on the computational complexity of the problem, and he has designed some efficient algorithms.

Another problem that I have studied is single-individual haplotype assembly, where we want to determine which variants a person actually has. My contribution on this problem has been the development of a software tool (HapChat) and the supervision of its experimental analysis. This approach has been incorporated into one of the most widely used tools for this problem. Most recently, we have extended this idea to viral data, where we want to quickly and effectively identify the viral strain in a sample. In this case I have supervised the entire work.

#### **Journal papers**

[J4] [J15] [J32] [J37] [J43] [J47]

#### **Software**

- HapChat <https://github.com/AlgoLab/HapCHAT>

### **Graph Algorithms**.....

Graph theory is one of the most important research fields that are common to Computer Science and Discrete Mathematics, as graphs are a mathematical device that is suitable for natural modeling of various real-world problems. One of the techniques that has been widely employed for designing efficient graph algorithm consists of decomposing the graph and then solving the problem on the smaller parts for finally recombining the partial solutions.

I have developed some efficient algorithms for computing the modular decomposition on hypergraphs and k-structures.

#### **Journal papers**

[J52]

### **Text Algorithms**.....

The sheer size of genomic data makes important to develop data structures to represent with a small amount of memory while querying them efficiently. The main such data structure is the Burrows-Wheeler Transform (BWT), and I have developed some algorithms to efficiently compute the BWT together with another auxiliary data structure (the Longest Common Prefix array) and for building the string graph (a data structure that represents how some strings overlap with each other). Moreover, I have supervised the implementation and the experimental analysis of those algorithms.

#### **Journal papers**

[J8] [J13] [J14] [J19] [J20] [J22]

#### **Software**

- bwt-lcp-parallel <https://github.com/AlgoLab/bwt-lcp-parallel>
- bwt-lcp-em <https://github.com/AlgoLab/bwt-lcp-em>
- FSG <https://github.com/AlgoLab/FastStringGraph>
- LSG <https://github.com/AlgoLab/LightStringGraph>

## **Research and Technology Transfer**

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### **Research and consulting contracts**.....

**2003–2004:** Consultant on the computational aspects of the project INTERREG IIIB (2000–2006) W.E.S.T. WOMEN EAST SMUGGLING TRAFFICKING (WP.2.2). **Proponent: Fondazione Ismu – Iniziative e Studi sulla Multietnicità.** I have been the only computer scientist involved in the project.

**2005:** Consultant on the computational aspects of the project “Indagine Finalizzata all’Analisi degli Effetti Prodotti dai Processi di Regularizzazione dei Lavoratori Extracomunitari, con Particolare Riferimento al Mercato del Lavoro e all’Integrazione Sociale nelle Regioni Ob. 1”, funded under Misura I.2 FESR “Adeguamento del Sistema di Controllo Tecnologico del Territorio”, PON Sicurezza 2000/2006. **Proponent: Fondazione Ismu – Iniziative e Studi sulla Multietnicità.** I have been the only computer scientist involved in the project.

## Publications

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### Journal articles.....

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- [J2] Jorge Avila Cartes, Santosh Anand, Simone Ciccolella, Paola Bonizzoni, and Gianluca Della Vedova. Accurate and fast clade assignment via deep learning and frequency chaos game representation. *GigaScience*, 12, 12 2022.
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- [J10] Simone Ciccolella, Giulia Bernardini, Luca Denti, Paola Bonizzoni, Marco Previtali, and Gianluca Della Vedova. Triplet-based similarity score for fully multilabeled trees with poly-occurring labels. *Bioinformatics*, 37(2):178–184, 2021.



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