Giorgio Valentini CURRICULUM VITAE

SYNTHETIC CURRICULUM

Degree in Biology and Computer Science, PhD in Computer Science (University of Genoa). Full Professor at the Department of Computer Science, University of Milan (UNIMI). National Qualification as full professor in 2017. Member of the UNIMI Doctoral School of Computer Science and scientific director for UNIMI of the European doctorate in Genomics and Bioinformatics in collaboration with the Joint Research Center of the European Union. Director of AnacletoLab, Computational Biology and Bioniformatics Laboratory of the Department of Computer Science of the University of Milano. Principal Investigator of 10 national and international research projects funded by public and private institutions in the area of Bioinformatics, Machine learning and Big Data analytics. In 2018 one of his papers has been awarded as one of the best the by the International Medical Informatics papers of year Association. His main research activity includes the development and application of Artificial Intelligence methods to bio-medical problems and Computational Biology, with a special focus on Machine Learning methods for Personalized and Precision Medicine, Network Medicine and Systems Biology, in the context of collaborations with various European and American research centers and universities. He is author of over 150 scientific publications with peer-review in journals, book chapters and international conferences in the field of Machine Learning, Bioinformatics and Computational Biology. He is member of CINI-AIIS, the Italian National Laboratory in Artificial Intelligence and Intelligent Systems.

ANALYTICAL CURRICULUM

Qualifications and courses of study

- National scientific qualification for the function of full professor sector 01 / B1 Informatica valid from 10/04/2017 to 10/04/2023 (article 16, paragraph 1, Law 240/10)
- PhD in Computer Science, University of Genoa (2003). Title: "Ensemble methods based on biasvariance analysis".
- Master Degree in Information Sciences, University of Genoa (1999), grade: 110/110 cum laude.
- MasterDegree in Biological Sciences, University of Genoa (1981), grade: 110/110 cum laude.
- Classical Maturity, Classical Lyceum of Savona (1977), grade: 56/60.

Actual position

Full Professor (scientific-disciplinary sector INF / 01) at the Department of Computer Science, University of Milan since 2019.

I teach didactic courses within the Computer Science teaching coordination board of the same University.

Previous positions

- Associate Professor at the Department of Computer Science, University of Milan (2010-2018)
- Permanent researcher at the Department of Computer Science, University of Milan (2005-2009)
- Post-doc researcher (research fellow) at the Department of Computer Science, University of Milan (2003-2004)
- PhD student at the University of Genoa (2000-2003)
- Contract researcher at the National Institute of Physics of Matter (1999), Genoa
- Lecturer in natural sciences, biology and chemistry and at several Italian high schools (1983-1998)

Research activity

My research activity is between Machine Learning and Bioinformatics, and is motivated by complex problems in Molecular Biology and Medicine. By modeling these problems in a Machine Learning context, I design and develop new machine learning algorithms or adopt suitable existing algorithms to address relevant problems in the field of Bioinformatics, with a particular interest, especially in recent years, in the development of Machine Learning methods for Medicine Genomics, and Precision and Personalized Medicine.

Although in my research activity the development of new Machine Learning methods is closely linked to real problems in the bio-medical field, I have also developed "pure" Machine Learning research lines, especially for the design and analysis of ensemble methods.

For this reason the general outline of my main research lines can be schematically articulated as follows:

I. Bioinformatics

A. Analysis, development and application of supervised Machine Learning methods to Bioinformatics

- A1. Machine Learning Methods for Personalized Medicine
- A2. Hierarchical ensemble methods for structured prediction in biological ontologies
- A3. Supervised ensemble methods for supporting bio-molecular diagnosis

B. Analysis development and application of semi-supervised Machine Learning methods to Bioinformatics

B1. Methods based on kernelised score functions for the analysis of complex biomolecular networks.

B2. Methods based on parameterized and cost-sensitive Hopfield networks for the prediction of gene function.

B3. Scalable methods for the analysis of big biomolecular networks.

C. Analysis development and application of unsupervised Machine Learning methods to Bioinformatics

C1. Stability analysis based methods for evaluating the reliability of clusters identified in complex bio-molecular data

- C2. Ensemble clustering methods for pattern research in bio-molecular data
- D. Methods for the integration of big biological and medical data.
 - D1. Algorithms for the massive combination of biomolecular networks
 - D2. Supervised ensemble methods for the integration of omics data
- II. Machine Learning
 - A. Analysis and design of ensemble methods
 - A1. Hyper-ensemble methods for unbalanced classification problems with big data
 - A2. Multiclass, multi-label and multi-path hierarchical ensemble methods
 - A3. Ensemble methods based on the decomposition of the error into bias and variance
 - A4. Supervised ensemble methods based on randomized projections
 - A5. Methods of ensemble with error correction codes for multiclass classification.
 - A6. Ensemble clustering methods

B. Design and implementation of machine learning software libraries

Description of the research lines

The research lines are summarized below with reference to the publications listed at the bottom of ththis document.

I. Bioinformatics

A. Analysis, development and application of supervised Machine Learning methods to Bioinformatics

A1. Machine Learning Methods for Personalized Medicine

The identification of genetic variants associated with human pathologies represents one of the fundamental challenges of "Personalized and Precision Medicine", and requires the development of a new generation of Machine Learning methods to select the rare potentially "deleterious" variants (i.e. causative or associated with the risk of disease) in the sea of "neutral" variants that represent the "physiological" genetic variability of each individual. To this end I developed hyperSMURF (hyper-ensemble of SMOTE under-sampled random forests), a new method that adopts learning strategies based on

resampling techniques and hyper-ensembling techniques to tackle the problem of the "imbalance" that characterize several types of genomic [R49, C77]. This Machine Learning method constitutes the "core" of Genomiser, a methodology and a software tool recently proposed in the context of an international collaboration, which uses both genotypic and phenotypic data to identify pathological variants causing Mendelian genetic diseases [R48].

HyperSMURF is a general method for the analysis of genetic variants and can be applied to the study of different pathologies, but its performance depends significantly on the tuning of learning parameters [C78]. For this reason we developed a highly parallel version of the algorithm for High Performance Computing architectures in the context of the HyperGeV project - Detection of Deleterious Genetic Variation through Hyper-ensemble Methods in order to fine-tune the parameters and analyze big data in the the context of relevant Genomic Medicine problems [paper submitted to GigaScience].

I am also developing imbalance-aware learning techniques based on mini-batch sizing and mini-batch imbalance-aware sampling techniques for predicting pathogenic genetic variants with deep neural networks [C84].

A2. Hierarchical ensemble methods for structured prediction in biological ontologies

Relevant concepts in the field of molecular biology (for example: the functions of genes and proteins) and medicine (abnormal phenotypes associated with human pathologies) are organized according to hierarchical ontologies structured as trees (eg: FunCat for the functional classification of genes) or as acyclic direct graphs (DAG) (eg: the Gene Ontology (GO) for the classification of genes and proteins and the HPO (Human Phenotype Ontology) for the classification of pathological human phenotypes.

In this context I developed hierarchical ensemble methods [R39] based on the true path rule (TPR) [R29, C44, C48, C51] and cost-sensitive Bayesian methods for probabilistic reconciliation of the output of the base learners [R25, C50, C53] and for the structured prediction of the function of genes and proteins in tree-structured ontologies. I have therefore shown that the combination of hierarchical ensemble methods, cost-sensitive learning strategies and the integration of different types of data significantly improve the performance in the prediction of gene functions at the level of the entire genome [R25, R32]. In the context of the prediction of abnormal human phenotypes according to the HPO, I have recently proposed new hierarchical ensemble methods for structured prediction based on DAG that have achieved state of the art results [R50, C76, C79, C70, C71]. Recently, new methods based on isotonic regression algorithms combined with the TPR algorithm led to state-of-the-art results in the prediction of protein function [paper in preparation].

A3. Supervised ensemble methods for supporting bio-molecular diagnosis

The bio-molecular classification of pathological phenotypes requires the development of methods wellsuited to the characteristics of the "omics" data used, often characterized by high dimensionality. In this context, various methods of supervised ensembles were explored, such as methods based on error correcting codes [R3, C11, C14], on the reduction of the dimensionality of data through randomized projections [R8, C22, C23], or methods of bagging and its variants [R5, C18, C19] and ither based on the analysis of data complexity [C37, C43].

Finally, I applied univariate feature selection and cost-sensitive SVM methods to the analysis of radiographic images (classification of pulmonary nodules) with results comparable with the best ones in the literature [R9, C25].

B. Analysis development and application of semi-supervised Machine Learning methods to Bioinformatics

In the field of Systems Biology and Network Medicine, I developed semi-supervised graph-based learning methods for the study of biological systems as complex entities, in which the biological functions derive fundamentally from the relationshipds between the parts that make up the system.

B1. Methods based on kernelised score functions for the analysis of complex biomolecular networks.

The problem of the association of genes or proteins or more generally of biomolecules with a specific biological property (eg: a biomolecular function, a diseases or a drug indication) can be modeled as a problem of node label ranking on a graph. Most of the methods proposed for graph analysis use local or global learning strategies to rank the nodes or predict the arcs of the graph itself. To integrate local and

global learning strategies, I proposed semi-supervised transduction methods that are able to learn both the global topology and the local characteristics of each node of the graph [R46] through the graph kernelization.

These methods have been successfully applied to the analysis of complex biomolecular networks for protein function prediction, to gene ranking problems (disease gene prioritization), and drug repositioning problems (search for new therapeutic indications of drugs originally designed for different therapeutic purposes) [R31, R33, R35, R38, R41, C61].

Finally, I recently proposed new semi-supervised network-based methods based on graph kernels which instead of analyzing the "biomarker space", as usually is the case in the context of Network Medicine, analyze the "patient space", through the construction of patient networks based on the similarity of their biomolecular profiles (eg: considering their expression or genetic profiles). The "patient networks" can be used both for diagnostic or prognostic purposes, to stratify the patients themselves into pathological subtypes, to identify biomarkers associated with specific pathologies or to predict the response to drugs [N21, paper submitted].

B2. Methods based on parameterized and cost-sensitive Hopfield networks for the prediction of gene function.

In the context of the prediction of the function of proteins and genes according to the GO, a significant problem is represented by the imbalance in the annotations. In fact for the majority of GO classes only a relatively small number of annotated genes (positive examples) is available. In this context, classical learning methods (including the Hopfield Networks) tend to always predict the majority class. For this reason we designed a new class of parameterized Hopfield Networks (COSNet) able to learn their own learning parameters from the data, explicitly taking into account the imbalance between positive and negative examples [R36, R44, C58].

A COSNet variant, designed to explicitly consider "categories" of neurons known a priori in the network has been successfully applied to the prediction of the function of proteins in a multi-species context [R45]. Finally, an "imbalance-aware" method of data integration coupled with the parametric Hopfield Networks has obtained state-of-the-art results for the prediction of protein functions [R43, C67]

B3. Scalable methods for the analysis of big biomolecular networks.

Big data analysis in large biomolecular networks represents a significant problem in computational biology. In this context, I developed vertex-centric algorithms and used technologies such as GraphChi based on the efficient use of secondary memory to process large graphs that cannot be directly loaded into primary memory. The goal is to analyze big data built with "omics" data, with relevant applications in Molecular Biology and Medicine, using simple stand-alone workstations. Promising experimental results have been obtained in the context of multi-species prediction of protein function [R42, C65, C74].

Another line of research concerns the use of GPU technology for the massively parallel implementation of node label prediction algorithms such as COSNet for efficient processing of large graphs [R52], with applications to the prediction of GO classes, using the integrated multi-species network of the STRING database [N20] which includes millions of proteins of different species. The construction of a network consisting of tens of millions of human SNPs (single nucleotide polimorphisms) for the search for potentially deleterious or pathological SNPs through the parallel version [R52] of COSNet is being developed.

C. Analysis development and application of unsupervised Machine Learning methods to Bioinformatics

C1. Stability analysis based methods for evaluating the reliability of clusters identified in complex biomolecular data

The validation of the clusters identified by clustering algorithms is a problem of great relevance in bioinformatics: genomics and proteomics present several problems in which it is fundamental to evaluate the reliability of the structures and patterns identified in complex biomolecular data.

The research activity was articulated in the development of algorithms for the analysis of the reliability and selection of the order of the model for unsupervised problems [R14, R15, R18, C29, C34, N10], and algorithms for the analysis of the reliability of the single clusters [R11, R12, C24], using a new approach

based on the analysis of the stability of the clusters obtained. We developed statistical tests based on the chi square distribution [R15, C29] and on the classical Bernstein inequality [R18, C34] for the search for multiple structures in complex data.

These methods were applied to the validation of pathological subclasses characterized at the biomolecular level and to the search for multiple structures in biomolecular data, using data generated by high-throughput bio-technologies [R12, R13, R15, R18, R22, C39, N9, N10].

C2. Ensemble clustering methods for pattern research in bio-molecular data

The search for bio-molecular patterns in data characterized by high dimensionality and low cardinality (for example: DNA microarray or spectrometric data related to proteins), led to the design and development of clustering ensemble methods specific for this type of data. In particular, unsupervised methods based on randomized projections have been developed to analyze data characterized by high dimensionality [C28]. These methods were subsequently applied to the analysis of gene expression data [C31]. We also developed clustering ensemble methods based on randomized projections that use a fuzzy approach both for the basic clustering constituting the ensemble, and for combining the clustering obtained on multiple instances of the data. From the initial algorithm [C33] a more general algorithmic scheme has been developed from which different fuzzy ensemble clustering algorithms can be derived [C36] and this approach has been applied to the analysis of gene expression data for the search of pathological subclasses characterized at bio-molecular level [R20].

D. Methods for the integration of big biological and medical data.

D1. Algorithms for the massive combination of biomolecular networks

The integration of biomolecular networks constructed with different types of "omics" data is a significant problem in the field of Systems Biology and Network Medicine.

My research activity has contributed to show that the integration with non-weighted and weighted methods of different biomolecular networks plays an essential role in identifying the genes associated with over 700 pathologies [R41]. I have also proposed several methodologies for the construction and integration of biomolecular networks, with applications to the prediction of protein function, and to the problems of disease gene prioritization and drug-target association [R33, R38, R43, C63, C66, C67, C75].

D2. Supervised ensemble methods for the integration of omics data

My research activity has shown that even relatively simple ensemble methods such as majority voting or decision templates can achieve results comparable with state of the art in "omics" data integration [R24, R26]. My other works have confirmed the effectiveness of ensemble methods for the integration of complex biomolecular data [C41, C45, C46, C47], also showing that the ensemble methods are also able to tolerate relatively high levels of noise in the data, without a significant deterioration in performance [R27]. I have also carried out studies on the application of ensemble methods for the integration of heterogeneous data and for the prediction of the subcellular localization of proteins [C54, C57], and on the use of XML for the integration of heterogeneous biomolecular data [R23, C28, C40].

II. Machine Learning

Although my research on new machine learning methods has been mostly motivated by complex problems in Molecular Biology and Medicine, I have also developed "pure" Machine Learning research lines, especially for the design and analysis of ensembles of learning machines.

A. Analysis and design of ensemble methods

A1. Hyper-ensemble methods for unbalanced classification problems with big data

Many relevant classification problems, not only in the field of Computational Biology and Genomic Medicine, are characterized by a strong imbalance of the examples between the classes. In this context

classical Machine Learning methods tend to be biased towards the majority class (negative class) and fail to learn the examples of the minority class (positive class). The hyperSMURF method, originally designed for strongly unbalanced genomic problems [R49] is general enough to be applied in other contexts characterized by a strong data imbalance. The algorithm, based on hyper-ensembling and data resampling techniques, has achieved state-of-the-art results in the field of Genomic Medicine [R48].

The parallelized version, parSMURF, is able to process big data, and further improves the performance of hyperSMURF, thanks to the automatic tuning of its learning parameters. The methods scales nicely with big data, and has been deployed in two flavours: a version for stand-alone multi-core architectures and another one for High performance computing clusters, depending on the level of complexity of the problem addressed [article submitted to GigaScience].

A2. Multiclass, multi-label and multi-path hierarchical ensemble methods

The problem of gene clssification has stimulated the research and development of multiclass classification algorithms (the functional classes of genes are of the order of hundreds or thousands), multi-label (a gene can belong to several classes) and multi-path (classes are structured according to trees or DAGs). The hierarchical ensemble algorithms that we developed for this type of problem [R25, R29, R32, R50] have a broader value and can be applied in other contexts characterized by hierarchical tree or DAG-structured taxonomies. These algorithms adopt a two-step learning strategy: at first the individual classes are learned in a "flat" manner by the base learners, and then the predictions of the trained models are combined according to the class hierarchy. Recently we developed an algorithm for DAG-structured taxonomies based on the integration of isotonic regression and bottom-up combination techniques (from the most specific classes to the most general classes) which guarantees the consistency of the predictions and systematically improves the predictions of the "flat" learning algorithms [article in preparation].

A3. Ensemble methods based on the decomposition of the error into bias and variance

In this line of research the decomposition of the error into bias and variance is used as a tool to analyze the property and characteristics of the learning algorithms.

Based on the Domingos theory, which generalizes to the 0/1 loss function the classical analysis based on the quadratic loss function, I analyzed the relationships between learning and decomposition of the error in bias and variance in the case of Support Vector Machines [R7].

The characterization of the learning processes of SVMs in terms of the decomposition of the error into bias and variance also offers a rationale for the development of new ensemble methods [R7, C16].

Taking advantage of the SVM bias-variance analysis, I proposed a new ensemble algorithm, called Lobag (Low Bias Bagging), which estimates SVM bias, selects SVMs with minor bias and then combines them through bootstrap-based aggregation mechanisms. This approach jointly reduces the bias and variance of the error, and can be interpreted as a "low-bias" variant of bagging [C20]. The method has been successfully applied to the classification of bio-molecular tumor diseases [C19]. The bias-variance error analysis was subsequently extended to resampling-based ensemble methods, showing the relationships and differences in the learning mechanisms of bagging methods, random aggregation and Lobag [R10, C21].

A4. Supervised ensemble methods based on randomized projections

In relation to the bio-molecular diagnosis of cancer, ensemble methods based on random subspaces have been developed, using SVM as a learner base [R8, C22]. An extension of the model, which includes a feature selection stage to eliminate the less relevant features for classification, followed by the application of the random subspace method on the remaining features, showed competitive results with state-of-the-art ensemble methods published in the literature [C23].

A5. Methods of ensemble with error correction codes for multiclass classification.

ECOC (Error Correcting Output Coding) ensemble methods allow to improve the reliability of the prediction for multiclass classification problems through the redundant coding of the class labels, realized with the decomposition of a multiclass problem into a series of dichotomous problems solved by a classifier ensemble.

In this context I analyzed the effectiveness of ECOC methods for multi-class problems in both ensembles of learning machines and single "monolithic" learning machines [R4, C2] and subsequently I experimentally analyzed the dependency between errors at the level of the single bits of the ECOC

codewords through measurements based on mutual information [R6], in order to compare different types of ECOC codings and different architectures of machine learning systems based on ECOC [C5, C7, C8]. In addition to applications in the bioinformatics field [R3, C11, C12], the ECOC ensembles (together with boosting methods) have been successfully applied also to multi-class problems with electronic noses [R1, C3, C6, C15].

A6. Ensemble clustering methods

Unsupervised ensemble methods based on randomized projections, motivated by clustering problems in spaces of high dimensionality and reduced cardinality that characterize different problems in the bioinformatics field, represent an unsupervised extension of the methods of supervised random subspace [C28].

In [C24] I showed how the random projections induced by the random subspace method can produce significant distortions in the gene expression data, while using randomized projections that obey the Johnson and Lindenstrauss lemma we can generate data of reduced dimensionality whose metric features are similar to those of the original space [R13]. In accordance with this analysis, we proposed clustering ensemble methods based on randomized projections [C31] that have been successfully applied to the analysis of DNA microarray data [C28].

A fuzzy extension of the ensemble method proposed in [C31] has been developed in [C36]: from the combination of different "crispization" mechanisms of basic fuzzy clustering and different types of fuzzy aggregation of basic clustering, we can derive an algorithmic scheme from which several fuzzy ensemble clustering methods can be obtained [C36]. These methods have been applied to the analysis of gene expression data [R20, C32, C33].

B. Design and implementation of machine learning software libraries

My research activity both in Machine Learning and in Bioinformatics has always been accompanied by the design and implementation of software libraries.

Several ensemble methods that I developed were made available and implemented in the C++ library, NEURObjects, initially conceived for implementing neural networks [R2, C1].

Later, in parallel with the increase in research activities in the bioinformatics field, I designed and implemented open source R libraries for the analysis of complex bio-molecular data. In particular the clusterv library [R11] allows to analyze the reliability of single clusters in high-dimensional bio-molecular data, the mosclust library [R14] allows to determine the "optimal" number of clusters and to identify multiple structures present in complex bio-molecular data, while the hcgene library [R17] allows to analyze the acyclic direct graphs of the Gene Ontology and the FunCat trees to support the functional classification of proteins. The RANKS library [R46] provides graph-based algorithms for node label ranking and classification problems. The COSNet library implements new models of parametric Hopfield networks [R44], while HEMDAG [R50] hierarchical ensemble methods and hyperSMURF [R49] "imbalance-aware" hyper-ensembling methods for the analysis of strongly unbalanced data, and its parallel extension (parSMURF) in C ++ is being released for the analysis of genomic big data [paper submitted to an international journal].

Indices of scientific productivity

The productivity indexes extracted on 7/4/2019 from the Scopus and Google Scholar database are presented below:

Index	Scopus	Google Scholar
Number of citations	1642	3261
H - index	23	32
i10 - index	50	75

Management or participation in the activities of a research group characterized by national or international collaborations

A) Direction of AnacletoLab, Laboratory of Computational Biology and Bioinformatics of the Department of Computer Science, University of Milan http://anacletolab.di.unimi.it/.

The laboratory brings together a group of professors / researchers, grant holders and PhD students and collaborates with different national and international research groups in the field of Computational Biology, Molecular Biology and Medicine, including:

a) Computational Biology Group of the Charite - University of Humboldt Universitat Berlin,

b) the Computational Biology group of the Berlin Institute of Health,

c) the Computer Science dept. of the Royal Holloway, University of London,

d) the Jackson Laboratory for Genomic Medicine, CT, USA,

e) The Artificial Intelligence department of the University of Granada, Spain

f) The Computer Science Dept of Aalto University, Helsiniki, Finland,

g) the Wellcome Trust Sanger Institute and the European Bioinformatics Institute (EBI) of Hinxton, UK, h) The Division of Environmental Genomics and Systems Biology, Lawrence Berkeley National Laboratory, Berkeley, CA, USA,

i) the European Center for Living Technologies of Venice,

l) Other national research groups both in the medical field (National Cancer Institute, and S. Raffaele Hospital of Milan, National Institute of Molecular Genetics) and computer science (University of Salerno and University of Cagliari).

(Period: from 2010 to today)

B) Participation as director of the AnacletoLab laboratory of UNIMI to the international challenge CAFA2 (Critical Assessment of Functional Annotation) within the Special Interest Group "Protein Function Prediction" of the ISCB (International Society of Computational Biology). The Special Interest Group brings together the main international research groups for the prediction of the function of proteins with computational methods (http://biofunctionprediction.org/).

In my position as head of the AnacletoLab research group, I participated in the research activity of the Special Interest Group "Protein Function Prediction" which led to the publication in Genome Biology (one of the main journal of computational biology) of a collective work involving the research activity of over 50 research groups from all over the world. In this context the laboratory was placed in the first 3 in the challenge CAFA2 for the prediction of human genes associated with pathological phenotypes. AnacletoLab participated in the new challenge CAFA3 for the function prediction of human proteins and of the main model organisms and in the prediction of genes associated with abnormal human phenotypes. The preliminary results of this challenge (the evaluation is still in progress) place AnacletoLab again among the "top ranked" groups at the international level for the prediction of the function of proteins. (Period: from 2013 to today).

C) Participation in the activity of an international research group for the search of mutations associated with genetic diseases in non-coding regions of the genome.

My contribution consists in the development of machine learning methods specific to this problem. The international group includes several research groups from Europe, North America and Australia, including Queen Mary University of London; Genomics England, UK; the Wellcome Trust Sanger Institute, Hinxton, UK; the Max Planck Institute for Molecular Genetics, Berlin; the Department of Biomedical Informatics and Intelligent Systems Program, University of Pittsburgh; the Division of Environmental Genomics and Systems Biology, Lawrence Berkeley National Laboratory, Berkeley, CA; the Department of Medical Informatics and Clinical Epidemiology, Oregon Health & Science University, Portland, USA; the Kinghorn Center for Clinical Genomics, Garvan Institute of Medical Research, Darlinghurst, Australia; the Charité-Universitätsmedizin

Berlin.

The computational methodology developed for the search of mutations associated with Mendelian genetic diseases in non-coding regions of the genome has been published in the American Journal of Human Genetics, a leading journal in this field. This method, called Genomiser, represents the state of the art at international level for the search for causative mutations of Mendelian genetic diseases. The Machine Learning method, developed specifically by me in collaboration with the Computational Biology group of the Charitè UniversitatzMedizin in Berlin, and which constitutes the core of Genomiser, was recently published in a journal of the Nature Publishing group. This method is general enough to be used in other contexts to search for genetic variants associated with genetic and tumor diseases, and in fact a collaboration is underway with the School of Medicine of the University of Utah to study the hereditary factors of various tumors. (Period: 2014 to present)

D) Collaboration in a joint project with the Berlin Institute of Health (BIH, which brings together research groups from the von Humboldt and the Freie Universität in Berlin and from the Max Delbrück Center for Molecular Medicine) for the systematic study of genetic variations in regulatory regions of the human genome and of their impact on genetic diseases and cancer. This project requires the development of innovative Artificial Intelligence methodologies for the analysis of data generated by new biotechnologies, such as Massive Parallel Reporter Assay (MPRA) for the study of the functional in vivo effect of genetic variants, and for the analysis and massive integration of epigenomic data recently made available by the International Human Epigenome Consortium (IHEC). The goal is to decode the "genetic code" at the base of gene regulation in the non-coding regions of the genome, and to study its alterations to predict the pathological effect of genetic mutations underlying cancer and genetic diseases.

The project is currently funded by the German DAAD and the MIUR, but the complexity of the problem requires collaboration with other European and American scientific partners and relevant funding for the generation and bioinformatic processing of biotechnological data. For this reason we are working together with the BIH and other European and American partners for the preparation of a European research or a NIH funded project.

(Period: 2017 to present)

E) Collaboration with the Jackson Lab for Genomic Medicine (CT, USA) and with the Wellcome Sanger Institute - Hinxton (Cambridge, UK) for the search of splicing mutations in Exonic Sequence Enhancers (ESE, that promote exon splicing) in the human genome. These mutations, which are extremely difficult to detect because they are apparently "silent", are the basis of various tumors. The idea we are working on is to build predictors based on artificial intelligence techniques capable of identifying these mutations on the basis of a set of appropriate genomic and epigenomic features and based on ESE manually labeled through the analysis of the medical literature. After the publication of the first results (which appear very promising) we will prepare an application for a grant from the NIH (National Institute of Health of the United States) to finance this line of research. (Period: 2018 to present)

F) Research collaboration with the Urology Department (URI) of the S. Raffaele Hospital (OSR) in Milan, regulated by a Confidentiality Agreement signed between UNIMI and OSR, for the project "Male infertility as a proxy of the progerois syndrome", of which I am the scientific director for UNIMI (Andrea Salonia, head of the URI unit is the scientific director for OSR). The purpose of this agreement is to start-up joint research activities for prparing a grant to the NIH. (Period: 2019 to present)

G) In the a.y. 2015/16 I enjoyed a sabbatical year during which I carried out research at the European Center for Living Technologies of Venice (Prof. Pelillo's research group), at the Charitè, Faculty of Medicine of the von Humboldt University of Berlin (research group of Computational Biology of prof. Robinson), and the Artificial Intelligence Department of University of Granada (prof. Blanco).

Scientific responsibility (Principal Investigator - PI) of international and national research projects, eligible for funding on the basis of competitive calls for proposals involving peer review

- Scientific coordinator for UNIMI of the EU Collaborative Doctoral Partnership in Genomics and Bioinformatics, funded by the European Commission in collaboration with the EU Joint Research Center (2018-2022). Our University has been selected among the top 5 European Universities for the area of Genomics and Bioinformatics. The five-year collaboration contract is renewable at the end of the five-year period, subject to agreement between the parties, and is aimed at training researchers in the field of Genomics and Bioinformatics, able both to carry out innovative research in these scientific disciplines and to scientifically support the European Commission in making decesions about European policies for Medical Genomics and Data Analytics in Health-care.
- PI of the project "Developing machine learning methods for the prioritization of regulatory variants in human disease" in collaboration with the Berlin Institute of Health, funded by MIUR and DAAD (Germany) (2018-2019)
- PI of "HyperGeV: Detection of Deleterious Genetic Variation through Hyper-ensemble Methods" (2016-2018), funded by CINECA and the Lombardy Region
- PI of "HPC-SoMuC: Development of Innovative HPC Methods for the Detection of Somatic Mutations in Cancer" (2017-2018), funded by CINECA and the Lombardy Region.
- Head of the UNIMI unit of the project "A composite predictive model of response to Fingolimod: integration of clinics, neuroimaging and genomics", funded by the Italian Multiple Sclerosis Foundation. The project is leaded by the Genetics Laboratory of the Complex Neurological Diseases of the S. Raffaele Hospital in Milan. The genetic biomarkers associated with the response to the Fingolimod drug, identified by the data of S. Raffaele and the predictive model developed by the UNIMI unit are currently being patented (2016-2018).
- Head of the UNIMI unit (via subcontracting) of the Finding-MS project within the European project ERA-PerMed Joint Transnational Call (JTC) 2018 on "Research projects on Personalized Medicine smart combinations for pre-clinical and clinical research with data and ICT solutions ". The project is in collaboration with the S.Raffaele Hospital of Milan, the CNR - Institute for Bio-medical Technologies, the Center Hospitalier Universitaire de Toulouse and geneXplain, a German bioinformatics company (2019-2022)

Participation in international and national research projects, admitted to the financing on the basis of competitive calls for peer review

- PRIN Project "Multicriteria Data Structures and Algorithms: from compressed to learned indexes, and beyond", coordinated by the University of Pisa (2019-2021)
- "Discovering Patterns in Multi-Dimensional Data" project (2016-2017) funded by the University of Milan.
- Participation in the Milan unit of the Network of Excellence "Pattern Analysis, Statistical Modeling and Computational Learning 2 (PASCAL2)", 7th European Framework Program financed by the European Union (2007-2013)
- Participation in the Milan unit of the Network of excellence PASCAL, 6th European Framework Program (2004-2006)
- Project "Computational methods for bio-medical pattern analysis" funded by UNIMI (2011-2013)
- MIUR project COFIN-PRIN Automata and formal languages: mathematical and applicative aspects (2010-11);
- PUR 2009 Project: "Automatic methods for pattern analysis in the biomedical field" funded by the University of Milan;
- PUR 2008 Project: "Innovative computational models" funded by the University of Milan; IEIIT-CNR 2007 Project: Machine Learning Techniques for Modeling and Growing Up " (2007-2008);
- PUR 2006 Project: "Stochastic and quantum models for computational and di problems bioinformatics " funded by the University of Milan;
- PUR 2006 Project: "Identification of transcriptional profiles in Acute Myeloid Leukemia by microarray on hematopoietic stem fractions " funded by the University of Milan;

- MIUR Project COFIN-PRIN Formal Languages and automata: methods, models and applications (2003-2005)
- MIUR Project COFIN-PRIN Machine Learning Techniques for Bioinformatics: Analysis and Modeling of Functional and Structural Data of Gene Expression (2001-2002)
- From Bits to Information: Statistical Learning Technologies for Digital Information Management Search USA, Project funded by the National Scientific Foundation (NSF) (2002).

Participation in editorial committees of international journals

- Member of the Editorial board of Scientific Reports, Nature (IF 2017: 4.122), indexed in Scopus and ISI Web of Science (from 1 June 2018).
- Member of the editorial board of Advances in Bioinformatics, Hindawi (IF 2017: 1.8), indexed in Scopus (from 2014)
- Guest Editor of Artificial Intelligence in Medicine, Elsevier (IF: 2.879) indexed in Scopus and ISI Web of Science for the Special issue "Computational Intelligence and Machine Learning in Bioinformatics" (2008-2009).
- I was also a member of the Editorial board of other minor journals in the Bioinformatics and Machine Learning area from 2008 to 2017.

Reviewer activity for international journals

I have worked as reviewer for the main international journals in Bioinformatics and Machine Learning, including:

- Journal of Machine Learning Research
- Machine Learning
- IEEE Transactions on Pattern Analysis and Machine Intelligence
- Pattern Recognition
- IEEE Transactions on Neural Networks and Learning Systems
- IEEE Transactions on Knowledge and Data Engineering
- Artificial Intelligence
- Computational Intelligence,
- Neurocomputing
- Neural Networks
- IEEE Transactions on Cybernetics,
- GigaScience
- Bioinformatics
- PLoS Computational Biology
- BMC Bioinformatics
- Briefings in Bioinformatics
- IEEE ACM Transactions on Computational Biology and Bioinformatics
- Artificial Intelligence in Medicine
- Journal of Bioinformatics and Computational Biology
- PLoS One

Conference and Workshop Organization

- Co-organizer of the workshop Soft computing methods for characterizing diseases from omics data - CIBB 2019, Bergamo 3-4 settembre 2019
- Chair of CIBB 2018 Computational Intelligence methods for Bioinformatics and Biostatistics, Lisboa (Portogallo) (Caparica 6-8 Settembre 2018)
- Chair of the Workshop BigTargets, European Conference on Machine Learning (ECML), Porto (Portogallo), 2015;

- Chair of the Fourth Italian Workshop on Machine Learning and Data Mining (MLDM 2015) XIV Conference of AIIA Pisa, 2015;
- Chair of SUEMA 2010, Third International Workshop on Supervised and Unsupervised Ensemble Methods and Their Applications at ECML European Conference on Machine Learning 2010, Barcelona, Spagna;
- Wokshop Learning from Multiple Sources with Applications to Robotics at NIPS 2009, Whistler, Canada, 2009;
- Chair of SUEMA 2008, European Conference on Artifical intelligence (ECAI) 2008 Patras, Grecia;
- Chair of CIBB 2007, Fourth International Conference on Computational Intelligence Methods for Bioinformatics and Biostatistics, Portofino, Italia
- Chair of SUEMA 2007, International Workshop on Supervised and Unsupervised Ensemble Methods and Their Application (nell'ambito di IbPRIA2007) a Girona (Spagna).

Participation in the scientific committee of international conferences.

Member of the Scientific Committee Program of over 50 international conferences within the scope of the machine learning and computational biology from 2007 to present, including:

- European Conference in Computational Biology
- European Conference on Machine Learning and Principles and Practice of Knowledge Discovery in Databases
- International Joint Conference on Artificial Intelligence (IJCAI) Machine Learning Track
- International Symposium on Foundations and Applications of Big Data Analytics
- Automatic Function Prediction Critical Assessment of Functional Annotation experiment (nell'ambito di ISMB Intelligent Systems for Molecular Biology)
- ICANN International Conference on Artificial Neural Networks
- SIAM International Conference on Data Mining
- ACM SIGKDD International Conference on Knowledge Discovery and Data Mining
- S+SSPR IAPR Joint International Workshops on Statistical + Structural and Syntactic Pattern Recognition
- International Conference on Pattern Recognition (ICPR)
- International Conference on Multiple Classifier Systems (MCS)

Invited speaker at international and national congresses and conferences

- Invited talk at the Statistical Workshop "Statistical approaches and validation in clustering: mixture models and nonparametric methods ", organized by the Department of Statistics of the University of Caen (France), entitled "Stability-based methods for cluster validation". (29-06-2007)
- Invited talk at CIBB 2008, FIFTH INTERNATIONAL MEETING ON COMPUTATIONAL INTELLIGENCE METHODS FOR BIOINFORMATICS AND BIOSTATISTICS IIASS - Vietri sul Mare, Salerno (Italy) entitled: "Unsupervised stability-based ensembles to discover reliable structures in complex bio-molecular data ". (04-10-2008)
- Invited talk at the Third Italian Workshop on Machine Learning and Data Mining XIII AI * IA Symposium on Artificial Intelligence, Pisa December 2014, "Analysis of bio-molecular networks through semi-supervised graph-based learning methods". (10-12-2014)
- Invited talk at Fifth Italian Workshop on Machine Learning and Data Mining XV AI * IA Symposium on Artificial Intelligence, Genoa, November 2016, "A hyper-ensemble approach for the genomewide prediction of disease and trait-associated genetic variants". (28-11-2016)

- Invited speaker at the Interdisciplinary Aspects of Biomolecular Modeling Workshop "Machine Learning approaches for Modeling Complex Biomolecular Systems" (26-6-2019)
- Invited talk at the Annual Workshop in Bioinformatics and Genomics of the Catalan Society of Biology and the Bioinformatics Barcelona association, Barcelona, Spain "Machine Learning for Computational Biology and Precision Medicine" (17-12-2019)

Invited speaker at Italian and European universities and research centers.

I held several seminars on Machine Learning and Bioinformatics topics in various Italian and foreign Universities and research centers since 2005, including:

- Department of Computer Science della Royal Holloway, University of London;
- Department of Computer Science, Aalto University, Helsinki;
- Department of Computer Science, Aristotle University of Thessaloniki;
- Charitè Universitatzmedizin Berlin
- European Center for Living Technologies, Venezia;
- Departamento de Sistemas Informáticos y Computación Universitat Politècnica de València;
- Computational Genomics Department del Centro de Investigacion Prencipe Felipe, Valencia;
- School in Bioinformatics, University of Brno (Czech Republic);
- Computer Science and Artificial Intelligence Department, University of Granada;
- Dipartimento di Informatica, Università di Pisa;
- Dipartimento di Matematica e Informatica dell' Università di Palermo;
- Dipartimento di Ingegneria ed Elettronica (DIEE) Universita' di Cagliari;
- IIASS Istituto Internazionale Alti Studi Scientifici "E. R. Caianiello di Vietri (Salerno);
- Dipartimento di Informatica dell' Università degli Studi di Salerno;
- IST Istituto Nazionale per la Ricerca sul Cancro, Genova.
- INT Istituto Nazionale Tumori, Milano
- Dipartimento di Biologia e Genetica per le Scienze Mediche, Università degli Studi di Milano

International awards.

- The paper M. Notaro, M. Schubach, P.N. Robinson, G. Valentini. Prediction of Human Phenotype BMT Bioinformatics, vol. 18 (1), 2017 has been awarded as one of the 5 best articles of the year by the International Medical Informatics Association (IMIA) for the "Knowledge Representation and Management" section.

Patents

A signature of 21 SNPs (Singe Nucleotide Polymorphism) for the prediction of the response of patients suffering from multiple sclerosis to the Fingolimod drug is being patented (UIBM -Ufficio Italiano Brevetti Marchi, number 102019000023154).

This signature has been carried out using an algorithm and a software I developed and applied to the analysis of genomic data provided by the Department of Neurology of the San Raffaele Hospital in Milan.

Evaluation activities in the context of national and international competitive selection procedures.

- Research expert for the European Commission (number: EX2014D182522) for the evaluation of proposals, projects and programs submitted for evaluation for European funding (from 2014)
- Auditor registered with REPRISE (register of scientific experts established at the MIUR) for the "basic research" section: Scientific-disciplinary sectors: Informatics (INF / 01) Sectors ERC: Computer Science and Informatics: Informatics and Information Systems, computer science, scientific computing and intelligent systems (PE6), Bioinformatics, biocomputing, and DNA and

molecular computation (PE6_13), Diagnostic tools (eg genetic, imaging) (LS7_2), Computational biology (LS2_11) (from 2018).

- Reviewer for proposals submitted to the Netherlands Organization for Scientific Research (NWO):
 Reviewer of the project "Collaborative experimentation for data mining" (2011-2012)
 - Reviewer of the project "Context-aware protein function prediction" (2012/13)
- Reviewer for proposals submitted to the Research Foundation Flanders (Fonds Wetenschappelijk Onderzoek Vlaanderen, FWO):

- Reviewer of the project "COCO-MULT: Constrained Constructive Machine Learning for Multiple Targets" (2018)

- Reviewer for SIR (Scientific Independence of young Researchers) projects funded by MIUR (2014/15)
- Reviewer for projects of the "Bloodwise" program for the study of hematological tumors, funded by 14M Genomics, a company for the diagnostic spin-out of the Wellcome Trust Sanger Institute of Hinxton, UK. (from 2012 to 2016)

Formal assignment of teaching or research fellowships from qualified foreign or supranational universities and research institutes

- Research fellowship Computer Science Dept. Oregon State University USA: development of ensemble methods based on bias-variance analysis. (from 01-03-2001 to 04-04-2001)
- Visiting researcher Principe Felipe Research Center, Valencia, Spain (2009) (from 09/14/2009 to 10/15/2009)
- Visiting professor at the Computer Science Department of the Aristotle University of Thessaloniki for common research activities in the area of machine learning methods for multi-target prediction problems characterized by high dimensionality both in the input space and in the output space.

(from 29-09-2014 to 10-28-2014)

- Visiting professor at the Computer Science Department of Royal Holloway, University of London for common research activities (development of methods for the integration and analysis of complex biomolecular networks) and for holding seminars to support the teaching of the master and PhD in Computer Science at Royal Holloway. (March 2015)
- Fellowship for visiting professors at the Computer Science Department of Aalto Univeristy (Helsinki) as part of the "Machine Learning for Metagenomics" project, funded by the Aalto Science Foundation of Helsinki (AScI Visiting Fellow Program 2014-15). (from 25-05-2015 to 25-06-2015)
- Visiting professor at the European Center for Living Technologies (ECLT). During the visit we strengthened joint research activities in the field of game theory and semi-supervised network-based methods for the analysis of bio-molecular networks. (from 01-02-2016 to 02/28/2016)
- Fellowship for visiting professors at the Computational Biology group of the Charité -Universitätsmedizin Berlin within the Funding program "Research Stays for University Academics and Scientists" funded by the DAAD (German Academic Exchange Service) for the project "Hierarchical ensemble methods for structured predictions in Biological Ontologies ". (from 01-03-2016 to 06/01/2016)
- Visiting professor at the Departamento de Ciencias de la Computación and Inteligencia Artificial de la Universidad de Granada. During the visit, common research activities were developed for the development of graph-based algorithms and their application to ranking problems of genes associated with specific pathologies, and to "repositioning" problems (alternative therapeutic use of drugs).

(from 07-06-2016 to 06-28-2016)

Responsibility for studies and scientific research entrusted by public or private institutions

- Research assignment for the development and modeling of machine learning methods with applications in computational biology, as a member of the CNR affiliated project "Machine Learning Techniques for Modeling and Growing Up ", entrusted by IEIIT-CNR (2007-2008)
- Consultancy contract with the company SIGEDA of Milan for the development of ensemble methods based on error correction codes for the multi-class classification of data coming from an artificial olfactory system (electronic nose) (2007)
- Responsibility of a research project for the prediction with machine learning methods of quantitative tumor phenotypes in model organisms and for the study of susceptibility to pulmonary and hepatic carcinogenesis, entrusted by the research unit of Genetic Epidemiology and Pharmacogenomics of the National Cancer Institute (INT) of Milan. (from 01-04-2015 to 1-10-2017)
- Commission of the Centro S.Raffaele Foundation of Milan for the development of a predictive model of response to treatment with II level drugs for patients with Multiple Sclerosis. (from June 2016 to June 2018)

Design and development of software libraries publicly available to the scientific community.

- HEMDAG: Hierarchical Ensemble Methods for Directed Acyclic Graphs https://cran.rproject.org/web/packages/HEMDAG
- HyperSMURF: Hyper-Ensemble Smote Undersampled Random Forests (software library for supervised prediction with highly imbalanced big data). https://cran.r-project.org/web/packages/hyperSMURF
- RANKS: Ranking of Nodes with Kernelized Score Functions (software library for node label learning in graphs).
- https://cran.r-project.org/web/packages/RANKS
- COSNET: Cost Sensitive Network for node label prediction on graphs with highly unbalanced labels.

http://www.bioconductor.org/packages/devel/bioc/html/COSNet.html;

- HCGene: an R package to support the hierarchical classification of genes. https://homes.di.unimi.it/valentini/SW/hcgene;
- Clusterv: an R package for cluster validation. https://homes.di.unimi.it/valentini/SW/clusterv;
- Mosclust: an R package for the discovery of significant structures in bio-molecular data. https://homes.di.unimi.it/valentini/SW/mosclust;
- NEURObjects a set of C++ library classes for neural networks development. http://homes.di.unimi.it/valentini/SW/NEURObjects;
- PerfMeas: an R package implementing different performance measure for classification and ranking tasks.
 - http://cran.r-project.org/web/packages/PerfMeas;
- NetPreProc an R package that implements preprocessing and normalization methods for network-structured data.
 - http://cran.r-project.org/web/packages/NetPreProc;
- Bionetdata an R data package that includes several examples of chemical and biological data networks.

http://cran.r-project.org/web/packages/bionetdata.

Participation in national and international scientific societies and associations

I am a member of ISCB - International Society of Computational Biology, BITS - Italian Society of Bioinformatics, INNS (International Neural Network Society), Data Mining and Big Data Analytics Technique Committee (DMTC) - IEEE Computational Intelligence Society (IEEE-CIS), of the CINI AIIS laboratory, and of the scientific committee of the working group on Machine Learning and Data Mining, within AI * IA, Italian Association for Artificial Intelligence.

Academic teaching

- My academic teaching activity is articulated both through the institutional courses, held at the University of Milan and through the doctorate courses, masters and courses held at the University of Milan and in other universities especially in the field of Bioinformatics and Machine Learning. In particular, I held courses for the three-year and master's degree in Computer Science, for the degree in Digital Communication, for the Master Degree in Molecular Biotechnology and Bioinformatics and for the degree in Industrial and Environmental Biotechnology, as well as basic Computer science courses for various three-year degrees in Medicine. I have also held and still maintain courses in English.
- I am currently also responsible for the General Computer Science course for the degree in Science and Technology for the study and conservation of cultural heritage and information media (the Computer Science Department is a department associated with this degree course).
- I am member of the teaching board of the PhD in Computer Science of the University of Milan since 2008. In this context I am responsible for the Machine Learning for Genomic Medicine course.
- From the academic year 2018/19 I am a member of the coordinating committee for the Department of Computer Science of the Master in Bioinformatics and Functional Genomics of the University of Milan, carried out in collaboration with the National Institute of Molecular Genetics (INGM) and the Policlinico 'University of Milan. In the academic year. 2017/18 I already held some lectures (Machine Learning for Personalized and Precision Medicine) within the same master.
- I am the scientific director for UNIMI of the EU Collaborative Doctoral Partnership in Genomics and Bioinformatics, funded by the European Commission in collaboration with the Joint Research Center (JRC) of the EU. The five-year plan of the collaborative doctorate between UNIMI and JRC is aimed at training researchers in the field of Genomics and Bioinformatics able both to carry out innovative research in this field and to scientifically support the political decisions of the Commission for Medical Genomics and Data Analytics in Health-care.

Academic year	course	degree	Liv.
18/19	Machine Learning for Genomic Medicine	Ph.D. course in Computer Science	D
18/19	Bioinformatica	Informatica	М
18/19	Informatica Generale	Scienze e tecnologie per lo studio e la conservazione dei beni culturali e dei supporti dell'informazione	Т
18/19	Machine Learning for Personalized and Precision Medicine	master in Bioinformatics and Functional Genomics UNIMI	Master
17/18	Bioinformatica	Informatica	М
17/18	Informazione Multimediale	Comunicazione Digitale	Т
17/18	Informatica Generale	Scienze e tecnologie per lo studio e la conservazione dei beni culturali e dei supporti dell'informazione	Т
16/17	Bioinformatica	Informatica	М
16/17	Bioinformatics Methods (in inglese)	Molecular Biotechnologies and Bioinformatics	М
16/17	Bioinformatica e Biostatistica	Biotecnologie Industriali ed Ambientali	Т
14/15	Bioinformatica	Informatica	М
14/15	Metodi Bioinformatici	Biotecnologie Biomolecolari e Bioinformatica	М
14/15	Informatica Avanzata	Biotecnologie Industriali ed Ambientali	Т
13/14	Bioinformatica	Informatica	М
13/14	Metodi Bioinformatici	Biotecnologie Biomolecolari e Bioinformatica	М

The following table summarizes the teaching assignments carried out in UNIMI starting from the a.a. 2004/05 to date:

13/14	Informatica Avanzata	Biotecnologie Industriali ed Ambientali	Т
12/13	Bioinformatica	Informatica	М
12/13	Metodi Bioinformatici	Biotecnologie Biomolecolari e Bioinformatica	М
12/13	Informatica Avanzata	Biotecnologie Industriali ed Ambientali	Т
11/12	Bioinformatica	Informatica	М
11/12	Metodi Bioinformatici	Biotecnologie Biomolecolari e Bioinformatica	М
11/12	Informatica Avanzata	Biotecnologie Industriali ed Ambientali	Т
10/11	Bioinformatica	Informatica	М
10/11	Metodi Bioinformatici	Biotecnologie Biomolecolari e Bioinformatica	М
10/11	Informatica Avanzata	Biotecnologie Industriali ed Ambientali	Т
09/10	Bioinformatica	Informatica	М
09/10	Metodi Bioinformatici	Biotecnologie Biomolecolari e Bioinformatica	М
08/09	Linguaggi di programmazione per la bioinformatica	Genomica Funzionale e Bioinformatica	М
08/09	Bioinformatica	Informatica	М
08/09	Informatica applicata ai processi biotecnologici	Biotecnologie Industriale e Ambientali	Т
07/08	Linguaggi di programmazione per la bioinformatica	Genomica Funzionale e Bioinformatica	М
07/08	Bioinformatica	Informatica	М
07/08	Informatica	Fisioterapia e Dietistica	Т
06/07	Linguaggi di programmazione per la bioinformatica	Genomica Funzionale e Bioinformatica	М
06/07	Bioinformatica	Informatica	М
06/07	Informatica	Podologia ed Igiene Dentale	Т
05/06	Linguaggi di programmazione per la bioinformatica	Genomica Funzionale e Bioinformatica	М
05/06	Bioinformatica	Informatica	М
05/06	Informatica	Podologia ed Igiene Dentale	Т
04/05	Linguaggi di programmazione per la bioinformatica	Genomica Funzionale e Bioinformatica	М
04/05	Algoritmi per la Bioinformatica	Genomica Funzionale e Bioinformatica	М
04/05	Laboratorio di Bioinformatica	Biologia	Т

M : "laurea magistrale" degree; T : "laurea" degree; D : Ph.D. course

• Previously I held the Functional Bioinformatics 1 course for the degree in Physics of the University of Genoa in the academic year 2002/03 and 2004/04 and the Neural Networks 1 exercises and the System Programming Laboratory (a.a 2001/02) for the degree course in Computer Science of the University of Genoa.

• I obtaiend teaching assignments for doctorate courses in various universities, including the University of Palermo (PhD in Computer Science and Mathematics), the University of Salerno (PhD in Computer Science), the University of Brno (Czech Republic) (Summer Doctoral School in Bioinformatics), the Computer Science Department of Royal Holloway, University of London.

• I was a lecturer and coordinator of courses on the R language for the analysis of biomedical data for Ph.D. students, research fellows and researchers in various research centers, including: "An R course for Oncology Bioinformatics", CINECA - Casalecchio di Reno, June 2008
"Course on the R language for the analysis of DNA microarray data with machine learning methods", IST Genova, November 2009

- "Analysis of biomolecular data with R", Parco Tecnologico di Lodi, 21-23 January 2009

I held several seminars and Ph.D. courses in various Italian and European universities, including

- "Discovering significant structures in bio-molecular data" Ph.D. course 3rd International Summer School on Computational Biology, Mikulov, Czech Republic, 13 -15 August 2007
- "Ensemble methods in bioinformatics" Ph.D. course il Dottorato in Matematica ed Informatica all'Università di Palermo (22-26 giugno 2008)
- "Supervised Gene Set Analysis for the molecular characterization of patients: a machine learning approach", seminar at Centro de Investigacion Principe Felipe di Valencia, 6-10-2009
- "True Path Rule hierarchical ensembles for genome-wide gene function prediction: a machine learning approach", seminar dor the Ph.D. course in Computer Science de l'Universitat Politecnica de Valencia, 13-10-2009
- "Hierachical ensemble methods for gene function prediction" seminar for the Ph.D. course in Information Engineering, Dipartimento di Ingegneria Elettrica ed Elettronica (DIEE) dell'Università di Cagliari (May 2010)
- "True Path Rule and H-Bayes hierarchical cost-sensitive ensembles for gene function prediction" held at 'Istituto Internazionale per gli Alti Studi Scientifici "E.R. Caianiello", Vietri sul mare (Salerno), 27 May 2010
- "True Path Rule hierarchical ensemble methods for gene function inference" seminar for the Ph.D. course in Copmuter Science, Dipartimento di Informatica, Università di Salerno, 6 june 2011
- "Stability-based methods for the assessment of clusters discovered in bio-molecular data" lecture for the Ph.D. course in Matematica ed Informatica all'Università di Palermo, june 2012
- "Ensemble methods for multi-target classification and regression", seminar at Informatics department Aristotle University of Thessaloniki, Grecia, 14 october 2014
- "Scalable methods for bio-molecular network analysis and for structured prediction in biological ontologies" seminar at the 'European Center for Living technologies Univ. Ca' Foscari, Venezia, 27 febbraio 2015
- "Analysis of bio-molecular networks through semi-supervised graph-based learning methods" seminar for the Ph.D. course in Computer Science, Computer Science Dept. Royal Holloway, University of London, 9 March 2015
- "Multi-label hierarchical prediction methods and their application to the automatic function prediction of proteins" seminario per il dottorato in Computer Science, Computer Science Dept. Royal Holloway, University of London, 10 March 2015
- "Scalable methods for the analysis of complex biomolecular networks", seminar at Dipartimento di Informatica, Università di Salerno il 5 Maggio 2015
- "Semi-supervised graph-based learning methods for the analysis of bio-molecular networks" seminar for the Ph.D. course in Computer Science, Aalto University, Helsinki, Finland, 13 June 2015
- "Semi-supervised graph-kernel methods for disease gene prioritization", seminar at Departamento de Ciencias de la Computación e Inteligencia Artificial de la Universidad de Granada, Spagna, 15-6-2016
- "A Machine Learning and High Performance Computing tool to prioritize pathogenic variants in the human genome", seminar for the Ph.D. course in Computer Science, Computer Science Dept. Royal Holloway, University of London, 26 Febbraio 2019

I was a member of various commissions for the final doctoral exam, and doctoral thesis reviewer for various Italian and European universities:

- Member of the final examination committee for the Doctorate in Computer Science of the University of Salerno (April 2010)
- Member of the final examination committee for the PhD in Information Engineering of the Department of Electrical and Electronic Engineering (DIEE) of the University of Cagliari (May 2010).
- Member of the final examination committee for the Doctorate in Computer Science of the University of Salerno (February 2011)
- Member of the final examination committee for the doctorate in computer science of the DIBRIS University of Genoa (May 2018)
- Member of the final exam for the PhD in Computer Science of the Freie Universitat Berlin (September 2018)
- Member of the final exam for the PhD in Computer Science of Royal Holloway University of London (February 2019)
- Reviewer of the doctoral thesis of Isabel Segura Bedmar (Computer Science Department, University Carlos III of Madrid) "Application of Information Extraction techniques to pharmacological domain: Extracting drug-drug interactions" (2010)
- Reviewer of the doctoral thesis of Francesco Iorio (Doctorate in Computer Science University of Salerno) "Automatic Discovery of Drug Mode of Action and Drug Repositioning from Gene Expression Data" (2010)
- Reviewer of the doctoral thesis of Luca Pinello (Department of Mathematics and Computer Science University of Palermo) "Multi Layer Analysis" (2012)
- Reviewer of the doctoral thesis of Carmen Navarro (Departamento de Ciencias de la Computación and Inteligencia Artificial, Universidad de Granada, Espana) "Approach to Personalized Medicine through the development of new Artificial Intelligence methodologies "(2017).
- Reviewer of the doctoral thesis of Guido Zampieri (Ph.D. Course in Biosciences Curriculum: Genetics, Genomics and Bioinformatics, University of Padua) "Prioritization of candidate disease genes via multi-omics data integration". (2017)
- Reviewer of the Master of Science thesis in Artificial Intelligence by Jeremy Borg (Faculty of ICT, University of Malta) "Improved Performance of Error Correcting Output Codes for Multiclass Classification" (2017)
- Reviewer of the doctoral thesis of Juan Caceres Silva (Department of Computer Science, Royal Holloway University of London), "Charting disease gene propagation of disease phenotypic similarities". (2019)
- Reviewer of the doctoral thesis of Samuele Fiorini (Department of Computer Science, Bioengineering, Robotics and Systems Engineering University of Genoa) "Challenges in biomedical data science: data-driven solutions to clinical questions". (2018)
- Reviewer of Max Schubach's doctoral thesis (Department of Mathematics and Computer Science, Freie Universitat Berlin) "Learning the Non-Coding Genome" (2018)
- Reviewer of the doctoral thesis of Juan Caceres Silva (Department of Computer Science, Royal Holloway University of London), "Network Medicine Characterisation of Genetic Disorders by Propagation of Disease Phenotypic Similarities". (2019)

Supplementary teaching and student support activities.

• Tutor, thesis supervisor or co-supervisor for the following PhD students for the Doctorate in Computer Science, University of Milan:

 Francesca Ruffino: "Supervised Learning Methods for the Analysis of Gene Expression Data"
 Raffaella Folgieri: "Ensembles based on Random Projection for gene expression data analysis"
 Roberto Avogadri: "Unsupervised clustering methods for high-dimensional data anslysis"
 Francesco Saccà (dottorato in Matematica e Statistica per le Scienza Computazionali): "Problemi di Clustering con vincoli: algoritmi e complessita'"
 Marco Frasca: "Graph-based approaches for imbalanced data in functional genomics"
 Rajab Ali Keshavarz Emami: "Machine learning methods for the prediction of epileptic seizures"
 Marco Notaro: "Hierarchical Ensemble Methods for Ontology-based Predictions in Computational Biology"
 Alessandro Petrini: "Machine learning methods for the the prediction of pathogenic variants in non coding regions of the human genome"

- Supervisor of the following research fellows: 1) Matteo Re; 2) Marco Frasca
- Supervisor of the graduate student Jessica Gliozzo, winner of a scholarship abroad at the Dept. of Computer Science of Royal Holloway University of London, for the purposes of preparing the Master's thesis a.a. 2015/2016, funded by the University of Milan.
- Speaker or co-supervisor of more than 40 degree theses (triennial or "magistrale" or of the old system), in Machine Learning and Bioinformatics, for the degrees in Computer Science, Information and Communication Technologies, Molecular Biotechnologies and Bioinformatics, Mathematics, from 2005 to date.
- I have worked as a tutor for dozens of students in the undergraduate and graduate degrees in Computer Science.

Institutional activities

- UNIMI contact person for Area 06 Computer Science for the organization and census, in collaboration with UniMITT, of the research activities in the IT area with implications in the biomedical and bio-technological fields. from 2005 to 2007).
- Associate Member of the CNR Institute of Electronics and Information and Telecommunications Engineering for the project "Machine Learning Techniques for Modeling and Growing Up" (2007-2008)
- Member of the Doctoral College in Computer Science, University of Milan since 2008
- Member of the evaluation commission for the Faculty of Science of the University of Milan for the ERASMUS INTENSIVE PROGRAM (IP) a.a. 2009/2010 "Interdisciplinary approaches to microarray data analysis" organized by the University of Helsinki, Warwick, Naples (Federico II) and Milan.
- Member of the transfer commission and responsible for transfers to the degree course in Computer Science F1X of the University of Milan from other degree courses since the academic year 2010/11 to 2014/15.
- Member of the committee of the Computer Science department for educational and scientific exchange with the University of Beijing and for the technology transfer with the Chinese genomics company Rose Genomics (Beijing) as part of an agreement between the UNIMI Foundation, UNIMI and Rose Genomics (from 2018)
- Member of the Data Mining and Big Data Analytics Technique Committee (DMTC) IEEE Computational Intelligence Society (IEEE-CIS) (from 2015)

- Member of the scientific committee of the working group on Machine Learning and Data Mining, within AI * IA, Italian Association for Artificial Intelligence (from 2015)
- Member of the IEEE Computational Intelligence Society 2017 PhD Thesis Award committee (from 2018)

Publications

International journals

R58. L. Cappelletti, T. Fontana, G. Di Donato, L Di Tucci, E. Casiraghi and G. Valentini <u>Complex Data</u> <u>Imputation by Auto-Encoders and Convolutional Neural Networks: A Case Study on Genome Gap-Filling</u>, Computers 9(2):37, 2020

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