### Sébastien Benzekry

French Born February 22nd, 1985 benzekry@phare.normalesup.org http://benzekry.perso.math.cnrs.fr/ Inria team MONC Institut de Mathématiques de Bordeaux University of Bordeaux 351 cours de la Libération 33405 Talence Cedex, France +33 6 95725707

#### Situation

2013 -	Research Scientist (CR1) at Inria Bordeaux, team MONC
	Grant bonus for doctoral management and research achievements (PEDR)
	Formation
2012	Postdoctoral fellow in mathematical modeling of cancer from January 2012 until November 2012 at the Center of <b>Cancer and Systems Biology</b> at Tufts University, School of Medicine, Boston, under the supervision of Philip Hahnfeldt
2008 - 2011	Doctorate of applied mathematics
	LATP and Laboratoire de Pharmacocinétique et Toxicocinétique, Marseille
Title	Modeling, mathematical and numerical analysis of anti-cancerous thera- pies for metastatic cancers
2008	Laureate of the <b>Agregation of mathematics</b> , option Scientific calculus (rank 105).
2007 - 2009	Student at the Ecole Normale Supérieure of Cachan, 3rd year entrance concourse.
2004 - 2007	Mathematics magister at the Ecole Normale Supérieure of Paris. Mention Good.
2006 - 2007	Master 2 Mathematics of modeling at the University Pierre et Marie Curie, Paris, specialty biology. Mention Good.
	Master 2 internship at the mathematics department of the UPC in <b>Barcelone</b> : Exploration of deterministic and stochastic neuron networks
2004 - 2006	Bachelor and Master 1 degrees of Mathematics, ENS Paris.

#### Research interests

My research is focused on applied mathematics to cancer research. Some topics of my research include

- Data-based modeling of the metastatic development and tumor-tumor interactions
- Scheduling optimization for anti-cancerous therapies. Pharmacokinetics/Pharmacodynamics modeling in oncology

### International journals

Mathematical modeling of tumor-tumor inhibition supports a systemic control of tumor growth

**S. Benzekry**, C. Lamont, D. Barbolosi, L. Hlatky, P. Hahnfeldt *Cancer Research*, in press, (2017)

Non-Standard Radiotherapy Fractionations Delay the Time to Malignant Transformation of Low-Grade Gliomas

A. Henares-Molina, S. Benzekry, P.C. Lara, M. Garcia-Rojo, V.M. Perez-Garcia and A. Martinez-Gonzalez

PloS One, in press, (2017)

Model Driven Optimization of Antiangiogenics + Cytotoxics Combination : Application to Breast Cancer Mice Treated with Bevacizumab + Paclitaxel Doublet Leads to Reduced Tumor Growth and Fewer Metastasis

S. Mollard, J. Ciccolini, D.C. Imbs, R. El Cheikh, D. Barbolosi, S. Benzekry *Oncotarget*, Volume 5, 10.18632/oncotarget.15484, (2017)

In vivo bioluminescence tomography for monitoring breast tumor growth and metastatic spreading : comparative study and mathematical modeling

S. Mollard, R. Fanciullino, S. Giacometti, C. Serdjebi, **S. Benzekry**, Joseph Ciccolini *Scientific reports*, Volume 6, 36173, (2016)

Mathematical modeling of cancer immunotherapy and synergy with radiotherapy R. Serre, S. Benzekry, L. Padovani, C. Meille, N. André, J. Ciccolini, F. Barlési, X. Muracciole, and D. Barbolosi

Cancer Research, 76(17), 4931-40, (2016)

Modeling spontaneous metastasis following surgery : An In vivo/ in silico approach S. Benzekry, A. Tracz, M. Mastri, R. Corbelli, D. Barbolosi, J.M.L. Ebos Cancer Research, 76(3), 535-547, (2016)

Computational modelling of metastasis development in renal cell carcinoma E. Baratchart, S. Benzekry<sup>\*</sup>, A. Bikfalvi<sup>\*</sup>, T. Colin<sup>\*</sup>, L.S. Cooley, R. Pineau, E. Ribot, O. Saut, W. Souleyreau *BlaS. Computational Biology* 11(11), a1004626 (2015)

*PloS Computational Biology*, 11(11), e1004626, (2015)

Metronomics reloaded : theoretical models bringing chemotherapy into the era of precision medicine

**S. Benzekry**<sup>+</sup>, E. Pasquier<sup>+</sup>, D. Barbolosi, B. Lacarelle, F. Barlesi, N. Andre and J. Ciccolini Seminars in Cancer Biology, 35, 53-61 (2015)

# Improving efficacy of the combination between antiangiogenic and chemotherapy : Time for mathematical modeling support

J. Ciccolini, **S. Benzekry**, B. Lacarelle, D. Barbolosi and F. Barlési Proceedings of the National Academy of Sciences USA, 112(27), E3453, (2015)

#### Design Principles for Cancer Therapy guided by changes in complexity of Protein-Protein Interaction Networks

**S. Benzekry**, J.A. Tuszynski, E.A. Rietman, G.L. Klement *Biology Direct*, 10 (32), (2015)

Host Age is a Systemic Regulator of Gene Expression Impacting Cancer Progression A. Beheshti, S. Benzekry, J.T. MacDonald, L. Ma, M. Peluso, P. Hahnfeldt, L. Hlatky

Cancer Research, 75 (6), 1134-43, (2015)

Classical Mathematical Models for Description and Forecast of Preclinical Tumor Growth S. Benzekry, C. Lamont, A. Beheshti, A. Tracz, J.M.L. Ebos, L. Hlatky, P. Hahnfeldt *PloS Computational Biology*, 10(8), e1003800, (2014)

Global dormancy of metastases due to systemic inhibition of angiogenesis S. Benzekry, A. Gandolfi, P. Hahnfeldt *PloS One*, 9(1), e84249 (2014)

# Maximum tolerated dose versus metronomic scheduling in the treatment of metastatic cancers

S. Benzekry, P. Hahnfeldt Journal of Theoretical Biology, 335, 235-244, (2013)

# Theoretical investigation of the efficacy of antiangiogenic drugs combined to chemotherapy in xenografted mice

F. Lignet , S. Benzekry, S. Wilson, F. Billy, O. Saut, M. Tod, B. You, A. Adda Berkane, S. Kassour, M.X. Wei, E. Grenier, B. Ribba *Journal of Theoretical Biology*, 320, 86-99, (2013)

Modeling the impact of anticancer agents on metastatic spreading S. Benzekry, N. André, A. Benabdallah, J. Ciccolini, C. Faivre, F. Hubert, D. Barbolosi Mathematical Modeling of Natural Phenomena, 7 (1), 306-336, (2012)

A new mathematical model for optimizing the combination between antiangiogenic and cytotoxic drugs in oncology

**S. Benzekry**, G. Chapuisat, J. Ciccolini, A. Erlinger, F. Hubert Comptes Rendus de l'Académie des Sciences - Mathématiques, 350, 23-28, (2012)

Mathematical and numerical analysis of a model for anti-angiogenic therapy in metastatic cancers.

**S. Benzekry** *M2AN*, 46 (2), 207-237, (2012)

### Passing to the limit 2D-1D in a model for metastatic growth

**S. Benzekry** Journal of Biological Dynamics, doi :10.1080/17513758.2011.568071, (2011)

# Mathematical analysis of a two-dimensional population model of metastatic growth including angiogenesis

#### S. Benzekry

Journal of Evolution Equations, 11 (1), 187-213, (2011)

#### Book chapters

Les lois de la croissance tumorale S. Benzekry Bibliothèque Tangente, Hors-série Maths et Médecine, (2016)

#### A Mathematical Model for Growing Metastases on Oncologists's Service

D. Barbolosi, A. Benabdallah, S. Benzekry, J. Ciccolini, C. Faivre, F. Hubert, F. Verga, B. You In Computational Surgery and Dual Training, Springer New York, 331-338, (2014)

### Proceedings and protocols

On the growth and dissemination laws in a mathematical model of metastatic growth S. Benzekry, J. ML  ${\rm Ebos}$ 

ITM Web of Conferences, 5, (2015)

Capturing the Driving Role of Tumor-Host Crosstalk in a Dynamical Model of Tumor Growth

**S. Benzekry**, A. Beheshti, P. Hahnfeldt, L. Hlatky *bio-protocol*, 5(21), e1644, (2015)

# Model-based optimization of combined antiangiogenic + cytotoxics modalities : application to the bevacizumab-paclitaxel association in breast cancer models

S. Mollard, S. Benzekry, S. Giacometti, C. Faivre, F. Hubert, J. Ciccolini, and D. Barbolosi Cancer Research, In : Proceedings of the 105th Annual Meeting of the American Association for Cancer Research, 74, (2014)

A mathematical model of systemic inhibition of angiogenesis in metastatic development S. Benzekry, A. Gandolfi and P. Hahnfeldt

ESAIM : Proceedings, 45, 75-87, (2014)

# International conferences (invited)

December 2017	Mathematical Methods and Modeling of Biophysical Phenomena, Rio de Ja- neiro, Brazil
May 2016	Metronomics @ Mumbai, Mumbai, India
December 2015	Present challenges of mathematics in oncology and biology of cancer, Marseille, France.
November 2015 November 2015	Dynamique et contrôle des croissances tumorales, Rouen, France. Contrôle des EDP et applications, Marseille, France.
October 2015 June 2015	Journées du Groupe de Métabolisme et Pharmacocinétique, Paris, France. Micro and Macro Systems in Life Sciences (MMSLS 2015), Bedlewo, Poland
March 2015	Workshop on hybrid and multiscale modelling in cell and cell population bio- logy, Laboratoire Jacques-Louis Lions, Paris, France. <i>Combined in vivo and</i> <i>in silico quantitative modeling of post-surgery metastatic development</i>
March 2015	Mathematical Methods and Modeling of Biophysical Phenomena, Cabo Frio, RJ, Brazil. A modeling study of metastatic initiation and tumor-tumor spatial interactions
January 2015	36th EORTC PAMM Winter meeting, Marseille, France. A translational in vivo/in silico quantitative modeling study of the impact of surgery on metas- tatic relapse and survival in breast cancer
October 2014	Autumn School by Japanese and French Mathematicians, Osaka, Japan. Ma- thematical modeling of tumor growth and metastatic spread. Data, theories and nredictions
July 2014	10th AIMS Conference on Dynamical Systems, Differential Equations and Applications, Madrid, Spain. A mathematical model of systemic inhibition of anaiogenesis in metastatic development
November 2013	French-Mexican Meeting on Industrial and Applied Mathematics, Villaher- mosa, México. Classical Mathematical Models for Description and Forecast of Preclinical Tumor Growth
May 2013	Bi-annual congress of the SMAI, Seignosse, France. Mathematical modeling of systemic inhibition of angiogenesis and metastatic dynamics
March 2013	Workshop on Mathematical Methods and Modeling of Biophysical Pheno- mena, Cabo Frio, Brazil. <i>Mathematical modeling of systemic inhibition of an-</i> giogenesis and metastatic dynamics
July 2012	2nd Annual Workshop on Cancer Systems Biology - Tumor Metronomics : Timing and Dose Level Dynamics, Center for Cancer and Systems Biology, Boston, USA. <i>Mathematical modeling of metastatic development and schedu-</i> <i>ling optimization of anti-cancerous therapies</i>
July 2011	7th International Congress of Applied and Industrial Mathematics (ICIAM 2011), Vancouver. <i>Mathematical modeling of the metastatic process and opti-</i> <i>mization of anti-cancerous therapies</i> , in the session Tumor growth modeling and system identification for clinical applications
July 2011	Workshop on Systems Biology of Tumor Dormancy. St. Elizabeth's Medical Center, Boston, USA. A modeling approach for therapies in metastatic cancers
May 2011	Journée Emergence de l'Institut Gustave Roussy (IGR), Paris. Un exemple de modélisation du processus métastatique

May 2010	2nd Workshop on Metronomic Anti-Angiogenic Chemotherapy in Paediatric
	Oncology, Marseille. Mathematical modeling of MTD and metronomic temo-
	zolomide
May 2010	8th conference of the American Institute of Mathematical Sciences (AIMS),
	Dresde. Modeling and mathematical analysis of metastatic growth under an-
	giogenic control, in the session Evolution equations and mathematical biology

# Accepted communications

March 2017	38th EORTC-PAMM Winter meeting, Split, Croatia. Optimization of the ti- ming of sequential administration of bevacizumab plus cutotorics in NSCLC
	by a mathematical model. (Oral communication)
October 2015	Journées du cancéropôle Grand Sud-Ouest, Bordeaux, France. Modeling spon- taneous metastasis following surgery : an in vivo-in silico approach (Poster)
June 2015	Mathematical Methods in Systems Biology, Dublin, Ireland. (Oral communication)
June 2014	European Conference on Mathematical and Theoretical Biology, Göteborg, Sweden. <i>Metastatic dynamics, systemic inhibition of angiogenesis and impli-</i> <i>cations for surgery</i> (Oral communication)
June 2014	International Tumor Dormancy Symposium, Lille, France. <i>Metastatic dyna-</i> <i>mics and systemic inhibition of angiogenesis. Implications for dormancy and</i> <i>surgery</i> (Poster)
October 2011	Journées du Groupe de Métabolisme et Pharmacocinétique. Paris. Biomathe- matical modeling for description of metastatic processes and optimization of combined anti-angiogenic + cytotoxic therapies (Poster and oral communica- tion)
June 2011	8th European Conference of Mathematical and Theoretical Biology, Krakow. Optimal schedules for therapies in metastatic cancers
June 2010	CANUM 2010, Gironde. A model of metastatic growth under angiogenic control
May 2010	CMPD 3, Bordeaux. A model of metastatic growth under angiogenic control
November 2009	Conference of biomathématics et biomechanics, Tozeur, Tunisie. Poster Ana- lysis and modeling of metastatic growth including angiogenesis

## Seminars

May 2017	Laboratoire d'Analyse Topologie et Probabilites, Institut de Mathematiques de Marseille, Mathematical Oncology : Theory Maste Reality, Marseille, France
November 2016	Integrated Mathematical Oncology Department, Moffitt Cancer Center. Ma- thematical Modeling of Metastasis : Theory Meets Reality. Tampa, Florida, USA.
October 2016	Department of Genetics, Roswell Park Cancer Institute (invitation by John ML Ebos). <i>Mathematical Modeling of Metastasis : Theory Meets Reality</i> . Buffalo, NY, USA.
October 2016	Robert Kerbel's laboratory at Sunnybrook Research Institute. <i>Mathematical Modeling of Metastasis : Theory Meets Reality</i> . Toronto, Canada.
October 2016	Mathematics Department Colloquium, Ryerson University. <i>Mathematical Mo-</i> deling of Metastasis : Predicting the invisible. Toronto, Canada.
May 2016	Seminar at the laboratory of Génétique, Immunothérapie, Chimie et Cancer" of the University of Tours, Tours, France. <i>Data-based mathematical modeling</i> <i>analysis of preclinical studies in oncology</i>
January 2016	Seminar at the laboratoire d'imagerie biomédicale, Paris, France. Classical Mathematical Models for Description and Prediction of Experimental Tumor Growth
May 2015	Biomathematics Seminar, University of Gothenburg, Sweden Modeling spon- taneous metastasis following surgery : An In vivo/ in silico approach
March 2014	Seminar of the inria team Dracula, Lyon, France. Classical Mathematical Mo- dels for Description and Prediction of Experimental Tumor Growth
April 2013	ANEDP team, Orsay, France. Mathematical modeling of systemic inhibition of angiogenesis and metastatic dynamics
February 2013	LATP. Marseille. A math walk in a biology lab
December 2011	Team MC2, Bordeaux. Modeling, mathematical and numerical analysis of anti- cancerous therapies
Novembee 2011	EDP Seminar of the Mathematics department, Besançon. Modeling, mathe- matical and numerical analysis of anti-cancerous therapies
Mai 2010	Math-Bio seminar Maths-Bio of the university Lyon 1. Modeling and mathe- matical analysis of metastatic growth under angiogenic control
October 2009	Maths-Cancer working group, School of Pharmacy, Marseille. <i>Modeling of angiogenesis</i>
June 2009	Bucolic days of the doctoral seminar, LATP, Marseille

# Foreign research visits

Fall 2016	Short sabbatical at the <b>Roswell Park Cancer Institute</b> (J. Ebos' lab), Buffalo, NY, USA
	Stay at the <b>Integrated Mathematical Oncology</b> department of the Moffitt
	Cancer Center, Tampa, FL, USA
September 2015	Mathematical Oncology Laboratory (Môlab), Ciudad Real, Spain. Col-
	laboration with V. Pérez-Garcia and A. Martinez-González
May 2011	Instituto di Analisi dei Sistemi ed Informatica "Antonio Ruberti", Consiglio
	Nazionale delle Ricerche, Rome. Collaboration with A. Gandolfi et A. d'Ono-
	frio

### PhD students

2016 - 2019	Chiara Nicolò : "Mathematical modeling of systemic aspects of cancer and cancer therapy" (co-supervision with O. Saut)
2013 - 2015	Etienne Baratchart : "Quantitative study of the metastatic process using mathematical modeling" (co-supervision with T. Colin and O. Saut)

#### Master 2 internships

2017	M. Bilous (Ecole Polytechnique, M2 of biomechanical engineering) : "Mathe- matical modeling and prediction of clinical brain metastases in non-small cell lung cancer"
2014	Aristoteles Camilo (IMPA, Rio de Janeiro, Brazil and University Pierre et Marie Curie, Paris, France) : "An experimentally-based modeling study of the effect of anti-angiogenic therapies on primary tumor kinetics for data analysis of clinically relevant animal models of metastasis"

#### Other internships

2016	Laura Lumale (INSA engineering school, 2nd year (M1)) : "Around a mathematical model of the concomitant tumor resistance phenomenon"
2015	Simon Evain ("Ponts et Chaussées" engineering school, 2nd year (M1)) : "Ma- thematical modeling of tumor and metastatic growth when treated with suni- tinib"
2015	Maxime Prodhomme (ENS Cachan, 1st year (L3))
Teaching	

- 2015 2017 Teaching assistant for ordinary differential equations in the first year of engineering school Matmeca, Bordeaux, France
- 2014 2017 Supervision of various projects in applied mathematics in Matmeca (L3) or University of Bordeaux (L3 and M1), Bordeaux, France
- 2009 2011 Teaching duties in mathematics (analysis, topology and differential calculus, numerical analysis, preparation to the agregation), 192 hours, University of Aix-Marseille, France.

Reviewer for grant applications submitted to : the Austrian Science Fund, the Centre d'Excellence Africain en Technologies de l'Information et de la Communication and the TEAM program (European Union and Foundation for Polish Science).

Invited editor of a special issue for the journal Complexity : "Mathematical Oncology : Unveiling Biological Complexity Using Mathematical Methods"

Reviewer for several international journals including **modeling journals** (Journal of Theoretical Biology, Bulletin of Mathematical Biology, PloS One, Mathematical Biosciences, Theoretical Biology and Medical Modeling, Mathematical Biosciences and Engineering, Journal of Biological Informatics, Journal of Biological Systems, ESAIM :Proc, Mathematics and Computers in Simulation) and **biological/medical journals about cancer and pharmacokinetics** (British Journal of Cancer, Clinical Pharmacokinetics, BMC Cancer, Breast Cancer Research and Treatment).

June 2018Member of the organizing committee of the 3rd Thematic school "Present<br/>challenges of mathematics in oncology and biology of cancer : modeling and<br/>mathematical analysis"July 2015Member of the scientific committee of the MB2 conference (Bio-<br/>mathematics modeling days of Besançon)March 2012Member of the organizing committee of the 2nd Thematic school "Present<br/>challenges of mathematics in oncology and biology of cancer : modeling and<br/>mathematics in oncology and biology of cancer : modeling and<br/>mathematics in oncology and biology of cancer : modeling and<br/>mathematical analysis"